Sexually Transmitted Disease
Treatment Tables

Federal Bureau of Prisons
Clinical Practice Guidelines

June 2011

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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 Table 1):
- 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 Table 2):
- In the case of pregnancy, consultation with an obstetrician or an infectious disease specialist is recommended.

Vaginitis (see Table 2):
- 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- For Candidiasis, treatment with Clotrimazole tablets has been replaced by treatment with one of the formulary creams: clotrimazole vaginal cream 1% or miconazole vaginal cream 2%.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 Table 4).
- 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 Table 4):
- 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 References page.
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### Table 1. Gonorrhea, Chlamydia, Nongonococal Urethritis, and PID

<table>
<thead>
<tr>
<th>Treatment (directly observed)</th>
<th>Comments</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Gonorrhea (GC) (N. gonorrhoeae) of the cervix, urethra, rectum, pharynx</strong></td>
<td></td>
</tr>
</tbody>
</table>
| Ceftiraxone 250 mg IM x1 | **Screening:** No routine screening at intake unless symptoms of gonorrhea are present, or unless syphilis or chlamydia have been diagnosed.  
**Diagnosis:** Nucleic acid amplification tests (NAAT) for symptomatic inmates. Follow-up tests to prove cure is not indicated unless symptoms persist.  
**Contacts:** Sex partners of patients with N. gonorrhoeae infections, whose last sexual contact with the patient was within 60 days before onset of symptoms or diagnosis, should be evaluated and treated for N. gonorrhoeae, C. trachomatis, syphilis, and HIV.  
**Note:** Ceftiraxone is available in 250 mg, 500mg, and 1- and 2-gram vials. **Recommendation:** 250 mg IM  
**Note:** Quinolones are no longer recommended for treatment of gonorrhea. |
| **Alternative treatments:** |  
- Ceftizoxime 400 mg orally x1 or  
**PLUS:** treat for chlamydial infection unless chlamydial infection is ruled out with sensitive test—nucleic acid amplification test (NAAT):  
- Azithromycin 1 g orally x1 or  
- Doxycycline 100 mg orally twice daily for 7 days |
| **Pharyngeal gonorrhea:** Treat with ceftriaxone.  
**HIV co-infection:** Treatment regimens are the same as for inmates without HIV co-infection.  
**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  
- Are age 25 and under, regardless of risk factors, or  
- Are older than age 25 with risk factors (e.g., new or multiple sex partners) and/or  
- Have HIV infection and/or  
- Have history of syphilis, gonorrhea, or chlamydia  
**Diagnosis:** For symptomatic inmates, confirm infection by 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. |
| Doxycycline 100 mg orally twice daily for 7 days | **Alternative treatments:** See CDC guidelines at [www.cdc.gov/stdtreatment/](http://www.cdc.gov/stdtreatment/)  
**Pregnant women:**  
- Azithromycin 1 g orally x1 or  
- Amoxicillin 500 mg orally three times daily for 7 days |
| Nongonococal Urethritis |
| Azithromycin 1 g orally x1 or | **Diagnosis:** All patients with confirmed or suspected urethritis should be tested for gonorrhea and chlamydia.  
**Treatment:** M. genitalium may respond better to azithromycin. **DOT dose.** |
| Doxycycline 100 mg orally twice daily for 7 days | **Contacts:** Refer all sex partners within preceding 60 days for treatment. |
| **Alternative treatments:** See CDC guidelines at [www.cdc.gov/stdtreatment/](http://www.cdc.gov/stdtreatment/) |
| Pelvic Inflammatory Disease (PID) – Outpatient Management |
| **Recommended regimen:**  
- Ceftiraxone 250 mg IM plus  
- Doxycycline 100 mg orally twice daily for 14 days | **Diagnosis:** Minimum criteria based on pelvic exam finding:  
- Cervical motion tenderness or uterine tenderness or adnexal tenderness  
**Hospitalization:** If surgical emergencies cannot be excluded, or if pregnancy, lack of clinical response to oral therapy, severe illness & unable to follow/delate an outpatient oral regimen, or has tubo-ovarian abscesses. |
| **Alternative regimen:**  
- Cefotaxime 2 g IM x 1 or other third-generation cephalosporin plus  
- Probenecid 1 g orally x1 plus  
- Doxycycline 100 mg orally twice daily for 14 days |  
**Note:** Both regimens can be given with or without metronidazole 500 mg orally twice daily for 14 days |
Table 2. Herpes Simplex Virus, Vaginitis, and Genital Warts

<table>
<thead>
<tr>
<th><strong>Herpes Simplex Virus (HSV)</strong></th>
<th><strong>Comments</strong></th>
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</thead>
<tbody>
<tr>
<td><strong>First episode:</strong>&lt;br&gt;Acyclovir 400 mg orally three times daily for 7–10 days</td>
<td>General: Genital herpes is a recurrent, lifelong infection. Sexual transmission of HSV occurs in asymptomatic persons. 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: 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 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. Pregnancy: Consult with obstetrician or infectious disease specialist.</td>
</tr>
<tr>
<td><strong>Recurrent episodes:</strong>&lt;br&gt;Acyclovir 400 mg orally three times daily for 5 days</td>
<td></td>
</tr>
<tr>
<td><strong>Suppressive therapy:</strong>&lt;br&gt;Acyclovir 400 mg orally twice daily for 1 year</td>
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</table>

**Vaginitis**

| **Bacterial Vaginosis (BV)** | **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 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. Treat if appropriate. Asymptomatic women at high risk of premature delivery should be screened during the second trimester and treated with metronidazole 500 mg orally twice daily x 7 days. |
| **Metronidazole 500 mg orally twice daily for 7 days** or **Metronidazole gel 0.75% 5 g intravaginally for 5 days** or **Clindamycin cream 2% 5 g intravaginally for 2 days** | |

**Candidiasis**

| **Clotrimazole cream 1% 5 g intravaginally for 7–14 days** or **Miconazole cream 2% 5 g intravaginally for 7 days** | **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. 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 (T. vaginalis)**

| **Metronidazole 2 g orally x1** | **Diagnosis:** Clinical diagnosis suggested by diffuse, malodorous, yellow-green vaginal 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. 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** or **Cryotherapy** or **Surgical excision** | **Diagnosis:** Made by visual inspection and may be confirmed by biopsy. Treatment: Response to treatment is often poor. Side effects of treatment are often worse than the condition itself. Treatment does not eliminate infection. Visible warts may resolve spontaneously. Therefore, within the BOP, treatment for genital warts is generally NOT provided. Large warts which require debulking (e.g., penile warts which are interfering 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 neoplasia. 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. |

Note: Metronidazole gel has a<50% cure rate and should not be used.
### Table 3. Syphilis Screening Guidelines and Diagnostic Tests

<table>
<thead>
<tr>
<th>BOP Syphilis Screening Guidelines</th>
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<tbody>
<tr>
<td><strong>Syphilis Intake Screening:</strong></td>
</tr>
<tr>
<td>All females.</td>
</tr>
<tr>
<td>Any males who have had sex with another man; or are HIV-infected; or have a history of syphilis, gonorrhea, or chlamydia.</td>
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</table>

**Syphilis Screening Tests:** Syphilis screening is performed utilizing nontreponemal tests (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 Diagnostic Tests" below in this table.

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

<table>
<thead>
<tr>
<th>Syphilis Diagnostic Tests</th>
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<tbody>
<tr>
<td><strong>Direct Methods:</strong> Used with skin lesions or pathologic specimens. Includes darkfield microscopy and the direct fluorescent-antibody (DFA-TP) test.</td>
</tr>
</tbody>
</table>

**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

<table>
<thead>
<tr>
<th>Stage: Primary/Secondary</th>
<th>Treatment</th>
<th>Monitoring</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Standard</strong>: Benzathine penicillin G 2.4 million units IM x 1.</td>
<td><strong>HIV-negative</strong>: Clinical evaluation &amp; RPR at 6 &amp; 12 months, retest for HIV after 3 months if the first HIV test result is negative.</td>
<td></td>
</tr>
<tr>
<td>Penicillin allergy: Doxycycline 100 mg orally twice daily for 14 days, or tetracycline 500 mg four times daily for 14 days.</td>
<td><strong>HIV-infected</strong>: Clinical evaluation &amp; RPR at 3, 6, 9, 12 &amp; 24 months.</td>
<td></td>
</tr>
<tr>
<td>If pregnant: Desensitize and treat with penicillin. Needs closer follow-up.</td>
<td><strong>Comments</strong>:</td>
<td></td>
</tr>
<tr>
<td>HIV: Same treatment as above. Lower threshold for CSF exam.</td>
<td>• 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.</td>
<td></td>
</tr>
<tr>
<td>Retreatment: Administer benzathine penicillin G 2.4 million units IM once weekly for 3 weeks, unless CSF exam indicates that neurosyphilis is present.</td>
<td>• Failure of RPR to decline fourfold in 6 months after initial treatment suggests possible treatment failure. Repeat HIV serology. If HIV-negative, consider CSF exam and consider retreatment as above.</td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>Stage: Latent</th>
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</thead>
<tbody>
<tr>
<td><strong>Early latent</strong>: Benzathine penicillin G 2.4 million units IM x 1.</td>
<td><strong>HIV-negative</strong>: Clinical evaluation &amp; RPR at 6, 12 &amp; 24 months.</td>
</tr>
<tr>
<td><strong>Late latent</strong>: Benzathine penicillin G 2.4 million units IM x 3 doses that are administered 1 week apart (for a total of 7.2 million units).</td>
<td><strong>HIV-infected</strong>: Clinical evaluation &amp; RPR at 6, 12, 18 &amp; 24 months. If early latent, then use lower threshold for CSF exam. If late latent, then perform CSF exam.</td>
</tr>
<tr>
<td>Penicillin allergy: Doxycycline 100 mg orally twice daily for 23 days, or tetracycline 500 mg four times daily for 28 days.</td>
<td><strong>Comments</strong>:</td>
</tr>
<tr>
<td>If pregnant: Desensitize &amp; treat with penicillin.</td>
<td>• Inmates should be retreated &amp; evaluated for neurosyphilis if:</td>
</tr>
<tr>
<td>HIV: Same treatment as above.</td>
<td>&gt; Titers increase fourfold, or</td>
</tr>
<tr>
<td></td>
<td>&gt; An initially high titer (≥1:32) fails to decline fourfold within 12–24 months, or</td>
</tr>
<tr>
<td></td>
<td>&gt; Signs &amp; symptoms of syphilis develop.</td>
</tr>
<tr>
<td></td>
<td>• HIV-infected: Use lower threshold for re-evaluating CSF.</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Stage: Tertiary (gumma and cardiovascular syphilis)</th>
<th></th>
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</thead>
<tbody>
<tr>
<td>Refer to specialist for treatment and follow-up for tertiary syphilis.</td>
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</tbody>
</table>

### 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. | If pleocytosis was present initially, repeat CSF exam every 6 months until the cell count is normal. |
| Penicillin allergy: Skin test to confirm allergy. Ceftriaxone may be an alternative, at 2g daily, either IM or IV, for 10–14 days. | Monitor RPR titers periodically. |
| If pregnant: Desensitize & treat with penicillin. | 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


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/mcresources.jsp):

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


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/HealthComms/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 (1st)

**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 (2nd)

**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 nonreproducible (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 nonrecombinant 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 syphils 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 nonrecombinant 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: