Sexually Transmitted Disease Treatment Tables

Federal Bureau of Prisons Clinical Practice Guidelines

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What's New in This Document?

The following are among the changes that have been made to the November 2008 version of these guidelines, in order to be current with the *Sexually Transmitted Diseases Treatment Guidelines* published by the CDC on December 12, 2010 (cited under *References*). Changes are highlighted in yellow throughout the document.

Gonorrhea (see Table 1):

- The recommended dose for ceftriaxone has been increased to 250 mg IM x1.
- Sex partners of patients with gonorrhea should be evaluated and treated for syphilis and HIV, in addition to *N. gonorrhoeae* and *C. trachomatis*.

Chlamydia (see <u>Table 1</u>):

• The BOP practice of routine intake screening for chlamydia in all females age 25 and under, *regardless of risk factors*, is clarified.

Nongonococcal Urethritis (see *Table 1*):

• It is recommended that azithromycin be given DOT.

Pelvic Inflammatory Disease (PID) (see *Table 1*):

- The dosage for doxycycline has been changed.
- Abundance of WBC on saline microscopy is added as evidence to support a PID diagnosis.
- More conditions indicating hospitalization have been added.

Herpes Simplex Virus (HSV) (see <u>Table 2</u>):

• In the case of pregnancy, consultation with an obstetrician *or* an infectious disease specialist is recommended.

Vaginitis (see <u>Table 2</u>):

- For Bacterial Vaginosis (BV), Metronidazole gel 0.75% and clindamycin cream 2% have been added as
 possible treatments.
- For Candidiasis, treatment with Clotrimazole tablets has been replaced by treatment with one of the formulary creams: clotrimazole vaginal cream 1% or miconazole vaginal cream 2%.

Syphilis – Primary/Secondary (see *Table 4*):

Tetracycline has been added as a treatment for patients with a penicillin allergy.

Syphilis - Latent:

- For patients with a penicillin allergy, treatment with doxycycline has been extended to 28 days. These patients can also be treated with tetracycline (see <u>Table 4</u>).
- The criteria for case classification for latent syphilis and the clinical description of late latent syphilis have been clarified (see *Attachment 1*).

Neurosyphilis and Syphilitic Eye Involvement (see <u>Table 4</u>):

- The standard treatments now include an approach with continuous infusion of aqueous crystalline penicillin.
- For patients with a penicillin allergy, dosing for ceftriaxone as a possible alternative is specified.

Syphilis Stages and Classification

• This information is now contained in *Attachment 1*.

Algorithms for Diagnostic Assessment and Management of Syndromes

• Seven recommended algorithms for diagnosing and managing STD-related syndromes have been posted on the BOP Clinical Practice Guidelines Web page. For more information, see the <u>References</u> page.

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Table 1. Gonorrhea, Chlamydia, Nongonococcal Urethritis, and PID

Treatment (directly observed) Comments Gonorrhea (GC) (N. gonorrhoeae) of the cervix, urethra, rectum, pharynx → Ceftriaxone 250 mg IM x1 Screening: No routine screening at intake unless symptoms of gonorrhea are present, or unless syphilis or Alternative treatments: chlamydia have been diagnosed. • Cefixime 400 mg orally x1 or Diagnosis: Nucleic acid amplification tests (NAAT) for Single-dose injectable cephalosporins. See CDC symptomatic inmates. Follow-up tests to prove cure is not guidelines at: http://www.cdc.gov/std/treatment/. indicated unless symptoms persist. PLUS, treat for chlamydia infection unless chlamydial Contacts: Sex partners of patients with N. gonorrhoeae infection is ruled out with sensitive test-nucleic acid infections, whose last sexual contact with the patient was amplification test (NAAT): within 60 days before onset of symptoms or diagnosis, • Azithromycin 1 g orally x1 or should be evaluated and treated for N. gonorrhoeae, • Doxycycline 100 mg orally twice daily for 7 days C. trachomatis, syphilis, and HIV. Pharyngeal gonorrhea: Treat with ceftriaxone. Note: Ceftriaxone is available in 250 mg, 500mg, and 1and 2-gram vials. Recommendation: 250 mg IM. HIV co-infection: Treatment regimens are the same as for inmates without HIV co-infection. Note: Quinolones are no longer recommended for treatment of gonorrhea. **Pregnant women:** Treat GC with ceftriaxone or cefixime: treat chlamydia with azithromycin or amoxicillin Chlamydia (C. trachomatis) → Azithromycin 1 g orally x1 or Screening: Routine intake screening for females who: → Doxycycline 100 mg orally twice daily for 7 days • Are age 25 and under, regardless of risk factors, or • Are older than age 25 with risk factors (e.g., have new or Alternative treatments: See CDC guidelines at: multiple sex partners) and/or http://www.cdc.gov/std/treatment/ • Have HIV infection and/or Pregnant women: • Have history of syphilis, gonorrhea, or chlamydia • Azithromycin 1 g orally x1 or Diagnosis: For symptomatic inmates, confirm infection by Amoxicillin 500 mg orally three times daily for 7 days. culture, NAAT, or other assay whenever feasible. Asymptomatic infection is common in men and women. Testing for cure is not indicated following treatment with azithromycin or doxycycline unless symptoms recur. Contacts: All sex partners in the 60 days preceding symptom onset should be evaluated and treated. Most recent sexual contact should be evaluated and treated, even if contact was >60 days before symptom onset. **Nongonococcal Urethritis** → Azithromycin 1 g orally x1 or Diagnosis: All patients with confirmed or suspected → Doxycycline 100 mg orally twice daily for 7 days urethritis should be tested for gonorrhea and chlamydia. Treatment: M. genitalium may respond better to Alternative treatments: See CDC guidelines at: azithromycin. DOT dose. http://www.cdc.gov/std/treatment/. Contacts: Refer all sex partners within preceding 60 days for treatment. Pelvic Inflammatory Disease (PID) - Outpatient Management Recommended regimen: Diagnosis: Minimum criteria based on pelvic exam finding: → Ceftriaxone 250 mg IM plus cervical motion tenderness or uterine tenderness or adnexal tenderness. The following additional evidence supports a → Doxycycline 100 mg orally twice daily for 14 days PID diagnosis: temperature >101°F, abnormal cervical or Alternative regimen: vaginal mucopurulent discharge, elevated erythrocyte • Cefoxitin 2 g IM x 1 or other third-generation sedimentation rate, elevated C-reactive protein, laboratory documentation of cervical infection with N. gonorrhoeae, cephalosporin plus • Probenecid 1 g orally x 1 plus

Note: Both regimens can be given with or without metronidazole 500 mg orally twice daily for 14 days.

Doxycycline 100 mg orally twice daily for 14 days

C. trachomatis, or abundance of WBC on saline microscopy. Hospitalize: If surgical emergencies cannot be excluded; or

if pregnancy, lack of clinical response to oral therapy, severe illness & unable to follow/tolerate an outpatient oral regimen or has tube-ovarian abscess.

Table 2. Herpes Simplex Virus, Vaginitis, and Genital Warts

Treatment	Comments		
Herpes Simplex Virus (HSV)		
First episode: Acyclovir 400 mg orally	General: Genital herpes is a recurrent, lifelong infection. Sexual transmission of HSV occurs in asymptomatic persons.		
three times daily for 7–10 days	Treatment: First episode: Treat with acyclovir, since treatment may reduce symptoms. Treatment does not eradicate herpes virus or affect the risk, severity, or frequency of recurrences.		
Recurrent episodes: Acyclovir 400 mg orally three times daily for 5 days	Recurrent episodes: Treatment must be initiated within one day of lesion onset or during the prodrome that precedes some outbreaks. Suppressive therapy: Consider on case-by-case basis, depending on the severity and		
Suppressive therapy: Acyclovir 400 mg orally twice daily for 1 year	frequency of recurrences and, upon release, returning to sex partner. Reconsider continuation of suppressive therapy after 1 year of therapy. Suppressive therapy does not eliminate asymptomatic viral shedding.		
	Note: Topical acyclovir is ineffective. Immunosuppression: Inmates with HIV infection or immunocompromised conditions may require higher doses of oral acyclovir or intravenous therapy for herpes infections.		
V	Pregnancy: Consult with obstetrician or infectious disease specialist.		
Vaginitis			
Bacterial Vaginosis (BV)			
Metronidazole 500 mg orally twice daily for 7 days or	Diagnosis: Clinical criteria require three of the following: 1) Homogenous, thin, white discharge smoothly coating vaginal wall 2) Presence of clue cells on microscopic exam 2) Public transition of this coating this coating vaginal wall		
Metronidazole gel 0.75% 5 g intravaginally daily for 5 days	 3) pH of vaginal fluid >4.5 4) Fishy odor of vaginal discharge before or after addition of 10% KOH ("whiff test") Pregnancy: BV is associated with adverse pregnancy outcomes. Test if symptomatic. 		
or Clindamycin cream 2% 5 g intravaginally daily for 7 days	Treat if appropriate. Asymptomatic women at high risk of premature delivery should be screened during the earliest part of the second trimester and treated with metronidazole 500 mg orally twice daily x 7 days.		
Candidiasis	, ,		
Clotrimazole cream 1% 5 g intravaginally for 7–14 days or	Diagnosis: Clinical diagnosis suggested by external dysuria, vulvar pruritis, pain, redness, vulvar edema, or thick curdy vaginal discharge. Diagnosis can be made by wet prep or gram stain demonstrating yeast or pseudohyphae or culture for yeast species.		
Miconazole cream 2% 5 g intravaginally for 7 days	Treatment: Uncomplicated vulvovaginal candidiasis (VVC) usually responds to short-course intravaginal therapy. Complicated VVC (e.g., recurrent or severe disease, non-albicans candidiasis, or presence of diabetes or other immunocompromised condition) usually requires more intensive treatment regime, sometimes with oral agents (i.e., fluconazole). Pregnancy: Use only topical agents.		
Trichomoniasis (<i>T. vaginalis</i>)			
Metronidazole 2 g orally x1	Diagnosis: Clinical diagnosis suggested by diffuse, malodorous, yellow-green vaginal		
Note: Metronidazole gel has a <50% cure rate and should	discharge with vulvar irritation. Diagnosis is made via microscopy of vaginal secretions and several new rapid tests. Culture is the most sensitive and specific available method of diagnosis.		
not be used.	Pregnancy: Vaginal trichomoniasis is associated with adverse pregnancy outcomes; however, metronidazole treatment does not appear to reduce perinatal morbidity. Clinicians should counsel patients about potential risks and benefits of treatment. Some specialists would defer treatment until after 37 weeks of gestation, using metronidazole 2 gm orally x1.		
External Genital Warts			
Topical agents	Diagnosis: Made by visual inspection and may be confirmed by biopsy.		
or	Treatment: Response to treatment is often poor. Side effects of treatment are often worse		
Cryotherapy	than the condition itself. Treatment does not eliminate infectivity. Visible warts may resolve spontaneously. Therefore, within the BOP, treatment for genital warts is generally NOT		
or	provided. Large warts which require debulking (e.g., perianal warts which are interfering		
Surgical excision	with hygiene) should be removed using electrocautery, with consultation. Inmate education regarding the facts of this condition is critical. Patients should be periodically evaluated for possible neoplasias.		
	Pregnancy: Imiquimod, podophyllin, and podofilox should not be used in pregnancy. However, because genital warts can proliferate and become friable during pregnancy, many specialists advocate their removal during pregnancy. Consult obstetrician.		

Table 3. Syphilis Screening Guidelines and Diagnostic Tests

BOP Syphilis Screening Guidelines

Syphilis Intake Screening:

- · All females.
- Any males who have had sex with another man; or are HIV-infected; or have a history of syphilis, gonorrhea, or chlamydia.

Syphilis Screening Tests: Syphilis screening is performed utilizing nontreponemal titers (e.g., RPR or VDRL). Note that these screening tests have relatively low sensitivity in patients with early primary syphilis or with tertiary syphilis. Nontreponemal test must be confirmed with a treponemal test (e.g., FTA-ABS). For details, see "Syphilis Diagnostics Tests" below in this table.

Note: All inmates diagnosed with syphilis should be tested for HIV.

Syphilis Diagnostic Tests

Direct Methods: Used with skin lesions or pathologic specimens. Includes darkfield microscopy and the direct fluorescent-antibody (DFA-TP) test.

Serologic Tests: A presumptive diagnosis of syphilis can be made by two different types of serologic, diagnostic tests: quantitative nontreponemal tests and confirmatory treponemal assays.

Note: Serologic tests for syphilis in persons with HIV infection are often more variable, but are still helpful diagnostically and for evaluating treatment response.

Quantitative Nontreponemal Tests

- VDRL: Venereal Disease Research Laboratory
 RPR: Rapid Plasmin Reagin
- ➤ Nontreponemal assays correlate with disease activity. A clinically significant difference between two tests requires at least a fourfold change in titer, e.g., 1:4 to 1:16; or 1:32 to 1:8. The same test should be used for comparisons, e.g., RPR to RPR. The RPR or VDRL titer should become non-reactive with treatment. However, some people will remain "serofast" with a low titer, despite adequate treatment. Serologic titers may decline more slowly for persons with recurrent syphilis.
- ➤ Reactive nontreponemal tests must be confirmed by a treponemal assay, because nontreponemal assays can be false-positive. See below.

Treponemal Assays

- FTA-ABS: Fluorescent treponemal antibody absorbed
 TP-PA: T. pallidum particle agglutination
- ➤ Treponemal assays usually remain positive for life; however, 15–25% of persons treated during primary syphilis may revert to a nonreactive status after 2–3 years.

• Cerebrospinal Fluid (CSF) Tests

- ➤ Neurosyphilis is diagnosed by clinical or laboratory findings.
- ➤ A positive CSF-VDRL (in the absence of significant blood contamination) is considered diagnostic of neurosyphilis; however, certain persons with neurosyphilis will have a negative CSF-VDRL.

- ➤ A negative CSF-FTA-ABS excludes nearly all cases of neurosyphilis.
- ➤ The CSF leukocyte count is usually elevated (>5 WBCs/mm³) in patients with neurosyphilis and is a helpful measure to assess treatment response.

Table 4. Syphilis Treatment and Monitoring

Treatment	Monitoring	
Stage: Primary/Secondary		
Standard: Benzathine penicillin G 2.4 million units IM x1.	HIV-negative: Clinical evaluation & RPR at 6 & 12 months; retest for HIV after 3 months if the first HIV test result is negative.	
Penicillin allergy: Doxycycline 100 mg orally twice daily for 14 days; or tetracycline 500 mg four times daily for 14 days.	 HIV-infected: Clinical evaluation & RPR at 3, 6, 9, 12 & 24 months. Comments: Inmates who have persistent or recurrent signs or symptoms, or who have a sustained fourfold increase in RPR titer, should be re-treated. Repeat HIV testing and perform a CSF analysis. 	
If pregnant: Desensitize and treat with penicillin. Needs closer follow-up.		
HIV: Same treatment as above. Lower threshold for CSF exam.	Failure of RPR to decline fourfold in 6 months after initial treatment suggests possible treatment failure. Repeat HIV	
Retreatment: Administer benzathine penicillin G 2.4 million units IM once weekly for 3 weeks, unless CSF exam indicates that neurosyphilis is present.	serology. If HIV-negative, consider CSF exam and consider retreatment as above.	
Stage: Latent		
Early latent: Benzathine penicillin G 2.4 million	HIV-negative: Clinical evaluation & RPR at 6, 12 & 24 months.	
units IM x1. Late latent: Benzathine penicillin G 2.4 million units IM x3 doses that are administered 1 week apart (for a total of 7.2 million units).	HIV-infected: Clinical evaluation & RPR at 6, 12, 18 & 24 months. If early latent, then use lower threshold for CSF exam. If late latent, then perform CSF exam. Comments:	
Penicillin allergy: Doxycycline 100 mg orally twice daily for 28 days; or tetracycline 500mg four times	 Inmates should be retreated & evaluated for neurosyphilis if: Titers increase fourfold; or 	

If pregnant: Desensitize & treat with penicillin.

HIV: Same treatment as above.

daily for 28 days.

- - > Titers increase fourfold; or
 - ➤ An initially high titer (>1:32) fails to decline fourfold within 12-24 months: or
 - > Signs & symptoms of syphilis develop.
- HIV-infected: Use lower threshold for re-evaluating CSF.

Stage: Tertiary (gumma and cardiovascular syphilis)

Refer to specialist for treatment and follow-up for tertiary syphilis.

Neurosyphilis and Syphilitic Eye Involvement

Standard: Aqueous crystalline penicillin G 3-4 million units IV every 4 hours for 10-14 days; or continuous infusion of 18-24 million units IV per day for 10-14 days.

Penicillin allergy: Skin test to confirm allergy. Ceftriaxone may be an alternative, at 2g daily, either IM or IV, for 10-14 days.

If pregnant: Desensitize & treat with penicillin.

- If pleocytosis was present initially, repeat CSF exam every 6 months until the cell count is normal.
- · Monitor RPR titers periodically.
- If the cell count is not normal in 2-3 years, consider retreatment.
- HIV-infected persons may have persistent CSF abnormalities and warrant close clinical follow-up.

Management of Contacts

Sexual transmission of T. pallidum occurs when lesions are present (usually in 1st year after infection). Syphilis cases whose partners are at-risk, and thus merit evaluation include:

· Primary syphilis: Partners exposed 3 months prior to treatment plus duration of symptoms. Secondary syphilis: Partners exposed 6 months prior to treatment plus duration of symptoms.

• Early latent syphilis: Partners exposed 1 year prior to treatment.

· Latent syphilis: Long-term partners.

Examine contacts clinically and serologically. If contact was exposed within previous 90 days, treat presumptively. If exposure occurred greater than 90 days ago, then treat based upon serologies.

References

Centers for Disease Control and Prevention. Sexually Transmitted Diseases Treatment Guidelines, 2010. *MMWR*. 2010;59(No. RR-12):1–110. Available from: http://www.cdc.gov/STD/treatment/2010/default.htm

Centers for Disease Control and Prevention (homepage on the internet). *Syphilis (Treponema pallidum)*–1996 case definition. Accessed November 6, 2008. Available from: http://www.cdc.gov/osels/ph surveillance/nndss/casedef/syphiliscurrent.htm

Centers for Disease Control and Prevention. Update to CDC's sexually transmitted diseases treatment guidelines, 2006: Fluoroquinolones no longer recommended for treatment of gonococcal infections. *MMWR*. 2007;56:332–336. Available from:

http://www.cdc.gov/mmwr/preview/mmwrhtml/mm5614a3.htm?s_cid=mm5614a3_e

Recommended Algorithms for Diagnostic Assessment and Management of Syndromes:

Highly useful algorithms for diagnosing and managing the following syndromes have been posted on the BOP Clinical Practice Guidelines Web page (http://www.bop.gov/news/medresources.jsp):

- Genital Ulcer Disease (Male/Female) Darkfield Unavailable
- Urethritis Gram Stain Unavailable
- Cervicitis
- Pelvic Inflammatory Disease
- Proctitis
- Vaginal Discharge
- Differential Diagnosis of Vaginitis

The algorithms originally appeared in *Managing STDs in the Correctional Setting: A Guide for Clinicians; 3rd Edition* (available at http://www.ratelleptc.org/Resources), and are posted with permission from the Sylvie Ratelle STD/HIV Prevention Training Center of New England.

Patient Educational Materials

The Centers for Disease Control and Prevention provides STD fact sheets in both English and Spanish (see http://www.cdc.gov/STD/HealthComm/fact_sheets.htm). Currently available CDC fact sheets cover:

- Bacterial Vaginosis
- · Chlamydia
- · Genital HPV Infection: HPV and Men
- Genital Herpes
- Gonorrhea
- Lymphogranuloma venereum (LGV)
- Pelvic Inflammatory Disease
- STD Detection and Treatment in HIV Prevention
- STDs and Pregnancy
- Syphilis: Syphilis and Men Who Have Sex with Men
- Trichomoniasis

Attachment 1. Syphilis Stages and Classification

Syphilis, primary (1°)

Clinical description: A stage of infection with *Treponema pallidum* characterized by one or more chancres (ulcers); chancres might differ considerably in clinical appearance. Primary syphilis is highly contagious. The disease can be transmitted from any contact with one of the ulcers.

Laboratory criteria for diagnosis: Demonstration of *T. pallidum* in clinical specimens by darkfield microscopy, direct fluorescent antibody (DFA-TP), or equivalent methods.

Case classification:

• Confirmed: A clinically compatible case that is laboratory confirmed.

Syphilis, secondary (2°)

Clinical description: A stage of infection caused by *T. pallidum* and characterized by localized or diffuse mucocutaneous lesions, often with generalized lymphadenopathy. The primary chancre may still be present. Approximately 25 percent of syphilis cases proceed to secondary syphilis, which lasts 4–6 weeks. This phase can include hair loss; a sore throat; white patches in the nose, mouth, and vagina; fever; headaches; and a skin rash. There can be lesions on the genitals that look like genital warts, but are caused by spirochetes rather than the wart virus. These wart-like lesions, as well as the skin rash, are highly contagious. Rash can occur on palms of hands. Infection can be transmitted by casual contact.

Laboratory criteria for diagnosis: Demonstration of *T. pallidum* in clinical specimens by darkfield microscopy, DFA-TP, or equivalent methods.

Case classification:

- Probable: A clinically compatible case with a nontreponemal (VDRL or RPR) titer greater than or equal to 4.
- Confirmed: A clinically compatible case that is laboratory confirmed.

Syphilis, latent

Clinical description: A stage of infection caused by *T. pallidum* in which organisms persist in the body of the infected person without causing symptoms or signs. Latent syphilis is subdivided into *early*, *late*, and *unknown* categories (see below) based on the duration of infection.

Case classification:

- Probable: No clinical signs or symptoms of syphilis and the presence of one of the following:
 - ➤ No past diagnosis of syphilis, a reactive nontreponemal test (i.e., VDRL or RPR), and a reactive treponemal test (i.e., FTA-ABS, TP-PA assay, various EIAs, and chemiluminescence immunoassays).
 - ➤ A fourfold change in titer, which is equivalent to a change of two dilutions and is considered necessary to demonstrate a clinical significant difference between two nontreponemal test results obtained using the same serologic test.

Syphilis, early latent

Clinical description: A subcategory of latent syphilis. When the initial infection has occurred within the previous 12 months, latent syphilis is classified as *early latent*.

Case classification:

- **Probable:** Latent syphilis in a person who has evidence of having acquired the infection within the previous 12 months, based on **one or more** of the following criteria:
 - > Documented seroconversion or a fourfold or greater increase in titer of a nontreponemal test during the previous 12 months.
 - ➤ A history of symptoms consistent with primary or secondary syphilis during the previous 12 months.
 - ➤ A history of sexual exposure to a partner who had confirmed or probable primary or secondary syphilis, or probable early latent syphilis (documented independently as having duration of less than 1 year).
 - ➤ Reactive nontreponemal and treponemal tests from a person whose only possible exposure occurred within the preceding 12 months.

(Attachment 1 continued on next)

Attachment 1. Syphilis Stages and Classification (continued from previous page)

Syphilis, late latent

Clinical description: A subcategory of latent syphilis, for persons whose only possible exposure occurred during the previous 12 months (in the absence of reactive nontreponemal and treponemal test indications), an asymptomatic person should be considered to have late latent syphilis or syphilis of unknown duration.

Case classification:

• **Probable:** Latent syphilis in a patient who has no evidence of having acquired the disease within the preceding 12 months, and whose age and titer do not meet the criteria specified for latent syphilis of unknown duration.

Syphilis, latent, of unknown duration

Clinical description: A subcategory of latent syphilis. When the date of initial infection cannot be established as having occurred within the previous year, and the patient's age and titer meet criteria described below, latent syphilis is classified as *latent syphilis of unknown duration*.

Case classification:

• **Probable:** Latent syphilis that does not meet the criteria for early latent syphilis, and the patient is aged 13-35 years and has a nontreponemal titer greater than or equal to 32.

Neurosyphilis

Clinical description: Evidence of central nervous system infection with T. pallidum.

Laboratory criteria for diagnosis: A reactive serologic test for syphilis and reactive VDRL in cerebrospinal fluid (CSF).

Case classification:

- Probable: Syphilis of any stage, a negative VDRL in CSF, and both of the following:
 - ➤ Elevated CSF protein or leukocyte count in the absence of other known causes of these abnormalities.
 - ➤ Clinical symptoms or signs consistent with neurosyphilis without other known causes for these clinical abnormalities.
- Confirmed: Syphilis of any stage that meets the laboratory criteria for neurosyphilis.

Syphilis, late, with clinical manifestations other than neurosyphilis

Includes: Late benign syphilis and cardiovascular syphilis.

Clinical description: Clinical manifestations of late syphilis other than neurosyphilis may include inflammatory lesions of the cardiovascular system, skin, and bone. Rarely, other structures (e.g., the upper and lower respiratory tracts, mouth, eye, abdominal organs, reproductive organs, lymph nodes, and skeletal muscle) may be involved. Late syphilis usually becomes clinically manifest only after a period of 15–30 years of untreated infection. The term **tertiary syphilis** refers to gumma and cardiovascular syphilis (but *not* neurosyphilis). Gumma are soft rubbery tumors often occurring in the mouth and are characteristic of tertiary syphilis.

Laboratory criteria for diagnosis: Demonstration of *T. pallidum* in late lesions by fluorescent antibody or special stains (although organisms are rarely visualized in late lesions).

Note: Analyze CSF for evidence of neurosyphilis when evaluating late syphilis with clinical manifestations.

Case classification:

- Probable: Characteristic abnormalities or lesions of the cardiovascular system, skin, bone, or other structures
 with a reactive treponemal test, in the absence of other known causes of these abnormalities, and without CSF
 abnormalities and clinical symptoms or signs consistent with neurosyphilis.
- Confirmed: A clinically compatible case that is laboratory-confirmed.

References:

CDC (home page on the internet). Syphilis (Treponema pallidum)—1996 case definition. Accessed November 6, 2008. Available at: http://www.cdc.gov/osels/ph_surveillance/nndss/casedef/syphiliscurrent.htm

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