



WHO WE ARE - THE ECHO HUB TEAM







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CHAGAS DISEASE 4-PART SERIES

Today's Session: Congenital and Pediatric Chagas Disease in the USA

Upcoming Sessions:

- February 7, 2024 Chagas Disease as a Migrant Health Issue
- **March 6, 2024** Interprofessional Team Approaches to Chagas Disease Management

Past Session:

December 6, 2023 - Chagas Disease in the USA: Screening, Diagnosis, and Treatment for Primary Care Clinicians

1.5 CME/CNE credit available for each session for total **6.0** credits for entire series provided by The American Academy of Family Physicians (AAFP)





PRESENTER



Dr. Morven S. Edwards, MD

 Dr. Edwards is an Attending Physician in the Division of Infectious Diseases at Texas Children's Hospital and Professor of Pediatrics at Baylor College of Medicine. Her research interests include group B streptococcal infections in infants, children and adults, and congenital Chagas disease.

PRESENTER



Nancy Piper Jenks, FNP

- Nancy Piper Jenks, MS, CFNP,MFTM RCPS (Glas), FAANP is a primary care practitioner in internal medicine at Sun River Health. She is a specialist in travel medicine, with a particular interest in tropical diseases and immigrant health.
- She has been the GeoSentinel site director at SRH for over 20 years, tracking infectious diseases in migrant populations. She has published in the peer-reviewed medical literature on topics that include Hepatitis E in travelers, enteric fever, strongyloidiasis in migrants, MRSA and Lyme disease in immigrant populations.

DISCUSSION FACILITATOR



Dr. Paula Stigler Granados, PhD

- Dr. Paula Stigler Granados is an Associate Professor in the School of Public Health and Division Head of the Environmental Health Division.
- She is a subject matter expert in Chagas disease and has been the PI for the last 8 years on a Center for Disease Control funded cooperative agreement award to raise awareness among healthcare providers in the U.S. about Chagas disease. She also works with the U.S. military on Chagas disease surveillance activities and helped launch the Texas Chagas Taskforce in 2015.







Congenital and Pediatric Chagas Disease in the USA

Chagas Disease ECHO Educational Series January 10, 2024



Morven S. Edwards, M.D. Professor of Pediatrics Division of Infectious Diseases Baylor College of Medicine Houston, Texas

Baylor College of Medicine

Objectives

- Cite the population at risk for congenital Chagas disease in the United States
- Describe the clinical features of congenital Chagas disease
- Understand how to establish the diagnosis of congenital Chagas disease
- Know which US children are at risk for Chagas disease

Medical News & Perspectives

Putting Chagas Disease on the US Radar Screen

Bridget M. Kuehn, MSJ

- "In the Los Angeles clinic of Sheba Meymandi, MD, about 20% of Latin American patients with heart failure can trace their illness to a cause many US physicians would never suspect: Chagas disease."
- "Chagas disease is joining an increasing list of infectious diseases such as dengue and chikungunya that are a concern in the United States."
- "It's not an exotic disease any more".

JAMA March 24/31, 2015; 313:1195.

What is Chagas Disease?

 Chagas disease is a vector-borne zoonosis with many animal reservoirs that is caused by the parasite, *Trypanosoma cruzi*. The parasite can be transmitted from mother to infant during pregnancy



- Most people who have Chagas disease live, or have lived in, Mexico, Central America or South America
- The parasite is only found in the Americas. An estimated 6 million people have Chagas disease
- Without treatment, Chagas disease is a lifelong infection. Approximately 1.2 million people have Chagas cardiomyopathy

Photo: Carlos Chagas in 1909 in his laboratory at the Instituto Oswaldo Cruz Bern C et al. *Clin Microbiol Rev* 2019;33(1):e00023-19



The triatomine bug, sometimes known as the kissing bug, is the vector for Chagas disease. The bug becomes infected after biting an animal or a person who is already infected with *T. cruzi*.

Triatomines defecate during or after taking a blood meal. A person bitten is inoculated by rubbing insect feces into the bite or on mucous membrane.



CDC Public Health Image Library









Trypanosoma cruzi amastigotes in infected heart muscle tissue

CDC DPDx-Laboratory Identification of Parasites of Public Health Concern. Available at: https://www.cdc.gov/dpdx/trypanosomiasisamerican/index.html





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Chagas Disease in the United States

- The largest group of women with Chagas disease living in the United States are immigrants from Mexico, El Salvador, Guatemala or Honduras
- An estimated 288,000 to 300,000 *T. cruzi-infected* persons live in the United States
- An estimated 43,000 women of childbearing age who have chronic Chagas disease live in the United States. Infants of these women are at risk for congenital Chagas disease
- Between 22 and 315 infants with congenital Chagas disease are born yearly in the United States

Bern C, Montgomery SP. *Clin Infect Dis* 2009; 49:e52. Irish A et al. *Emerg Infect Dis* 2022; 28:1313.







Chagoma or Romaña sign is thought to be from parasite penetration of the conjunctiva. The swelling is firm and lasts weeks

Photos from CDC

Chagas Cardiomyopathy

- Chagas heart disease results from chronic inflammation of the heart chambers and damage to the conduction system
- The pathogenesis is thought to involve parasite persistence in cardiac tissue and immune-mediated myocardial injury
- Early manifestations include conduction system abnormalities and segmental left ventricular wall motion abnormalities
- Later findings can include ventricular tachycardia, atrioventricular block or apical aneurysm with risk of sudden death

Photo by Dr. Anis Rassi, Jr.

Chagas Gastrointestinal Disease

Digestive Chagas disease is thought to be caused by parasitic damage to intramural neurons. The effects on the esophagus range from motility disorders to severe megaesophagus.



Photos by Dr. Anis Rassi Jr.

Involvement of the colon can cause constipation, abdominal pain and fecaloma.



Gastrointestinal Chagas disease occurs predominantly in patients infected in Argentina, Bolivia, Chile, Paraguay, Uruguay, and southern Brazil. This pattern is likely linked to differences in predominant *T. cruzi* genotypes.

Modes of Transmission

- Vector-borne: Contact with an infected triatomine bug is the most common mode of transmission
- Bloodborne: Contaminated blood products, organs or tissue
- Food or waterborne: In endemic regions, drinking water contaminated with triatomine bug feces or eating contaminated foods
- Laboratory accidents: Rare mode of transmission
- Congenital: Mothers with Chagas disease can transmit infection to their infants. An estimated 23% of infections occur through congenital transmission

Bern C et al. *Clin Microbiol Rev* 2019; 33:e00023-19. Photo: *Trypanosoma cruzi* parasite in a thin blood smear. CDC photo.



Mother-to-Child Transmission of T. cruzi

- Transmission occurs transplacentally in the 2nd or 3rd trimester of gestation. There is little evidence to suggest intrapartum or postpartum transmission
- Mothers usually asymptomatic
- Mother-to-infant transmission rates are 1% to 5%
- Transmission rates are higher (5%) in countries where *T. cruzi* is endemic than in those where it is not (3%)*

*Howard et al. BJOG 2014; 121:22.

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Congenital Chagas Disease

- An estimated 43,000 infected women of childbearing age live in the United States; an estimated 22-315 infected infants are born each year*
- Most congenitally infected infants appear at healthy at birth; untreated, they are at risk for developing life-threatening cardiac or GI disease decades later
- 10% to 40% of infants have clinical signs at birth with findings that can include prematurity, hepatosplenomegaly, jaundice, anemia and thrombocytopenia; none is specific for Chagas disease

Bern & Montgomery. *Clin Infect Dis* 2009; 49:e52. Beukens et al. *Mat Child Health* 2008; 12:283.

Congenital Chagas Disease: Initial U.S. Report

- Congenital Chagas disease in the United States was first reported in a boy born in Virginia in 2010. His mother had moved recently to the United States from Bolivia.*
- The infant was born at 29 weeks' gestation by C-section for fetal hydrops. His birth weight was 1,840 g. APGAR scores were 6 at 1 and 9 at 5 minutes. He had ascites and pleural and pericardial effusions
- Blood smear in week 2 of life revealed *T. cruzi* trypomastigotes and *T. cruzi* PCR was strongly positive; serologic tests for *T. cruzi* antibodies were positive
- He received benznidazole for 60 days and was cured

*CDC. Congenital transmission of Chagas disease- Virginia, 2010. MMWR 2012; 61:477.

Feature	Frequency of Finding
Low birth weight (<2500 g)	++++
Prematurity	++
Respiratory distress	+++
Hepatomegaly	++++
Splenomegaly	+++
Sepsis	++
Cardiomegaly/heart failure	++
Myocarditis	++
Cardiac arrhythmia	++
Meningoencephalitis	++
Neurologic signs	++
Edema/anasarca	++
Petechiae	++
Anemia	+

Congenital Chagas Disease Differential Diagnosis

- Congenital Chagas disease can mimic congenital infections such as syphilis, cytomegalovirus (CMV) and toxoplasmosis or fetal hydrops from intrauterine parvovirus B19 infection
- Prematurity or low birth weight, hepatosplenomegaly and petechiae are common in infants with congenital syphilis or CMV
- Congenital Chagas disease can mimic noninfectious conditions including congestive heart failure or respiratory distress syndrome

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Pregnancy-Based Screening for Chagas Disease Benefits Mother and Infant

- Pregnancy is the optimal access point for identifying Chagas disease at risk family units because delivery is the most likely time for contact with the healthcare system
- Women at risk have migrated from an endemic region
 - Risk is enhanced by having lived in a rural region
 - Having lived in a mud or thatched-roof home also increases risk
- Women who have visited and lived in an endemic region for 6 months or longer are also at risk
- Women at risk for Chagas disease should have pregnancy-based screening for *T. cruzi* IgG





Photo from WHO at http://www.who.int/chagas/resources/photo_gallery/en/



Chagas Disease Positive Persons in Los Angeles County, 2008-2014



Meymandi SK et al. Clin Infect Dis 2017; 64:1182.

Among 4,755 Latin American-born residents of Los Angeles County, 59 had Chagas disease for an overall prevalence of 1.24%.

Prevalence was highest among Salvadorans (3.45%) and, among those born in Mexico, from the states of Oaxaca (4.65%) and Zacatecas (2.2%).

>30,000 people living in Los Angeles county may have Chagas disease.

Challenges to Identifying Mothers and Infants with Chagas Disease

- Identifying maternal infection is key but mothers have no symptoms
- None of the findings in infant infection are specific for Chagas disease
- The diagnosis must be considered
- Screening during pregnancy or at delivery is key to identifying women and infants at risk but maternal screening is not standard of care
- The prevalence of infection among women of child-bearing age in the US is not known

Screening for Chagas Disease during Pregnancy

- Screening for Chagas disease can be performed during any trimester
- A commercially-available ELISA should be ordered to test for Trypanosoma cruzi IgG
- Chagas disease screening is a send-out test from most hospital laboratories. Results are available within days
- Chagas screening can be included with routine maternal screening
- It is not necessary or appropriate to screen for T. cruzi IgM

Pregnancy-Based Screening for Chagas Disease is Cost-Saving

- Pregnancy-based screening has the advantage that results are known at delivery. Screening at admission for delivery or screening of neonates are alternative approaches
- At current costs, targeted screening, including the cost of treatment, would result in savings of \$1,314 per birth and \$670 million in lifetime savings per birth-year cohort
- Universal screening and treatment would also be cost-saving

Perez-Zetune V et al. Am J Trop Med Hyg 2020; 102;1086

Evaluation for Suspected Congenital Chagas Disease

- Direct detection: Diagnostic if positive; less sensitive than PCR
- PCR: The most sensitive test for early diagnosis
 - PCR for *T. cruzi* is available at the CDC; testing is under CLIA
 - Initial negative should be repeated at 1 month of age as parasites multiply in the first weeks of life
- Maternal Serology: Order *T. cruzi* IgG if not performed during pregnancy
- Infant Serology: If PCR is negative and maternal serology is positive, follow infant's *T. cruzi* IgG. Negative serology at 9-12 months of age excludes congenital infection



Treatment of Chagas Disease

- Treatment is always indicated for congenital Chagas disease. Treatment early in life kills the parasite and prevents long-term complications from heart and intestinal disease; cure rates exceed 90%*
- Treatment is always indicated for women in the childbearing years**, both for the health of the woman and for the sake of her children
- Infection can be transmitted congenitally in sequential pregnancies among women chronically infected with *T. cruzi*

* *MMWR* 2012; 61:477-9. **Bern C. Antitrypanosomal therapy for chronic Chagas' disease. *N Engl J Med* 2011; 364:2527.



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Epidemiology of congenital Chagas disease 6 years after implementation of a public health surveillance system, Catalonia, 2010 to 2015



33,469 pregnant women from endemic countries and those who had lived in a rural area of an endemic country were screened

The overall prevalence of maternal Chagas disease was 2.8%. Rates were highest in women from Bolivia (15.8%), El Salvador (1.4%) and Paraguay (1.2%)

28 infants were diagnosed with congenital Chagas disease (transmission rate 4.2%)

Basile L et al. Euro Surveill. 2019;24(26):pii=1900011. https://doi.org/10.2807/1560-7917.ES.2019.24.26.19-00011

Epidemiology of congenital Chagas disease 6 years after implementation of a public health surveillance system, Catalonia, 2010 to 2015

- Children born to *T. cruzi*-positive women before the current pregnancy were screened
- Among 178 children, 14 (7.9%) were diagnosed with Chagas disease. The children ranged in age from 3-18 (median 10) years of age
- Siblings of an index case of Chagas disease should also undergo screening

Basile L et al. Euro Surveill. 2019;24(26):pii=1900011. https://doi.org/10.2807/1560-7917.ES.2019.24.26.19-00011

Evaluation of Family Members

- If a mother is diagnosed with Chagas disease, her other children should undergo serologic testing; treatment is always indicated for children <18 years of age
- Serologic testing of family members of an infant with congenital Chagas disease should also include:
 - The maternal grandmother
 - The mothers siblings

Chagas Disease Prevention

Chagas disease fact sheets for the public are available on-line in English and Spanish through CDC

Other printable resources include, "What happens to blood donors who test positive for Chagas disease?" and, "Chagas disease fact sheet for the public"

www.cdc.gov/parasites/chagas/printresources.html



Chagas Disease Prevention

Chagas disease printable resources are available through CDC

This poster is available without charge at this website, laminated with English on one side, Spanish on the other:

https://www.cdc.gov/parasites/chagas/printresources.html

Protect Your Baby From Chagas Disease

A COC

Chagas disease is an illness that can lead to serious heart and stomach problems, and even death.

Most people get Chagas disease from a bug. Mothers who have Chagas disease can give it to their unborn babies.

You could be at risk for Chagas disease if you have:

Lived in rural areas of Mexico,
Central America, or South America

Seen this bug
Stayed in a house with walls that have

cracks or crevices

If you think you may have Chagas disease, talk to your doctor. He or she can help you get tested and if you or your baby has Chagas disease, you both can get treated.

For more information on Chagas disease, please vis www.cdc.gov/parasites/chagas or call 404.718.4745

Acknowledgements

I am grateful to Susan P. Montgomery, DVM, MPH, epidemiology team lead in the CDC's parasitic disease branch for providing some of the slides in this presentation and for her example as a mentor and teacher.

Selected References

Bern C et al. Chagas disease in the United States: a public health approach. *Clin Microbiol Rev* 2019; 33(1):e00023-19. Comprehensive and current review.

Centers for Disease Control and Prevention. Access at: <u>http://www.cdc.gov/parasites/chagas/index.html</u> Excellent source for general information and diagnosis and treatment information.

Edwards MS et al. Evaluation and management of congenital Chagas disease in the United States. Provides algorithms for evaluation of mothers and infants. *J Pediatr Infect Dis Soc* 2019; 8:461.

Stillwaggon E et al. Congenital Chagas disease in the United States: Cost savings through maternal screening. *Am J Trop Med Hyg* 2018; 98:1733. Maternal screening is cost saving, whether targeted to at-risk pregnant women of performed universally.





Sun River Health



From Left to Right: Willie Mae Jackson, Pearl Woods, Rev. Jeannette Phillips, Anne K. Nolon, Mary Woods.

In the early 1970's a group of residents and religious leaders addressed the lack of appropriate health services in their community. In particular, a group of four women, fondly referred to as our founding mothers, spearheaded the efforts and have remained committed to the organization since its inception.







GeoSentinel Data from Sun River Health 2001-2022

Chagas Disease – one case in Mexican immigrant who donated blood and was then informed of infection, came to us for treatment, significant heart disease

Quality Improvement Project

Presented to our QI council and approved January 2023

Importance of identifying and treating Chagas in our at-risk patients Framing initiative as quality improvement, not research





Countries of Origin OB Patients SRH Brentwood Site, Long Island, NY

Informatic system review 1/9/2023 • El Salvador 63 (59%) • Honduras 17 (16%) • Ecuador (8.4%) 9 Colombia 6 (5.6%) • DR 3 (2.8%) 2 (1.8%) • Peru 2 (1.8%) • Mexico 2 (1.8%) • Chile (2.8%) 3 • USA Sun River Health

Our Team

Sun River Health

Nancy Piper Jenks, MS, CFNP, MFTM RCPS (Glasg), FAANP Quratulain Zeeshan, MD, Quality Medical Director WH Aarathi Nagaraja, MD, CPH, Medical Director HIV/Hep C Vasanthi Arumugam, MD, Infectious Diseases Marianne Boyce, RN, VP Infection Prevention, Lab Services Roberta Kelly, FNP-BC, CIC, Chief Nursing Officer, SVP Carlos Ortiz, Deputy Chief Operating Officer Amanda Ascher, MD, CMO Rachel Barnett, Program Director

Organization of pilot (January 2023)

Guidance from experts

Training/educating staff

Creating protocols for screening and tracking

Working with IT to create EMR templates for Chagas screening

Organizing system to bring in family members and spouses of patients for screening





	March 2016
	23-year-old woman from Honduras presents to our health center for prenatal care
Case 1	P2011 Has an 8-year-old child
	Has been in USA for 10 months
	Sun River Health











• July 2023 Patient is screened for T Cruzi AB, sent to commercial lab

Sun River Health 🍳



1

Case 1

Commercial serology positive 7/2023 2 tubes of whole blood and serum sent to NYS DOH Wadsworth 10/2023

<u>Test</u> T. cruzi AB EIA*

Comments and Disclaimers • Reference value: Non-reactive/negative

<u>Test</u> T. cruzi AB IB (TESA)*

Comments and Disclaimers * Reference value: Non-reactive/negative

<u>Test</u> T. cruzi Interpretation

Comments and Disclaimers

Comments and Disclaimers • The test results are considered **Positive** and are consistent with T. cruzi infection. Diagnosis of chronic Chagas disease is established after concordant positive results are obtained with at least two different types of T. cruzi serologic assays. Clinical consultation is available at CDC (phone: 404-718-4745; parasites@cdc.gov) for interpretation of these serologic results and other diagnostic indicators, in the context of additional Information.

<u>Interpretation</u>

Weak Positive

Reactive

<u>Result</u>

<u>Result</u>

Positive*

T. cruzi AB EIA Test Comments: This test detects antibodies against T. cruzi recombinant antigens.

T. cruzi AB IB (TESA) Test Comments: This test is an immunoblot assay which detects antibody reactivity to the 150-160kDa transialidase antigens of T. cruzi.

The performance characteristics of the T. cruzi AB IB (TESA) and T. cruzi AB IFA tests were determined by validations performed at the Division of Parasitic Diseases and Malaria (DPDM) at the Centers for Disease Control and Prevention. They have not been cleared or approved by the U.S. Food and Drug Administration.





















2					
Whole blood and serum sent to Wadsworth		Kinatoplastid Identification No parasites observed T. cnuzi (Mc) DNA by RT-PCR (*): T. cnuzi (sDNA) DNA by RT-PCR (*): Leishmania sp. (tAP3) DNA by RT-PCR (*): Leishmania sp. (AP3) DNA by RT-PCR (*):	Not Detected Not Detected Not Detected Not Detected Not Detected		10/ 2023 10/ 2023 10/ 2023 10/ 2023 10/ 2023 10/ 2023
		Specimen Id: IDR2300061301-02 Diagnostic Immunology Laboratory Phone: (518) 485-3845 Fay: (518) 486-7971	Spec	imen Type: Serum	
		Charge (T. cruzi) Disease Serology			
	new	Suggests evidence of infection at an unde	termined time.		101 /2023
		T. cruzi total Antibody ELISA			
	new	IDR2300061301-02 collected 10/19/23 Result:	Reactive		10/ /2023
		T. cruzi IgG ELISA			
	new	IDR2300061301-02 collected 10/19/23 Result:	Positive		10 //2023
		Diagnosis of chronic Chagas disease is establish of T. cruzi serologic assays. For more information please see: https://www.cdo	ed after concordant positive results .gov/parasites/chagas/health_pro!	s are obtained with at least two different ressionals/dx.html	types
				Sun River	Health



2

Assessments

- 1. Diabetes mellitus without complication E11.9
- 2. Chagas disease B57.2 (Primary)
- 3. Urinary tract infection without hematuria, site unspecified N39.0

Treatment

1. Chagas disease

Start Benznidazole Tablet, 100 MG, 2 tab, Orally, Twice a day, 30 days, 120 Tablet, Refills 1 <u>IMAGING: ECHOCARDIOGRAM</u>

IMAGING. LEHOCARDIOGRA

Notes: She is here for treatment

Start benzinidazole 200mg po q12 x 60 days

2 T. Cruzi/Chagas Disease: Screening Screening : Adult ... Counseling information The patient is from an endemic region of North, South, or Central Am and is requesting screening for T.cruzi., The patient/ parent or care giver verbalized understanding screening positive will require further evaluations, including diagnostic testing and possible treatm Country of origin/birth: El Salvador, undefined ... Region: ... Prior treatment or testing in the past? No .. Do other family members want to be screened? Yes ... **Diagnostic Evaluation and Treatment** Diagnostic evaluation and treatment for: Adult Why was the patient referred for testing: screening during pregnancy Did other family members test positive on screening or diagnostic labs: Yes ... Details: Father died of cardiomyopathy at age 50 (told T.cruzi infection) Country of origin/birth: El Salvador, undefined ... Housing: Mud, Adobe, Thatch, undefined Location of bathroom: Outside of the home, suggest strongyloides ... Does the patient or parent/care giver recognize a kissing bug? Yes ... How does the patient refer to the kissing bug ? El bicho besador Does the patient or parent/care giver recall family with Chagas disease or heart disease in their country of origin? Yes ... Details: Father and uncle died of cardiomyopathy and told because of T.Cruzi infection Is the patient currently pregnant or recently delivered ? Recently Delivered ... Date of delivery: 07/2023 Mode of delivery: C-Section The patient was counseled that the recently delivered child needs to be tested. Yes Lab Results: Bio-Reference: T. Cruzi Screening Date: 10/2023... Sun River Health 🤗

Sun River Health

	Start	Start small with screening initiatives in order to organize process
Take home points	Consider	Consider costs associated with screening, prenatal and pediatric patients have benefit of insurance
	Take	Take advantage of tremendous resource networks and individuals to support Chagas screening and treatment
	LOOK	LOOK for Chagas

