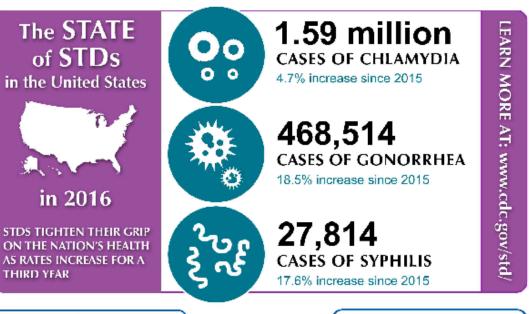


STI-HIV "Hot Topics" 2018

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Departments of Medicine, Microbiology, and Epidemiology
University of Alabama at Birmingham



Anyone who has sex is at risk, but some groups are more affected

- YOUNG PEOPLE AGED 15-24
- GAY & BISEXUAL MEN
- PREGNANT WOMEN

LEFT UNTREATED, STDS CAN CAUSE:



INCREASED RISK OF GIVING OR GETTING HIV



LONG-TERM
PELVIC/ABDOMINAL PAIN



INABILITY TO GET PREGNANT OR PREGNANCY COMPLICATIONS

HELP INTERRUPT THE STEADY CLIMB IN STDS WITH THESE THREE STEPS:



TEST II

Get tested. It's the only way
to know if you have an STD.

If you have an STD, work with your provider to get the right medicine.





Hot(est) Topics

- 1. Embracing Complexity / Rising STD Rates —Tensions between HIV Management & Prevention and STIs.
- 2. Syphilis-HIV

3. STI Pathogen Antimicrobial Resistance and Therapeutic Limitations

Other "Hot Topics" Not Discussed Today

- 1. Sexual transmission of Hepatitis C
- 2. Mycoplasma genitalium
- 3. Bacterial vaginosis as a dysbiosis
- 4. Diagnostics
 Syndromic
 Point of Care
 Multiplexed
- 5. Vaccines for STD Prevention Hepatitis B and A HPV In Development
- 5. Sexual transmission of Zika and Ebola

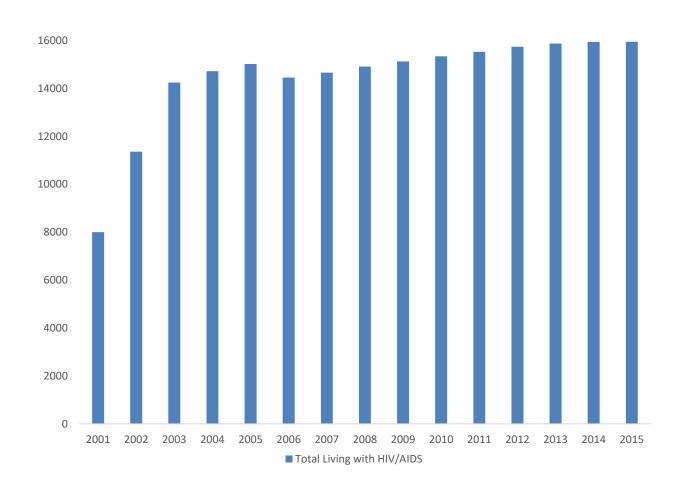


Hot(est) Topics

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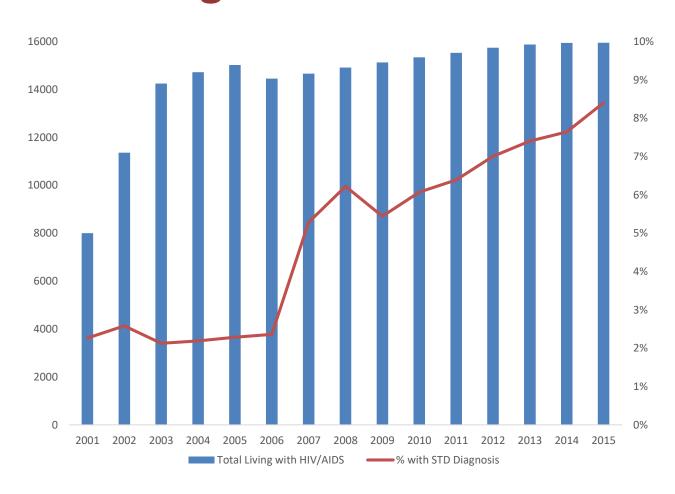
3. STI Pathogen Antimicrobial Resistance and Therapeutic Limitations

More People Living with HIV



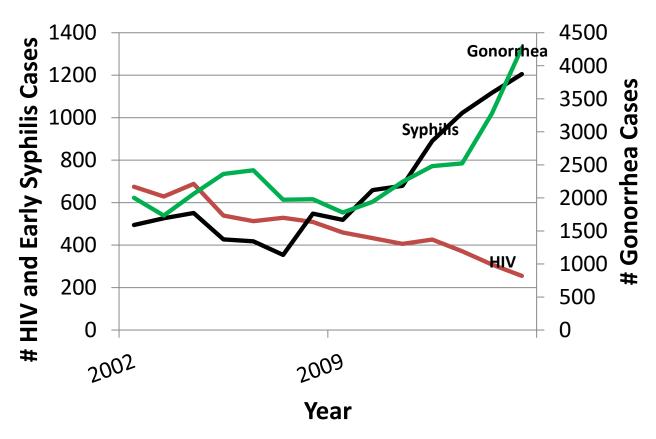
Courtesy of Susan Phillip and the SFDPH Population Health Division, Applied Research Community Health Epidemiology and Surveillance Branch

More People Living with HIV are Being Diagnosed with an STD



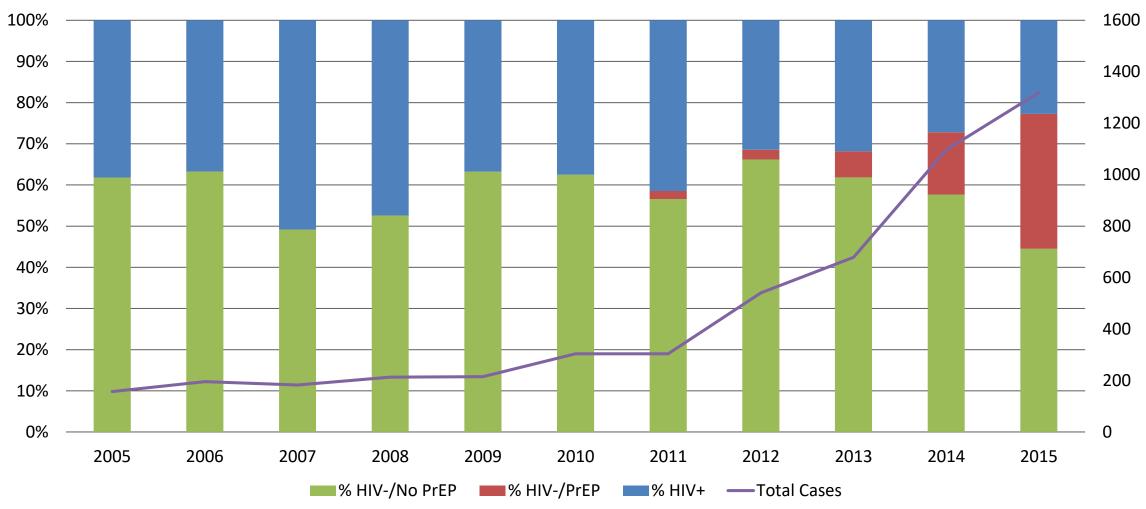
Courtesy of Susan Phillip and the SFDPH Population Health Division, Applied Research Community Health Epidemiology and Surveillance Branch

San Francisco: STD Increasing while HIV Diagnoses Decline



Courtesy of Susan Phillip and the SFDPH Population Health Division, Applied Research Community Health Epidemiology and Surveillance Branch

Frequency of Any Bacterial STI infection by HIV Status and PrEP Use among Male Patients, Fenway Health





Hot(est) Topics

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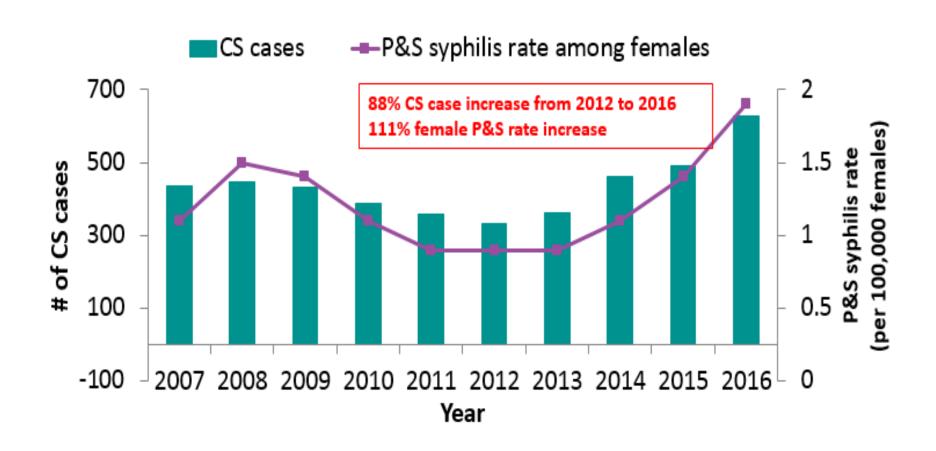
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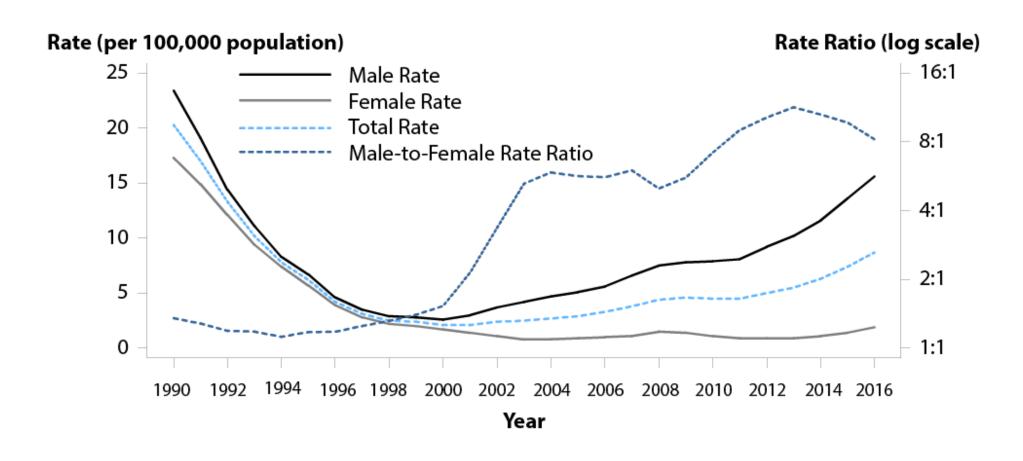
Syphilis: Global Impact

- 1. Global Perspective
 - a. 17-20 million cases
 - b. Most prevalent on the margins of society
 - c. Leading preventable cause of adverse pregnancy outcomes
- 2. North America/Western Europe
 - a. Recurring epidemics
 - b. Critical consideration GUD diagnosis
 - c. A diagnostic challenge- protean manifestations
 - d. A driving force for incident HIV
 - e. A public health embarrassment

Congenital Syphilis (CS) Cases and Rate of Primary and Secondary (P&S) Syphilis Among Females, U.S., 2007–2016

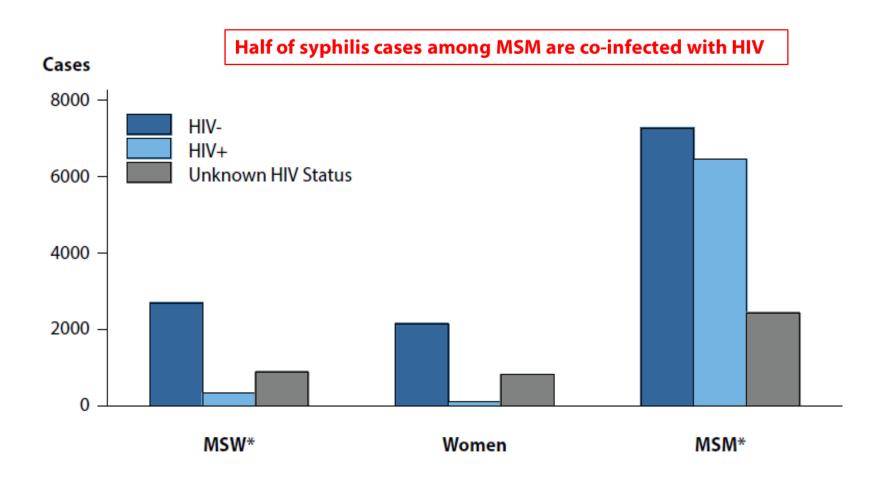


Primary and Secondary Syphilis — Reported Case Rates by Sex and Male-to-Female Rate Ratios, United States, 1990–2016





Primary and Secondary Syphilis — Reported Cases by Sex, Sexual Behavior, and HIV Status, United States, 2016



^{*} MSM = Gay, bisexual, and other men who have sex with men (collectively referred to as MSM); MSW = Men who have sex with women only.

Syphilis-HIV Interactions: Epidemiologic Considerations

P&S Syphilis 2016: 27,814 (+17%) EL Syphilis 2016: 28,173 (+20%) ~50% HIV Infected

~20% Previously Infected

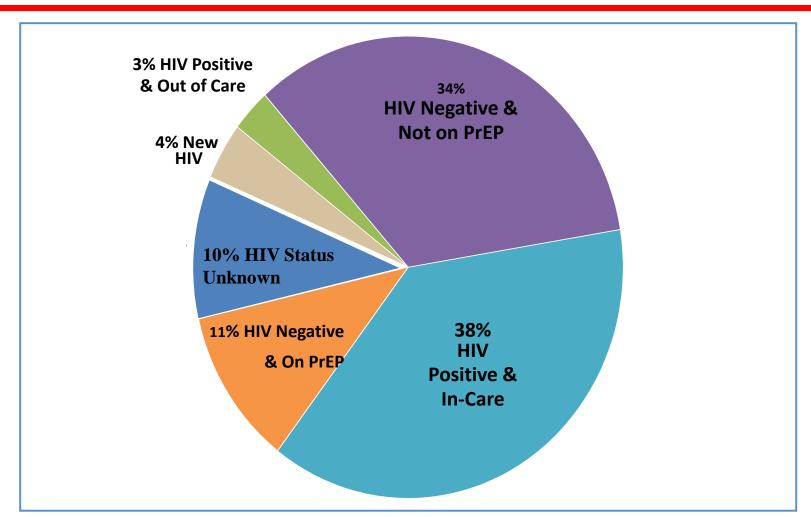
HIV Incidence Following Syphilis Diagnosis

1 Year = ~3.5%

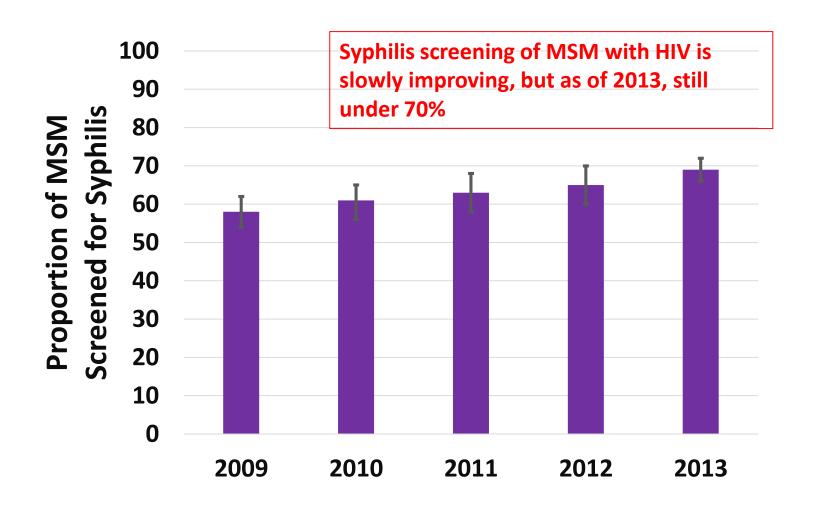
10 Years= 17.5%

- Peterman et al. Public Health Reports 2014; 129: 184-169
- Pathela et al. Clin Infect Dis 2015; 61: 281-7

HIV Prevention & Care Opportunities, California MSM, with Early Syphilis, 2017



Proportion of Sexually Active HIV-Infected MSM in Medical Care Screened for Syphilis in the Prior 12 Months, Medical Monitoring Project 2009-2013



Test for trend p<0.01

2015 CDC STD TREATMENT GUIDELINES Early Syphilis

- Recommended
- Benazthine Penicillin G, 2.4 Mu IM
 - Penicillin Allergy
- Doxycyline 100 mg PO, BID x 14d
 - Limited Data
- Ceftriaxone 1.0 g IM or IV x 8-10d
 - Azithromycin 2.0g PO



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The NEW ENGLAND JOURNAL of MEDICINE

Perspective FEBRUARY 9, 2012

The Emerging Threat of Untreatable Gonococcal Infection

Gail A. Bolan, M.D., P. Frederick Sparling, M.D., and Judith N. Wasserheit, M.D., M.P.H.

It is time to sound the alarm. During the past 3 years, the wily gonococcus has become less susceptible to our last line of antimicrobial defense, threatening our ability to cure gonorrhea and prevent severe sequelae.

Gonorrhea is the second most commonly reported communicable disease in the United States, with an estimated incidence of more than 600,000 cases annually. It disproportionately affects vulnerable populations such as minorities who are marginalized because of race, ethnic group, or sexual orientation. Unfortunately, Neisseria gonorrhoeae has always

Control and Prevention (CDC) are now limited to third-generation cephalosporins.²

But susceptibility to cephalosporins has been decreasing rapidly.³ The proportion of GISP isolates for which the minimum inhibitory concentration (MIC) of cefixime is elevated ($\geq 0.25 \mu g$ per milliliter) has increased by a factor of 17 — from 0.1% in 2006 to

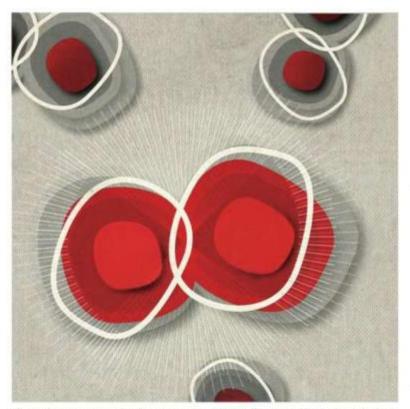
(0.04% of those in the GISP) had a MIC of ceftriaxone of 0.25 μg per milliliter in the first half of 2011, the proportion of GISP isolates with an elevated ceftriaxone MIC ($\geq 0.125 \mu g$ per milliliter) has increased by a factor of 10 since 2006 (from 0.05% to 0.50%). Again, increases were greatest in the west (from 0.04% to 1.90%) and among men who have sex with men (from 0.0% to 1.0%). These geographic and demographic patterns are worrisome because they mirror those observed during the emergence of fluoroquinolone-resistant N. gonorrhoeae.

MEDICAL DISPATCHES

SEX AND THE SUPERBUG

The rise of drug-resistant gonorrhea.

BY JEROME GROOPMAN



Gonorrhea mutates in the pharynx, making oral sex far more risky than people think.

The New Yorker, October 1, 2012

GONORRHEA THERAPY – HISTORICAL PERSPECTIVE

Previously Recommended Medications For Gonorrhea Therapy

Sulfonamides

Penicillins

Macrolides

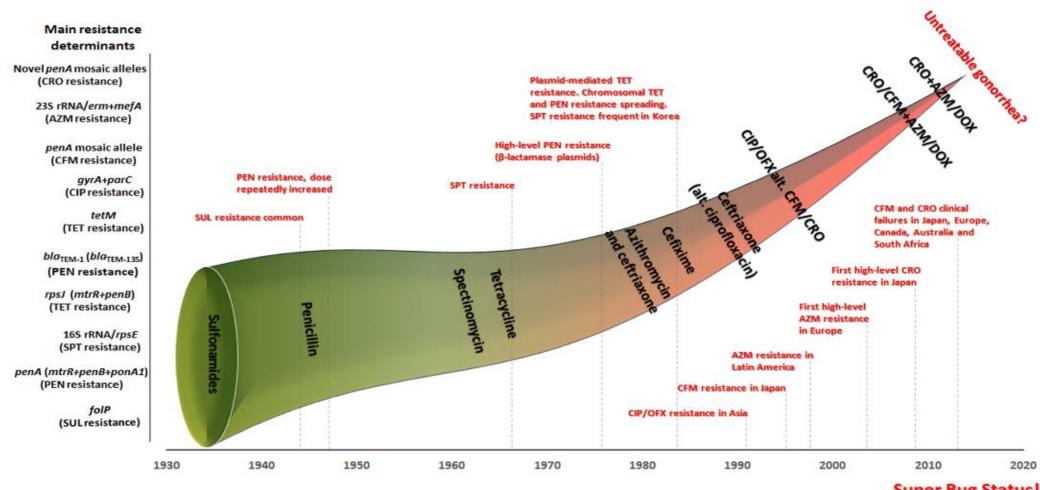
Tetracyclines

Aminoglycosides

Spectinomycin

Fluroquinolones

Gonorrhea Therapy: The Shrinking Pipeline



Super Bug Status!

Unemo & Shafer 2014. CMR

2015 CDC STD TREATMENT GUIDELINES Uncomplicated Gonorrhea

Ceftriaxone 250 mg IM

PLUS

Azithromycin 1.0 g Single Dose PO

Gonorrhea Treatment-What's Next

Salvage Therapy:

Gentamicin 240 IM/ Azithromycin 2.0g PO (IM Administration/Toxicity)
Gemifloxacin 340 mg/Azithromycin 2.0g PO (GI Toxicity)

On The Horizon:

Solithromycin

Delaflexacin

Zoliflodacin (ETX 0914)

Gepotidacin (BZT 116570)

Others

A New Model for STD Treatment Trials

Collaborative, multinational trial of Zoliflodacin, a candidate antibiotic for treatment of uncomplicated gonorrhea.

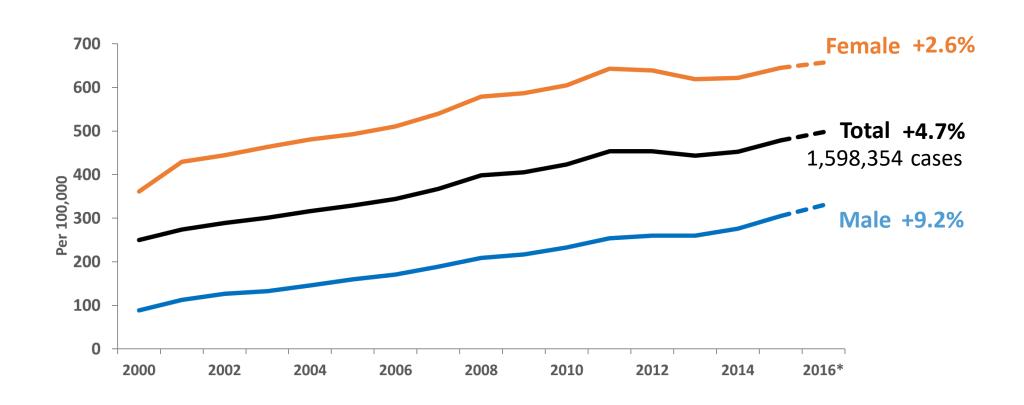
Led by GARDP (Global Antibiotic Research and Development Partnership)

Collaborators: Entasis Pharmaceutics, NIAID, WHO

Study sites in : U.S. Thailand, South Africa, E.U.

Goal: Approval of a non-beta-lactam antibiotic effective vs. resistant gonorrhea

Chlamydia — Rates of Reported Cases by Sex, United States, 2000–2016



STI Pathogen Drug Resistance and Therapeutic Limitations

- 1. Gonorrhea:
 - a. Progressive resistance, limited alternative therapeutics
 - b. Variation is therapeutic efficacy by site of infection
- 2. Chlamydia trachomatis:
 - a. Azithromycin efficacy
- 3. Syphilis:
 - a. Global penicillin shortages
 - b. Lingering question as to optimal therapy
 - i. Impact of HIV infection
 - ii. Serological response to therapy
- 4. Trichomoniasis
 - a. Optimal therapy
- 5. Bacterial vaginosis
 - a. Poor therapeutic response, high recurrence rates



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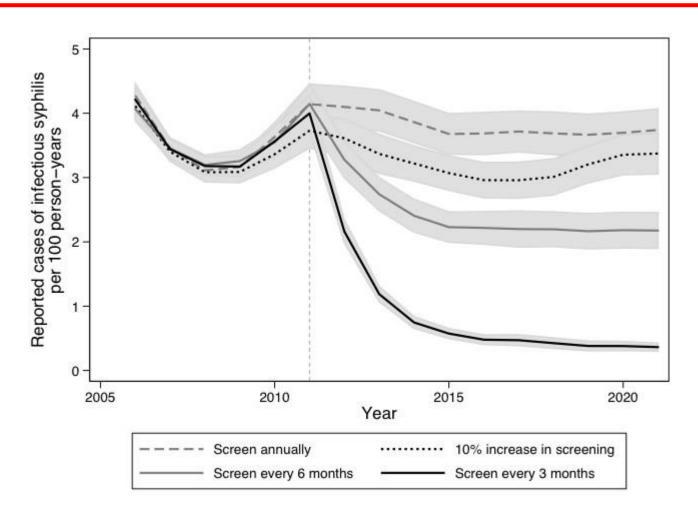
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Thank You

Impact of Screening Frequency on Syphilis Incidence



Tuite AR, Fisman DN, Mishra S. Screen more or screen more often? Using mathematical models to inform syphilis control strategies. BMC Public Health 2013;13:606.

TREATMENT OF EARLY SYPHILIS IN HIV-INFECTED AND UNINFECTED PERSONS

Proportion of Subjects with RPR Decline ≥2 Dilutions

		3 Mo.	6 Mo.	12 Mo.
Treatment (Group			
	Usual	75% (175)	76% (157)	82% (137)
	Enhanced	71% (189)	81% (172)	83% (144)
HIV-Status				
	Positive	62% (76)*	82% (69)	79% (61)
	Negative	76% (287)	81% (259)	84% (219)
	*P < 0.05			

From Rolfs et al, NEJM 1997; 44: 307-14