

# **NPQIC Summit Meeting:** **The Re-Emergence of Syphilis: New Screening Recommendations and Maternal / Neonatal Case Reviews**

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Neonatology, PGY-6

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Obstetrics and Gynecology, PGY-4



# Learning Objectives & Goals

1. Identify syphilis as a re-emerging health epidemic.
2. Define proper screening recommendations for pregnant people and neonates in your community.
3. Implement proper screening for pregnant people and neonates in your community (we'll show you how we did it!).
4. Recognize congenital syphilis in the neonate and when to seek Infectious Disease consultation.

# Disclosures

The following have no relevant financial relationships to disclose:

- Jennifer Berger, MD
- David Greiner, MD

# LEARNING OUTCOME

Identify syphilis as a re-emerging health epidemic.

# Syphilis is Back...With a Vengeance!

Daily Briefing



## Syphilis rates have hit their highest since 1950. Why?

Posted on February 01, 2024

Updated on February 01, 2024

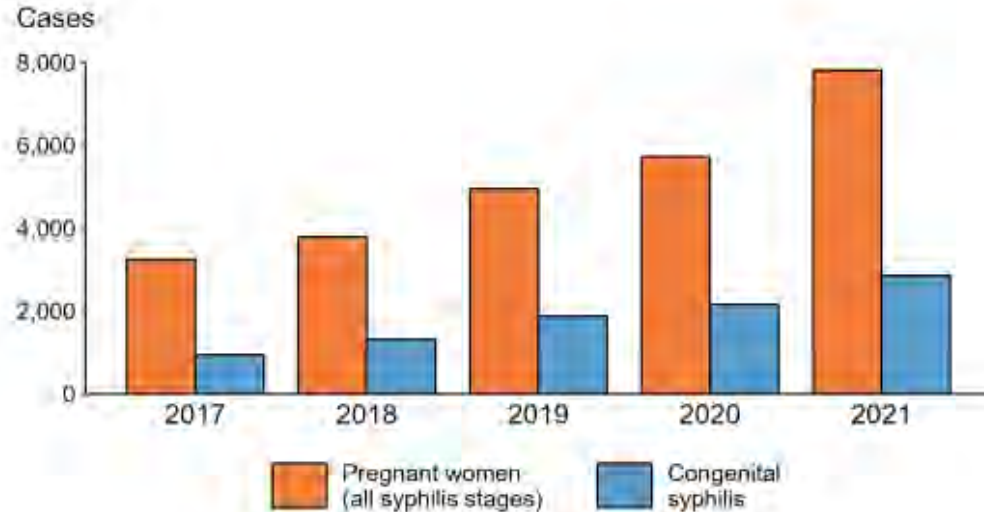
According to a CDC report for 2022:

- 207,255 syphilis cases in US
- 17% increase in 1 year, 80% increase in 5 years
- **3,761 infants** born with syphilis

(CDC National Notifiable Disease Surveillance System)

*Syphilis rates have hit their highest since 1950. Why?* (n.d.). Retrieved March 25, 2024, from <https://www.advisory.com/daily-briefing/2024/02/01/syphilis>

# National Syphilis Rates in Pregnant People and Neonates



- 10x as many babies born with syphilis in 2022 than a decade prior
- Rates at highest level in 30 years
- 231 related stillbirths
- 51 infant deaths

Jenco, M., & Editor, N. C. (2023). *Ten times as many babies born with syphilis in 2022 compared to 2012.*

<https://publications.aap.org/aapnews/news/27259/Ten-times-as-many-babies-born-with-syphilis-in>

# Syphilis is Curative and Preventable!

AAP News™

## Ten times as many babies born with syphilis in 2022 compared to 2012

November 7, 2023

Melissa Jenco, News Content Editor

88% of cases could have been **prevented** with timely testing and treatment

- 37% untimely or no testing
- 11% no documented treatment
- 40% inadequate treatment



Jenco, M., & Editor, N. C. (2023). *Ten times as many babies born with syphilis in 2022 compared to 2012.*

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# Congenital syphilis cases are highest in...



*Syphilis rates have hit their highest since 1950. Why?* (n.d.). Retrieved March 25, 2024, from <https://www.advisory.com/daily-briefing/2024/02/01/syphilis>



# Why is syphilis re-emerging?

Daily Briefing

## Syphilis rates have hit their highest since 1950. Why?



Posted on February 01, 2024

Updated on February 01, 2024

- **We aren't looking for it**
- Lack of resources
- Barriers to Care
- Lack of prenatal care
- Lack of funding
- Cases multiplying
- Substance Use Disorder
- Healthcare Disparities

**AAP News™**

**Ten times as many babies born with syphilis in 2022 compared to 2012**

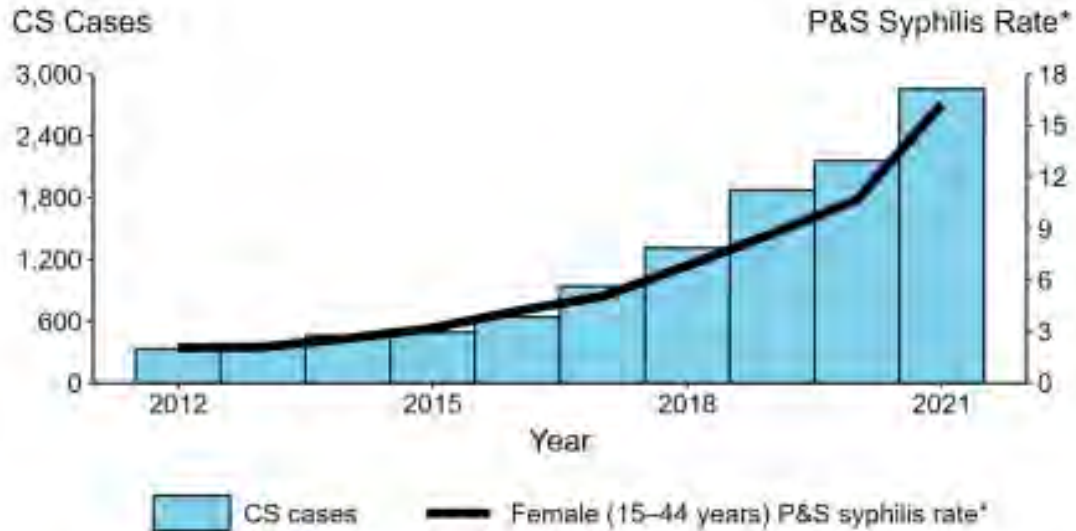
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UNIVERSITY OF NEBRASKA MEDICAL CENTER | CHILDREN'S NEBRASKA

# Congenital Syphilis is a Re-Emerging Health Epidemic



- **10x as many babies born with syphilis in 2022 than decade prior**
- **88% of cases could have been prevented w/ timely testing and treatment**
- **231 stillbirths, 51 infant deaths**

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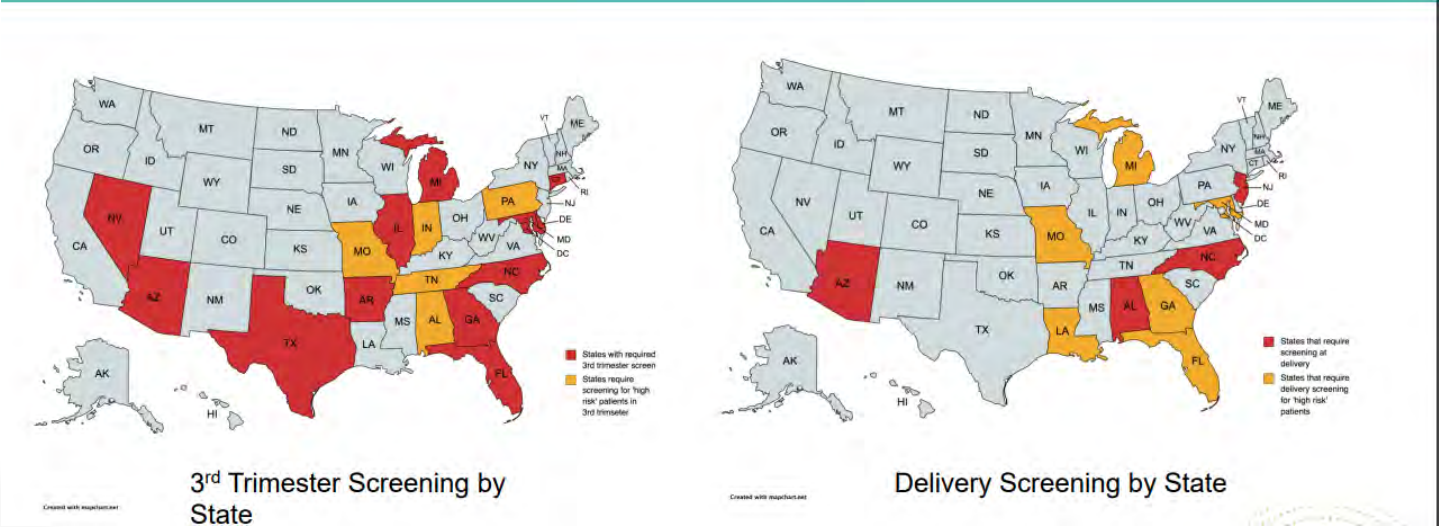
# LEARNING OUTCOME

Define proper screening recommendations for pregnant people and neonates in your community.

# Previous ACOG Recommendations

- Universal screening at the first prenatal visit
- Pregnant women at **high risk** of syphilis **may** be retested in the 3rd trimester and at delivery.
- Some states **require** re-testing at delivery even for women who are not at high risk for infection

# Mandatory Screening By State



# Screening at First Visit

- For many patients this may be in the ED
- Encouraged screening as early as possible to help detect and treat cases

# How do you define "High Risk"

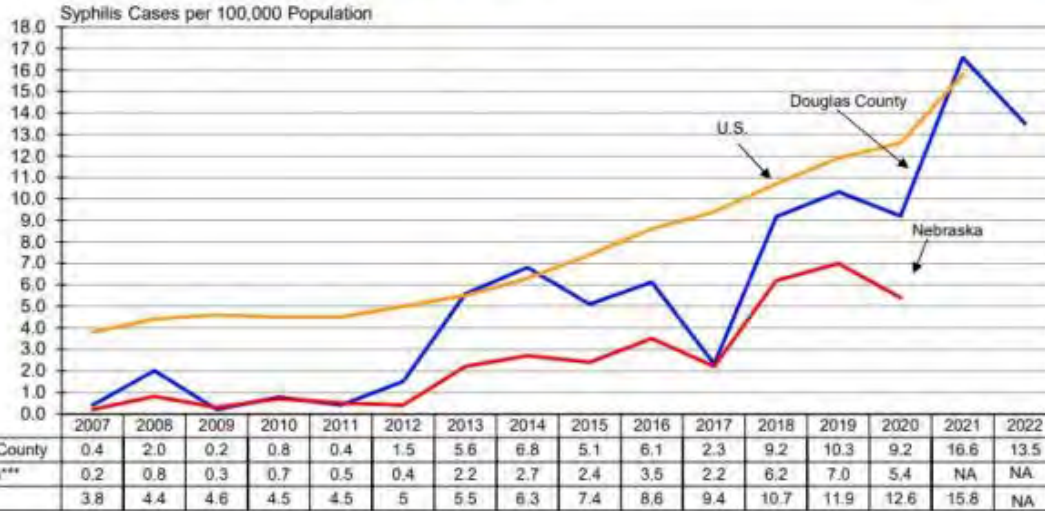


# Community Risk Factors

- No previously set cutoff
- Reasonable to use the CDC "Healthy People 2030" goal
- **4.6/100,000** reproductive age females



## Syphilis\* Rates\*\* Douglas County, Nebraska, and US 2007-2022



\* Primary and Secondary Syphilis.

\*\* Populations used for rate calculations are Projection Estimates for Douglas County from Woods & Poole Economics except for 2010 which uses the US Decennial Census, and 2011-2020 which use the US Census Bureau's Intercensal Population Estimates Program. 2022 rates are provisional because they use the 2021 population estimates. 2022 estimates are not yet available.

\*\*\* Source: CDC Sexually Transmitted Disease Surveillance

NA - Data is not yet available.

Source: DCHD Sexually Transmitted Disease Surveillance

Based on Date of Report

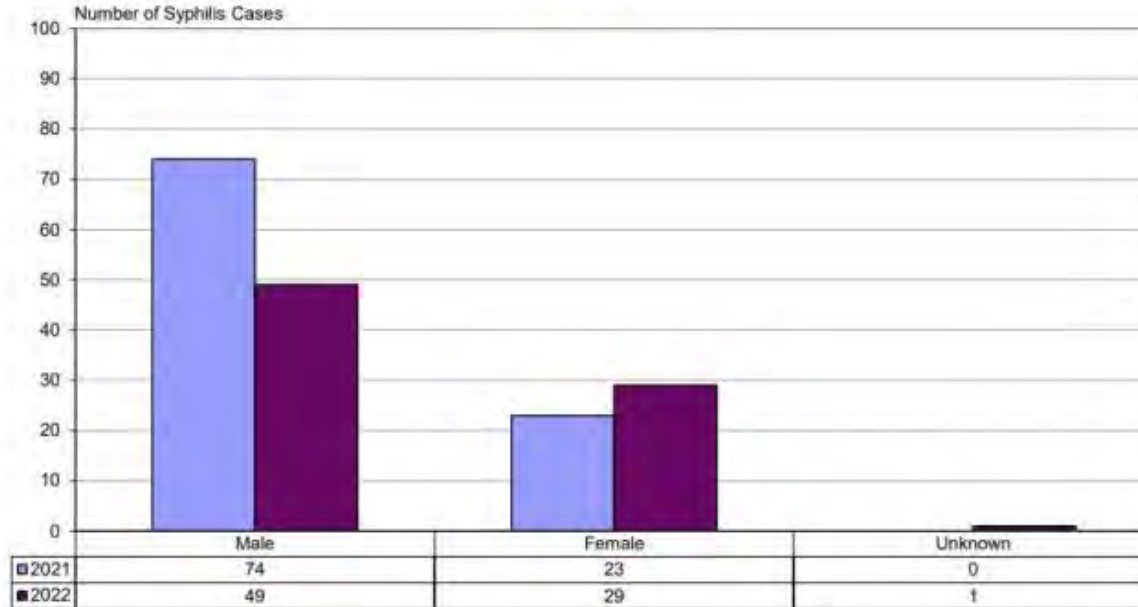
Douglas County Health Department

04/12/2023

## Number of Syphilis\* Cases by Gender

### Douglas County, NE

### 2021-2022



\* Primary and Secondary Syphilis  
Source: DCHD Sexually Transmitted Disease Surveillance  
Based on Date of Report

Douglas County Health Department  
04/12/2023



cdc sti atlas



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About 220,000 results (0.25 seconds)



Centers for Disease Control and Prevention (.gov)

<https://www.cdc.gov/nchhstp/atlas>



## NCHHSTP AtlasPlus

NCHHSTP AtlasPlus gives you the power to access data reported to CDC's National Center for HIV, Viral Hepatitis, STD, and TB Prevention (NCHHSTP).

[AtlasPlus](#) · [About the Atlas](#) · [Buttons](#) · [What's New](#)



1. Indicator    2. Geography    3. Year    4. Demographics

Start over    Previous    Create my table

Hispanic/Latino, then you must also select those two race/ethnicity groups. Not all data are available at all stratification levels. If a stratification is not available, the option will be greyed out.

**Age Group**

All age groups

Select specific age groups

- 0-14
- 15-19
- 20-24
- 25-29
- 30-34
- 35-39
- 40-44
- 45-54
- 55-64
- 65+
- Unknown

**Race/Ethnicity**

All races/ethnicities

Select specific races/ethnicities

- American Indian/Alaska Native
- Asian
- Black/African American
- Hispanic/Latino
- Multiracial
- Native Hawaiian/Other Pacific Islander
- Unknown
- White

**Sex**

Both sexes

Select specific sexes

- Male
- Female

Start over    Previous    Create my table

**Current selections**

**Indicator (3 selected)**

- Primary and Secondary Syphilis \* ‡
- Early Non-Primary, Non-Secondary Syphilis \* ‡
- Unknown Duration or Late Syphilis \* ‡

**Geography (1 selected)**

- Douglas County, NE

**Year (1 selected)**

- 2022

**Age Group (6 selected)**

- 15-19
- 20-24
- 25-29
- 30-34
- 35-39
- 40-44

**Race/Ethnicity (1 selected)**

Indicator	Year	Geography	Cases	Rates per 100,000
Unknown or Late	2022	Douglas County, NE	55	44.9
Early, Non-primary Non-Secondary	2022	Douglas County, NE	9	7.3
Primary and Secondary	2022	Douglas County, NE	27	22.0

# What about UNMC?

- Does not represent the entire community
- May be falsely elevated as these are patients seeking care

# UNMC Rates

- Total reproductive age females (Age 15-45) at UNMC in 2023: 85,801
- Positive Syphilis (Syphilis antibody and RPR): 132
- $132/85,801 = 0.001538$  or 0.15%
- **153/100,000 Reproductive Age Females**

# UNMC Pregnancy Rates

- Total pregnancies screened in 2023 = 3,713
- Positive syphilis = 26
- $26/3713 = 0.007002$  or 0.7%
- **700/100,000 Pregnant patients**

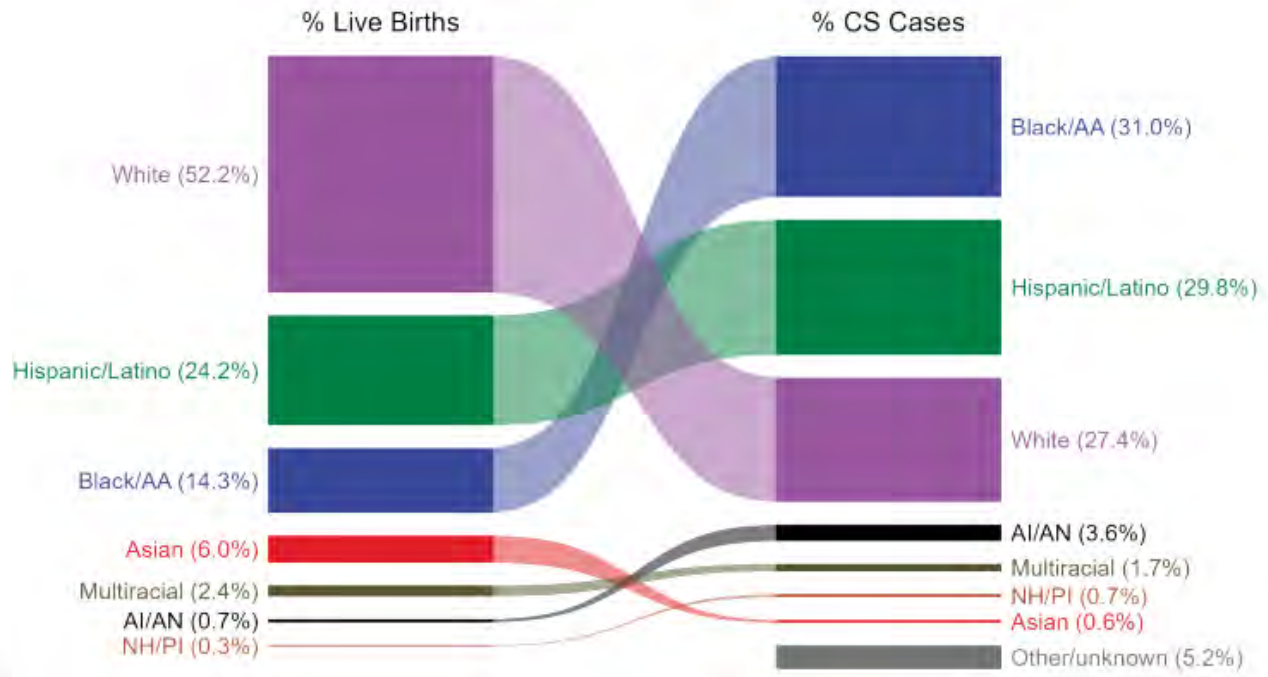


# April 2024 ACOG Recommendations

- Universal screening at first prenatal visit, third trimester, and at delivery
- 2 out of 5 infants with congenital syphilis had no access to prenatal care
- It's impossible to talk about syphilis without recognizing bias

# Rates of Reported Congenital Syphilis Cases by Race/Hispanic Ethnicity of the Mother, United States, 2022

Race or Ethnicity	Rates of Reported Congenital Syphilis Cases, 2022 (per 100,000 live births)
American Indian or Alaska Native	644.7
Asian	10.4
Black or African American	214.5
Hispanic or Latino	124.1
Multiracial	79.0
Native Hawaiian or Pacific Islander	404.4
White	54.1



# LEARNING OUTCOME

Implement proper screening for pregnant people and neonates in your community.

This is how we did it.



# QI Project

- EPIC Changes live May 13
- Collect data on compliance w/ screening between June 1, 2024 – August 30, 2024

## Results Console

### 1st Trimester

+ Order All

Blood Type/ Rh	+ Order
Antibody Screen	+ Order
HCT	+ Order
HGB	+ Order
Platelet count	+ Order
Rubella	+ Order
Varicella	+ Order
Syphilis	+ Order
HBsAg (NMC result)	+ Order
HBsAg (outside result)	+ Order
Hep C	+ Order
HIV	+ Order
Glucose tolerance 1 hr	+ Order
HGBA1C	+ Order
TSH	+ Order
Urine Culture	+ Order

### 2nd Trimester

+ Order All

HCT	+ Order
HGB	+ Order
Glucose Tolerance 1 hr	+ Order
HGBA1C	+ Order
Quad Screen	+ Order
Alpha Fetoprotein	+ Order
Syphilis	+ Order

### 3rd Trimester

+ Order All

Antenatal Rhogam	+ Order
HCT	+ Order
HGB	+ Order
Glucose Tolerance 1 hr	+ Order
HGBA1C	+ Order
GBS	+ Order
Syphilis	+ Order

## Order Sets


### ▼ Labs / Imaging

#### ▼ Lab

OB Lab Panel - If prenatal lab unavailable.

Type and screen OB profile (\$40)

Indication: OB profile

 Once, today at 0948, For 1 occurrence  
No Phase of Care, Sign

CBC with differential, platelet

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

Rubella antibody, IgG

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

Varicella zoster antibody, IgG qualitative

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

Treponema pallidum (syphilis) antibody

No Phase of Care

Hepatitis B surface antigen

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

CBC without differential, platelet

No Phase of Care

CBC with differential, platelet

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

Comprehensive metabolic panel

No Phase of Care

HIV antigen/antibody panel

No Phase of Care

Rapid HIV 1,2 Screen (\$45)

No Phase of Care

DIC screen (\$48)

No Phase of Care

T3, free

No Phase of Care

POCT vaginal wet prep

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

POCT fern test

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

POCT vaginal pH for rupture of membranes

Once, today at 0948, For 1 occurrence

No Phase of Care, Sign

Treponema pallidum (syphilis) antibody

Once, today at 0948, For 1 occurrence

Pt. Portal result release timeframe: Auto Release Standard  
No Phase of Care, Sign

Diabetic Specific Labs - NMC

- Ob Labor Admission
- Ob Antenatal admission
- NERAS Ob Pre-surgery
- Ob Postpartum admission

# LEARNING OUTCOME

Recognize congenital syphilis in the neonate  
and when to seek Infectious Disease  
consultation.



# Case #1

Gestational Age: 39w2d at birth

Reason for Admission: Concern for congenital syphilis

Maternal History: 28 yo G5P4014 with pregnancy notable for maternal substance use, limited prenatal care, and syphilis infection.

- 1st syphilis test done on a triage visit
  - o + treponemal test and + RPR titers
- Unable to get ahold of patient with positive test results
- Next presentation was to L&D for imminent delivery

Delivery: SVD

# How should we look for syphilis?

Recommendations and Reports

## CDC Laboratory Recommendations for Syphilis Testing, United States, 2024

John R. Papp, PhD<sup>1</sup>; Ina U. Park, MD<sup>1,2</sup>; Yetunde Fakile, PhD<sup>1</sup>; Lara Pereira, PhD<sup>3</sup>; Allan Pillay, PhD<sup>1</sup>; Gail A. Bolan, MD<sup>1</sup>

<sup>1</sup>*Division of STD Prevention, National Center for HIV, Viral Hepatitis, STD, and TB Prevention, CDC, Atlanta, Georgia;*

<sup>2</sup>*University of California San Francisco, San Francisco, California;* <sup>3</sup>*The Task Force for Global Health, Decatur, Georgia*

# Testing

Treponemal tests: detect antibody response to antigens specific to *T. pallidum* (TP-PA, FTA-ABS, MHA-TP, TPHA, CLIA)

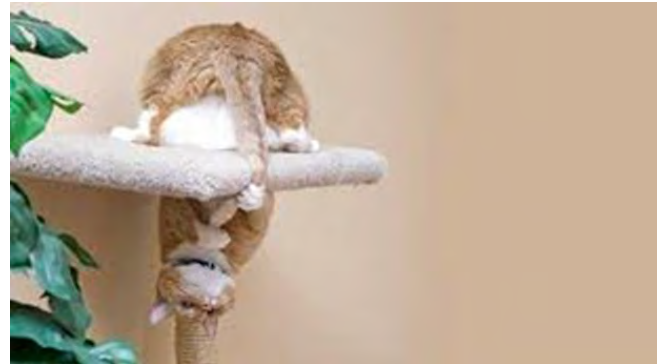
Non-treponemal (**lipoidal antigen tests**): RPR and VDRL

→ Detect damage to host cells (cardiolipin, lecithin, cholesterol)  
(damage to cells kind of unique to syphilis – 89% of the time)

→ A **titer** can be calculated

→ Used for screening, treatment outcomes, reinfection detection

BUT, false positives can happen.

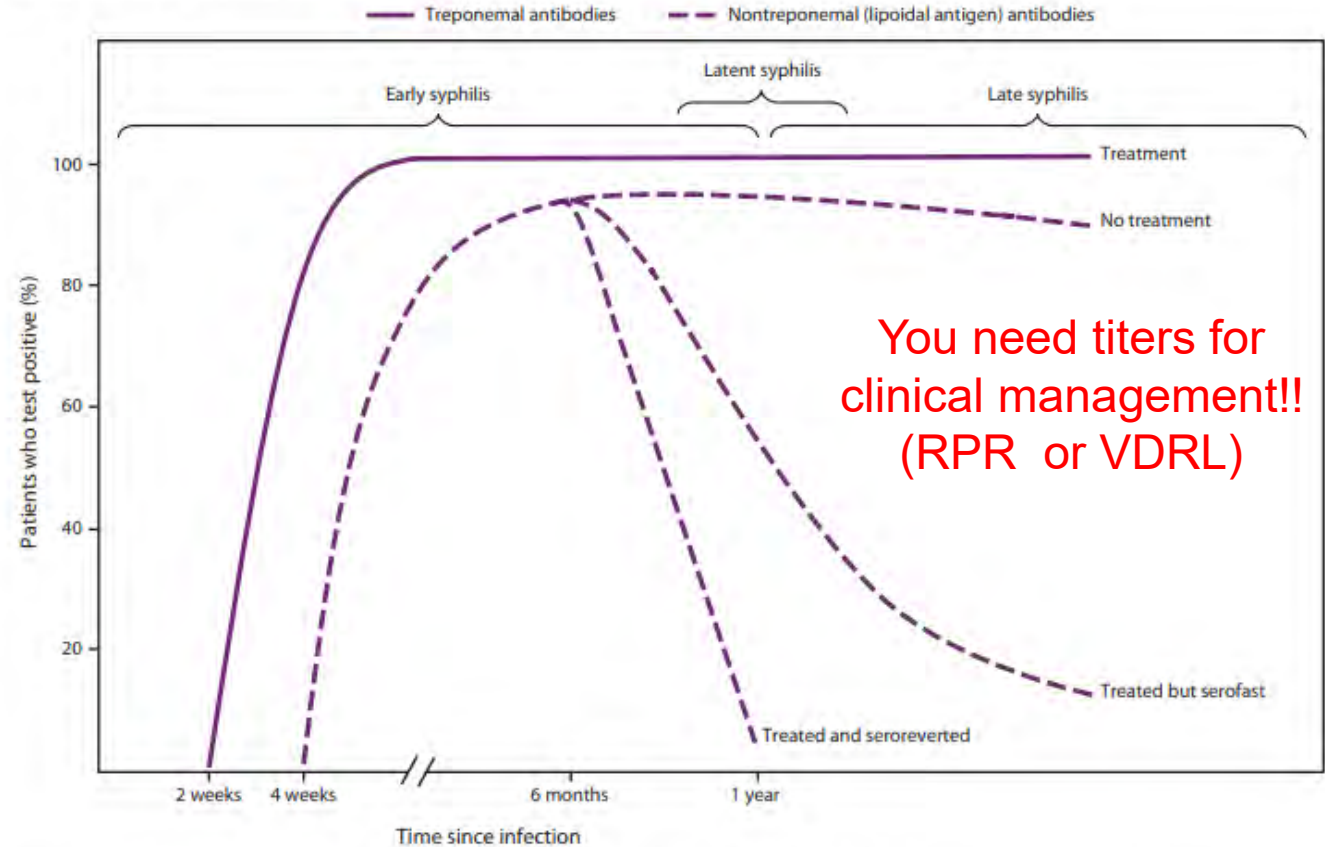


# Testing

## Treponemal Tests

- Detect antibodies to *T. pallidum*
- In general, positive for life!!

FIGURE 1. Serologic response to infection with *Treponema pallidum*, the causative agent of syphilis



Source: Adapted from Peeling RW, Mabey D, Kamb ML, Chen X-S, Radolf JD, Benzaken AS. Syphilis. Nat Rev Dis Primers 2017;3:17073. Used with permission.

Papp, J. R., Park, I. U., Fakile, Y., Pereira, L., Pillay, A., & Bolan, G. A. (2024). CDC Laboratory Recommendations for Syphilis Testing, United States, 2024. MMWR. Recommendations and Reports, 73(1), 1–32. <https://doi.org/10.15585/mmwr.rr7301a1>

# How do we test babies?

- Passive transfer of maternal antibody can cause positive treponemal test results in neonates and infants for up to 1 year (hematogenous spread)
- Performing treponemal tests on neonatal serum is not currently recommended -- not even IgM based tests
- Quantitative nontreponemal lipoidal antigen tests ARE recommended
  - Perform VDRL or RPR on neonatal serum NOT umbilical cord blood
  - Use the same test used on the mother so titers can be compared

# Case #1

## What does the baby need?

Mom had an RPR – we need an RPR.

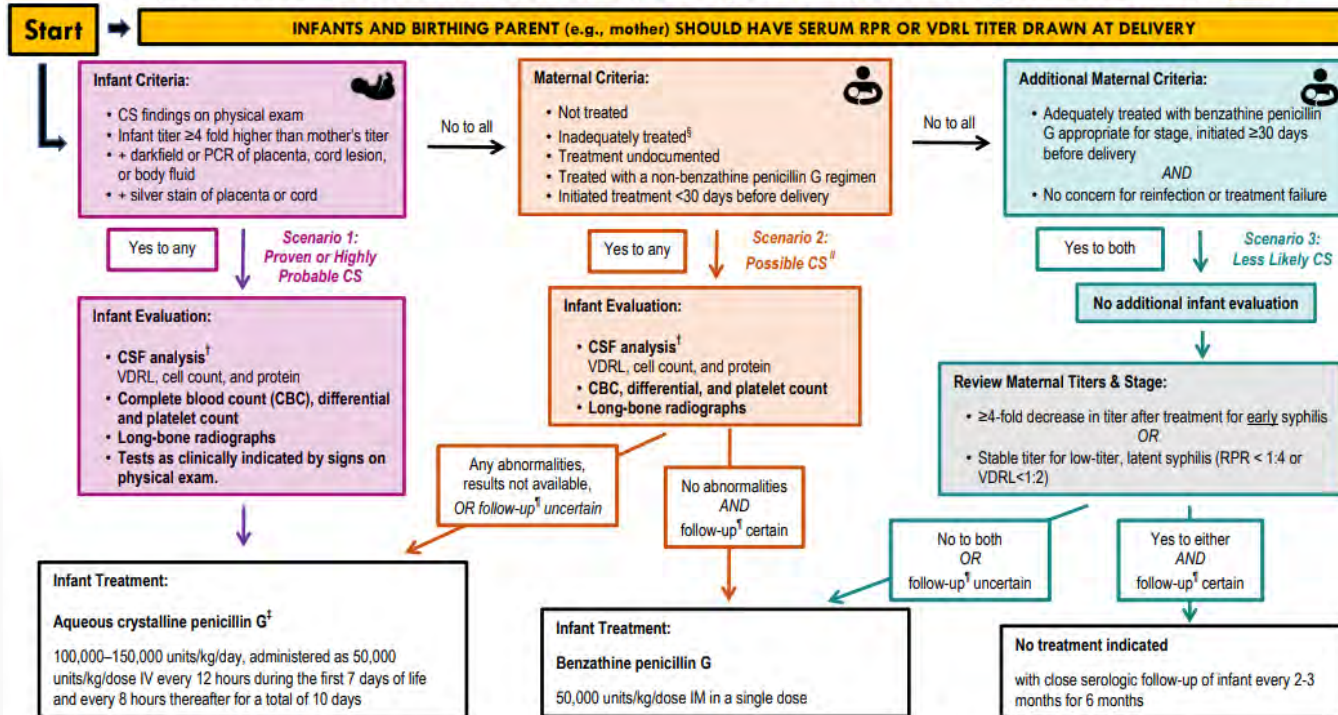
- RPR elevated with titer 1:4

Do we need any other workup? YES

Should we consult ID? YES

# CONGENITAL SYPHILIS (CS)

## Evaluation and treatment of infants (<30 days old) exposed to syphilis in utero\*



New tool for clinicians unveiled to ensure appropriate treatment of congenital syphilis. (n.d.). *California PTC*. Retrieved September 25, 2024, from <https://californiaptc.com/in-the-news/new-tool-for-clinicians-unveiled-to-ensure-appropriate-treatment-of-congenital-syphilis/>

\* Scenario 4 – in which an infant at delivery has a normal physical exam and titer < 4-fold mother's titer, AND the mother was adequately treated prior to becoming pregnant and sustains RPR titers <1:4 or VDRL <1:2 throughout pregnancy – is not included.

<sup>†</sup> CSF test results obtained during the neonatal period can be difficult to interpret; normal values differ by gestational age and are higher in preterm infants.

<sup>‡</sup> Alternative: Procaine penicillin G 50,000 units/kg/dose IM in a single daily dose for 10 days.

<sup>§</sup> Benzathine Penicillin G (BPG or Bicillin-LA), administered according to stage of disease and initiated at least 30 days prior to delivery is the only adequate treatment for syphilis during pregnancy.

<sup>¶</sup> Evaluation is not necessary if a 10-day course of parenteral therapy is administered, although such evaluations might be useful. If the neonate's nontreponemal test is nonreactive and the mother's risk for untreated syphilis is low, a single IM dose of BPG can be considered without evaluation.

<sup>¶¶</sup> All neonates with reactive nontreponemal tests should receive careful follow-up examinations and serologic testing (i.e., a nontreponemal test) every 2–3 months until the test becomes nonreactive. Neonates with a negative nontreponemal test at birth whose mothers were seroreactive at delivery should be retested at 3 months to rule out serologically negative incubating congenital syphilis at the time of birth.

FOR MORE INFORMATION ABOUT SCENARIO 4 MANAGEMENT, TREATMENT OF SYPHILIS IN PREGNANCY, NEONATAL CSF INTERPRETATION, AND CS INFANT FOLLOW-UP, PLEASE REFER TO THE CDC 2021 STI TREATMENT GUIDELINES.

# Case #1

Mom was positive for syphilis and **NOT** treated appropriately per state disease investigators.

Completed 10 days of penicillin G treatment.

Repeat RPR at 3 and 6 mos outpatient.

- \*CBC including differential and platelets: wnl
- \*LP with negative VDRL, normal cell count
- \*Ophthalmologic Exam: normal (repeat Q3mos until seronegative)
- \*Long Bone Scan with early changes of congenital syphilis
  - bone destruction occurs in up to 60%-80% of cases and can be sole manifestation<sup>1</sup>



- Wimberger Sign: metaphyseal destruction
- Periosteal reaction



1. Koliou, M., Chatzicharalampous, E., Charalambous, M., & Aristeidou, K. (2022). Congenital syphilis as the cause of multiple bone fractures in a young infant case report. *BMC Pediatrics*, 22, 728. <https://doi.org/10.1186/s12887-022-03789-y>

2. Stephens, J. R., & Arenth, J. (2015). Wimberger Sign in Congenital Syphilis. *The Journal of Pediatrics*, 167(6), 1451. <https://doi.org/10.1016/j.jpeds.2015.09.005>



# Case #2

Gestational Age: 37w3d at birth

Reason for Admission: Newborn Infant

Maternal History: 31 yo G5P3023 with pregnancy notable for maternal substance use and limited prenatal care. Mom intoxicated on arrival.

- Treponemal syphilis test + in pregnancy and at delivery
- RPR negative x2
- Repeat treponemal test negative (different test than 1st) x2
- The state investigators find evidence of a prior known syphilis infection with documented treatment prior to pregnancy.

Delivery: SVD

# Case #2

## What does the baby need?

Mom had an RPR – we need an RPR.

- RPR NEGATIVE

Whoops! A treponemal antibody test was done on this baby too.

- FTA-ABS is POSITIVE

Do we need any other workup? No

Should we consult ID? We did.

No treatment or further workup recommended. Follow-up at Q3mos with repeat RPR until 6 mos of age.

# Case #2

## If this baby were to have syphilis, how would it present?

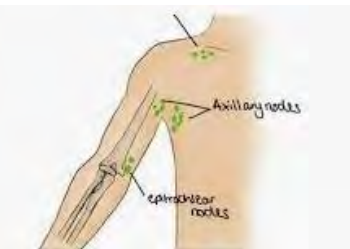
Early: symptoms before 2 years of age

- usually within the first 5 weeks after birth and by 3 mos
- **hepatomegaly** and bony abnormalities most common
- NO isolated splenomegaly (compared to other TORCH)
- bony abnormalities present at birth/early – heal by 6 mos

Late: after 2 years of age – at risk if not treated even if asymptomatic at birth

# Early vs Late Congenital Syphilis

Early: symptoms before 2 years of age



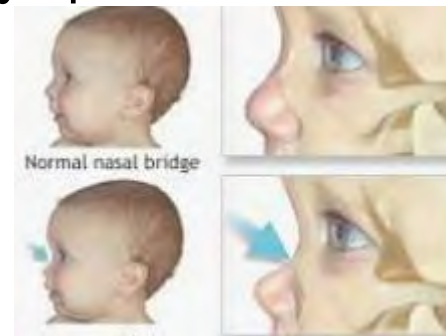
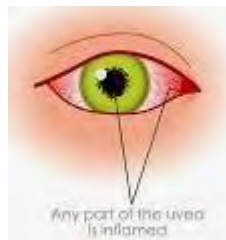
Snuffles/rhinitis



Late: after 2 years of age – at risk if not treated even if asymptomatic at birth



Deafness



Molly Crimmins Easterlin, Rangasamy Ramanathan, Theodore De Beritto; Maternal-to-Fetal Transmission of Syphilis and Congenital Syphilis. *Neoreviews* September 2021; 22 (9): e585–e599. <https://doi.org/10.1542/neo.22-9-e585>

# Case #3

Gestational Age: 38w5d at birth

Reason for Admission: Newborn Infant

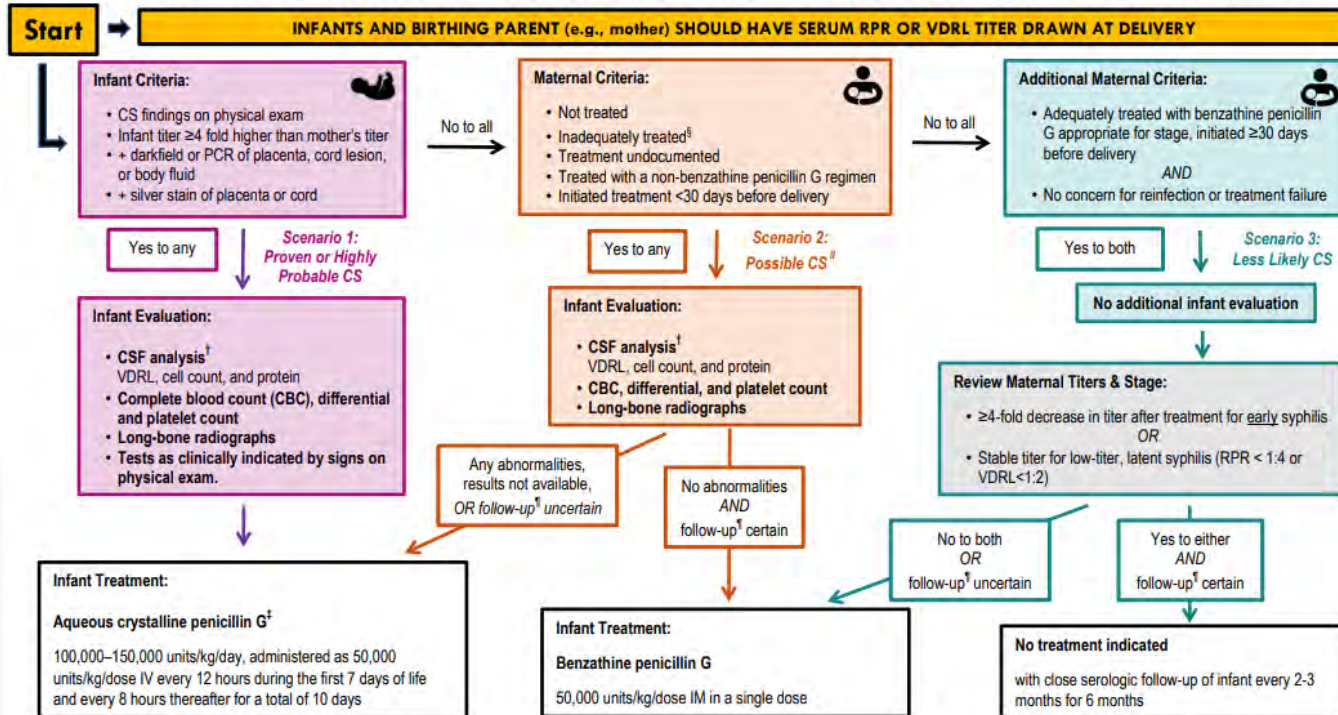
Maternal History: 18 yo G1P1001 with pregnancy notable for syphilis infection and delayed prenatal care.

- VDRL + at 2nd trimester screen at 28 weeks.
- FTA-ABS +
- Completed treatment with benzathine penicillin G.

Delivery: SVD

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‡ Alternative: Procaine penicillin G 50,000 units/kg/dose IM in a single daily dose for 10 days.

§ Benzathine Penicillin G (BPG or Bicillin-LA), administered according to stage of disease and initiated at least 30 days prior to delivery is the only adequate treatment for syphilis during pregnancy.

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# Case #3

Was appropriate treatment initiated greater than or equal to 30 days prior to birth? Yes

Does mom need another test?

-Yes - at delivery.

What test? VDRL

-VDRL with 2-fold decrease in titers

Does baby need a syphilis test?

-Yes, VDRL. VDRL obtained and negative. Physical exam normal.

Does the infant need further workup? No

Does the infant need treatment?

-Yes, a single benzathine penicillin G 50,000 units/kg IM (because mom's titers didn't drop 4-fold)

Does the infant need ID consultation and follow-up?

-Yes, follow-up VDRL titer at 3 mos

## Case #4

Doesn't exist because we eradicated syphilis in the only known host – humans.

We identified it.

We appropriately treated it.

We eradicated it.



**THANK YOU FOR YOUR TIME!**



**University of Nebraska  
Medical Center™**



**Children's  
NEBRASKA**

# References

- Molly Crimmins Easterlin, Rangasamy Ramanathan, Theodore De Beritto; Maternal-to-Fetal Transmission of Syphilis and Congenital Syphilis. *Neoreviews* September 2021; 22 (9): e585–e599. <https://doi.org/10.1542/neo.22-9-e585>
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