Sexually transmitted diseases in children: chlamydial oculo-genital infection

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Introduction

Three species of chlamydia responsible for a wide spectrum of diseases (table 1), are currently recognised: *C trachomatis*, *C psittaci* and *C pneumoniae*; sexual transmission is thought not to occur with the last two. There are at least 15 serovars of *C trachomatis*. In general, serovars A, B, Ba and C cause hyperendemic trachoma, serovars D to K, oculogenital infections and related diseases (table 2) and serovars L1–3 cause lymphogranuloma venereum (LGV); the last two groups of serovars are sexually transmissible. Infection by the D–K serovars affects children of all ages but is uncommon in the prepubertal period. It ranges from perinatal and neonatal infection acquired from the mother, to prepubertal infection as a consequence of long term carriage or voluntary or involuntary sexual intercourse, to sexually transmitted adolescent infection. Infection by the LGV serovars is seen mainly in adolescents with occasional cases reported in prepubertal children.

Historical review

The aetiology of ophthalmia neonatorum was poorly understood until the discovery of the gonococcus by Neisser¹ in 1879 which enabled Kroner² in 1884 to differentiate the nongonococcal “amicrobial” form. Halberstaedter and von Prowazek³ first demonstrated the intracytoplasmic inclusions of chlamydia in conjunctival scrapings of trachomatous lesions of man and orangutans in 1907. By 1909, similar inclusions were found in conjunctival scrapings from babies with ophthalmia^{4–6} and in the genital tract of the parents of such babies.^{4,7} In 1957, Tang *et al*⁸ isolated the chlamydial agent in egg yolk sacs. The introduction of a cell culture technique by Gordon and Kuan⁹ in 1965 and the microimmunofluorescence method for the detection of antibody and immunotyping by Wang and Grayston¹⁰ in 1970 enabled widespread study of the epidemiology and clinical

spectrum of chlamydial infections. *C trachomatis* was first cultured from the cervix of a mother whose baby had nongonococcal ophthalmia in 1959 by Jones, *et al*.¹¹ Pioneering work by Dunlop and associates^{12,13} in the 1960s confirmed that *C trachomatis* was an important and common oculogenital pathogen. Extraocular manifestation in infants was probably first described under the entity of pertussoid eosinophilic pneumonia by Botsztein¹⁴ in 1941. However, it was not until 1975 that Schachter *et al*¹⁵ suggested that *C trachomatis* could cause afebrile pneumonitis. This syndrome was further characterised by Beem and Saxon¹⁶ in 1977. *Lymphogranuloma venereum* (LGV) was characterised as a separate venereal disease in 1913 by Durand, *et al*¹⁷ who suggested the term “lymphogranuloma inguinale”. The introduction of the Frei skin test¹⁸ in 1925 for the diagnosis of LGV enabled the recognition of its many clinical presentations. The first case report in a child was probably that described by Weiss and Cain in 1925¹⁹ of a 14 year old girl presenting with suppurative inguinal bubo. Additional cases in prepubertal children were reported from 1932 onwards.^{20–22}

Epidemiology

C trachomatis is one of the commonest bacterial sexually transmitted diseases worldwide. This is more so in industrialised countries such as the USA and England but may reflect the accessibility of diagnostic facilities.^{23,24} Studies on the prevalence of chlamydia in different settings vary considerably and depend on many factors such as the study population and socioeconomic circumstances.

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Table 1 Clinical spectrum of chlamydia

Species	Serovars	Diseases
<i>C psittaci</i>	A no. of serovars	Psittacosis, ornithosis Abortion Endocarditis Pneumonia
<i>C pneumoniae</i>	L1, L2, L3	Lymphogranuloma venereum
<i>C trachomatis</i>	A, B, Ba, C	Hyperendemic trachoma
<i>C trachomatis</i>	D,E,F,G,H,I,J,K	Oculo-genital and other infections (table 2)

Table 2 Diseases in children caused by *C trachomatis*

Neonate: Conjunctivitis Rhinitis Pharyngitis Pneumonia Otitis media Proctitis Premature birth ?Stillbirth	Adolescent female: Cervicitis Bartholinitis Salpingitis Endometritis Ectopic pregnancy and infertility ?Cervical intraepithelial neoplasia
Children: Anogenital infection Croup Otitis media Myocarditis	Adolescent male and female: Conjunctivitis Punctate keratitis Endemic trachoma Cystitis Proctitis Perihepatitis Reiter's disease Pneumonia Peritonitis ?Appendicitis
Adolescent male: Nongonococcal urethritis Epididymitis	

Infection in pregnancy and the neonate

The prevalence of chlamydial infection in pregnant women is extremely varied. Reports range from 2–37%^{25–33} in the USA, 0–33% in Europe,^{34–37} and 1.3–29% in Africa,^{38–43} the higher figures tending to be amongst inner city adolescents, socially deprived ethnic groups or patients attending STD clinics. In Sweden⁴⁴ the prevalence of chlamydial infection in pregnant women has declined from 19% in 1984/5 to 2.4% in 1988. *C trachomatis* was isolated from 14% of women seeking abortion in Singapore⁴⁵ and 13% and 45% from pregnant women in Thailand⁴⁶ and Fiji⁴⁷ respectively.

The annual incidence of chlamydial ophthalmia neonatorum in Kenya is 81/1000 live births⁴³ compared with less than 4/1000 live births in the USA.²⁶ The prevalence of chlamydia in babies with ophthalmia neonatorum is 49% in England⁴⁸ and 17–35% in Africa.^{39 43 49}

Infection in prepubertal children

The prevalence of chlamydia in children is mainly based on seroepidemiological studies. In the USA,^{50–52} prevalences range from 10–15% starting between ages 2 to 9 and increasing with age.^{53–55} This is in contrast to a study in England of children aged 1–15 years which showed a low prevalence below the age of seven with no antibodies above that age.⁵⁶ Correlating serological responses to specific chlamydial infection in seroepidemiological studies is, however, difficult because of the cross reactivity between and within many of the chlamydial species.

Infection in adolescents

This sexually active group is at greatest risk for chlamydial infection and more so in socially deprived groups and in urban areas. In the USA the prevalence in adolescent males^{57 58} varies from 9–35% and in non-pregnant females^{58–63} from 8–25%. Sixteen percent of adolescent girls in residential care in England⁶⁴ and 19% of sexually active teenage girls in Sweden,⁶⁵ were found to have chlamydial infection.

Lymphogranuloma venereum

There are very few reported cases of lymphogranuloma venereum in children, where most have been diagnosed on clinical grounds supported by a positive Frei test. However, the antigen used in the Frei test is common to all species of chlamydia, and false positive results may occur. This test is no longer in use. Before 1939, less than 15 possible cases in children aged 3 to 14 years were reported in the literature.⁶⁶ In 1953, Greenblatt *et al*⁶⁷ presented a series of 197 consecutive cases from the USA. These included two children aged less than 10 years and 31 patients aged between 11 and 20 years. In 1962, Annamunthodo and Stewart,⁶⁸ reported eight cases in children aged 16 years or under from Jamaica. Buboec were present in two boys and five girls whilst one girl had proctitis. The prevalence of LGV in pregnant women is

unknown apart from isolated case reports.^{69–73}

LGV is endemic in tropical and subtropical areas of southeast Asia, India, east and west Africa, the Caribbean, and certain parts of the United States and South America. The few sporadic cases reported in Europe are mainly imported. There have been no studies of antenatal and paediatric cases in these endemic areas over the past three decades despite the introduction of more specific tests for LGV.

Routes of transmission

Transmission may be transplacental, perinatal, accidental or sexual including sexual abuse.

Transplacental

This is controversial because it is difficult to exclude ascending infection from the cervix and also the evidence is circumstantial, based on the probable effect of chlamydia on the foetus. The transplacental route of transmission was first suggested by the isolation of *C trachomatis* from spontaneously aborted foetuses⁷⁴ and serological studies of preterm babies with low birth weight but without ophthalmia born to infected mothers.^{75 76} This was supported by case reports of intrauterine chlamydial pneumonia in a stillbirth with intact amniotic membranes and in a premature infant who died when aged 4 days.^{77 78} Blood-borne infection from mother to foetus is a possibility as chlamydia is capable of systemic spread as evidenced by a report of *C trachomatis* endocarditis in a pregnant woman⁷⁹ and some cases of Reiter's disease in which there was evidence of dissemination from the genital tract to the joints.^{80–82}

Perinatal transmission

This is the major route of transmission in neonates. The infection is acquired at birth from the cervix of infected mothers during vaginal delivery. However, infection has been reported in babies delivered by Caesarian section,⁸³ usually in association with early rupture of the amniotic membrane where presumably the infection had ascended from the cervix. Babies born to infected women have a 18–50% chance of developing ophthalmia,^{25–27 30 84–86} 8–20% may develop pneumonia^{25–27 30 84 86} whilst 60–70% will have some evidence of infection.^{26 27 84 85}

Different sites such as conjunctiva, nasopharynx and vulvo-vagina can be seeded independently. Once infected, involvement of other sites is usually via contiguous spread or in the case of the rectum, by ingestion of infected ocular or nasopharyngeal material.

Accidental transmission

Accidental inoculation of infected material from the patients themselves or from infected partners or family members to the eyes or genital tract can lead to ocular or genital infection. A case of chlamydial conjunctivitis in an 18 month old sister of a baby with

chlamydial ophthalmia neonatorum has been described.⁸³ Non-sexual transmission in girls with chlamydial vulvo-vaginitis is thought to occur^{88 89} but sexual abuse cannot be excluded.

Child sexual abuse

This should be borne in mind in children with chlamydial infection because of the serious implications. Of sexually abused children 4%-17% in American studies have been found to have chlamydial infection. This may be concomitant with gonorrhoea.⁹⁰⁻⁹³

Voluntary sexual intercourse

Sexual transmission is directly or indirectly responsible for the spread of infection. Genital infection in older children generally indicates that they are either sexually active or abused. When considering the possibility of child sexual abuse, persistent infection has to be excluded.

Persistent infection

Chlamydial infection may persist for a long time after perinatal or sexual transmission. Perinatal infection may lead to persistent infection in the vagina, rectum or both for up to three years, and in the pharynx for up to 29 months.⁹⁴ A 17 month old toddler and a 6 year old girl with persistent neonatal chlamydial ocular infection and a 3 year old girl with recto-vaginal infection have been reported; the older toddler also had chlamydial vulvo-vaginitis.^{13 95}

Lymphogranuloma venereum

Lymphogranuloma venereum in children is acquired through sexual intercourse or non-venereal accidental transmission.^{68 96} In the latter, it is difficult to exclude the possibility of sexual abuse. Five cases, all girls aged 4-9 years with rectal involvement, were reported where transmission was probably from sharing the same beds or contaminated fomites from their infected mothers.⁶⁶ A case was reported in an 8 year old girl presenting with proctitis and malaena where an enema tip used by her infected mother was also used for the child.⁹⁷ Transmission through homosexual anal intercourse has been reported in two boys aged 11⁹⁸ and 17.⁶⁸ The older boy had a history of homosexual assault when aged 10 and symptoms of proctitis from age 14. A girl of eleven who acquired both syphilis and LGV following sexual assault has also been described.⁶⁸

There is little evidence of transplacental or perinatal transmission. However, symptomatic and asymptomatic cervical infection have been reported in women. Whether this can lead to perinatal transmission is unknown. The report of the possibility of congenital LGV in a 2 week old healthy baby with a positive Frei test born to an infected mother⁶⁹ is dubious as the Frei test is not specific for LGV. Other isolated reports do not support transplacental transmission.^{70 71}

Clinical presentation

Infection in the neonate

Perinatal infection may lead to prematurity, low birth weight^{29 75 76 99} and possibly stillbirth and neonatal deaths.^{84 99} Although some studies have not shown a relationship with low birth weight,^{28 84} it may be primary chlamydial infection during pregnancy, as evidenced by maternal chlamydial IgM or IgG seroconversion, that is important.^{100 101} The IgM may also indicate an active or invasive process. The presence of IgM in chlamydia-culture-positive women has been related to prematurity.¹⁰² Co-infection with trichomonas was associated with prematurity and low birth weight,³³ whereas with *M hominis*, it resulted in increased incidence of spontaneous abortion, prematurity, still birth and neonatal deaths.¹⁰³ Martin *et al*⁹⁹ reported a 10-fold increase in stillbirths and neonatal deaths based on 18 pregnant women infected before 20 weeks' gestation. This was not confirmed by a larger study which also controlled for age, race, socioeconomic status and other coexisting infections.¹⁰²

The commonest neonatal presentation is purulent conjunctivitis, with an onset at 5-13 days. Neonates with chlamydial ophthalmia may be premature with low birth weight.^{83 100 101} Gonorrhoea may coexist.^{48 49} Chemosis and conjunctival pseudomembranes may be present. If untreated, this may rarely lead to conjunctival scarring and corneal involvement.^{88 106} Other sites such as the nasopharynx, lung, middle ear, genital tract and rectum may also be infected¹⁰⁷ and occasionally lead to symptomatic rhinitis neonatorum,¹⁰⁸ afebrile pneumonia and otitis media.^{27 109 110}

Afebrile pneumonia usually presents in the first three months of life with staccato cough, tachypnoea, inspiratory crepitations and hyperinflation.^{16 17} It may be subclinical without the characteristic symptoms¹¹¹ and there may not be a history of conjunctivitis. Clinical signs are usually minimal, but a chest radiograph may show diffuse interstitial involvement. There may be an associated eosinophilia or increased serum immunoglobulins. The pneumonia may be severe in babies with low birth weight¹¹² and Ig G2/Ig G4 antibody deficiency¹¹³ and may lead to chronic lung disease.^{75 114} Fifty nine percent of infants less than 6 months of age with chlamydial pneumonitis developed middle ear disorders.¹¹⁵ Although afebrile pneumonia is more commonly reported in the United States, it is rarely described in the United Kingdom. This may reflect the size of reservoir of chlamydial infection resulting from inadequate chlamydial diagnostic and treatment facilities, the use of ocular prophylaxis against chlamydia after birth which allows extraocular infection to occur and possibly, the high index of suspicion in those hospitals in the United States where the studies were performed.

Infection in prepubertal children

Prepubertal children can present with urethri-

tis or vulvovaginitis.^{13 50 89 116 117} In one study¹¹⁷ nine children age 4–11 were found to have dual infection with *C trachomatis* and *N gonorrhoeae*. Chlamydia was isolated from either the urethra or rectum in boys and from the vagina or rectum or both in girls. One boy had symptoms of urethritis and two girls had vulvo-vaginal discharge, the rest were asymptomatic. In two cases, asymptomatic chlamydial infection was also seen in four other family members age 4–11 years. A 6 year old girl with chlamydial vulvovaginitis whose mother had chlamydial cervical infection and father had non-specific urethritis has been reported.¹³ Chlamydial recto-vaginal infection in children tends to be asymptomatic, being discovered by routine tests in suspected cases. Sexual abuse has to be excluded.

Whether chlamydia causes otitis media in children and in infants more than 7 months' old is controversial. While most studies^{118–120} show that it plays no role, *C trachomatis* has been isolated from the middle ear aspirates of children 1–8 years old with otitis media.¹²¹ Chlamydia-associated myocarditis in children aged one to six years, which was fatal in some cases has been reported.¹²² *C trachomatis* and *Staphylococcus aureus* were isolated from the subglottic exudate in a three and a half year old boy with tracheitis presenting with a barking cough, inspiratory stridor and respiratory distress after a two days afebrile upper respiratory tract infection.¹²³ Reiter's disease⁸⁰ has been reported in two Canadian Indian boys, aged 2 and 14 years, who also had positive chlamydial isolation from the synovial fluid and urethra respectively. The first boy had arthritis, conjunctivitis and pyuria which was preceded by diarrhoea. The older boy who was sexually active had dysuria, arthritis, enthesopathy, conjunctivitis, balanitis, sacroiliitis and pleurisy.

Infection in adolescents

Presentation of sexually acquired chlamydial infection in adolescents is similar to adults. Asymptomatic infection is common. Adolescent males present with nongonococcal urethritis (NGU) complaining of urethral discharge and painful micturition. *C trachomatis* is responsible for up to 50% of cases of NGU. Epididymitis and Reiter's disease may complicate the infection. Proctitis may result from anal intercourse in males.

The cervix is the commonest site of infection in adolescent girls. There may be mucopurulent cervicitis with follicles, or the cervix may appear normal. Bartholinitis, urethritis, cystitis and follicular proctitis may be seen. The most important complication is salpingitis which may lead to chronic pelvic pain, ectopic pregnancy and sterility. Perihepatitis, leading to right hypochondrial pain, which may or may not be associated with salpingitis, has been reported in adolescent girls age 14–16 years.¹²⁴

Chlamydial pneumonia has been described in two immunosuppressed adolescents aged 15 and 17 years.¹²⁵ One had a renal allograft, the other had chemotherapy for acute

lymphocytic leukaemia. Both had *C trachomatis* isolated from the lung tissue, eye and nasopharynx.

Lymphogranuloma venereum

Inguinal lymphadenopathy and proctitis are the commonest presentation in prepubertal children.⁶⁸ It is rare to find the primary lesion. Occasionally the children may present because an adult member is also affected. LGV may be associated with arthropathy; four cases of arthritis of the knees, one of whom also had swollen ankles, have been reported in a boy aged 11 years⁹⁸ and three girls aged 7, 8 and 12 years.⁶⁶ As in adults cases, erythema nodosum may occur in children.⁶⁶

Sexually active adolescents may present with a primary lesion manifesting as a small transient genital ulcer within 5 weeks of exposure but usually between 2 and 5 days. This is followed by inguinal lymphadenopathy, usually unilateral, which may be multiloculated. Enlargement of the lymph nodes above and below the inguinal ligament may lead to the "groove" sign. The bubo may resolve or rupture forming sinuses. Primary infection of the posterior vaginal wall may involve the rectovaginal septum. Perirectal involvement with proctocolitis may lead to fibrosis, scarring and stricture formation. Genital elephantiasis (esthiomene) may develop with chronic infection.

Diagnostic methods

The choice of diagnostic method depends upon the expertise and resources available, the prevalence of infection in the population to be tested, the workload, the sites of infection and the purpose of the test.

Microscopy

Giemsa-stained smears for chlamydial inclusions are only useful for conjunctival specimens because of its poor sensitivity in other sites.

Culture

Cell culture technique is specific but its sensitivity varies.^{126–129} Confirmation of infection should always be by direct visualisation of stained inclusions and not by the detection of antigen using enzyme immuno assay (EIA) as this can lead to false positives.¹³⁰ The latter should not be considered a "cell culture" technique. Cell culture, although labour intensive, is the method of choice for diagnosing infection in low prevalence populations, for sites other than the conjunctiva, cervix and urethra, for medico-legal purposes and in suspected sexual abuse cases because of its high specificity and positive predictive value.

Antigen detection tests

Antigen detection tests for clinical specimens using the direct immunofluorescent (DIF) method, (for example Microtrak (Syva, UK)), and EIA (such as Chlamydiazyme

(Abbotts), IDEIA (Novo Nordisk Diagnostics, UK)), have provided rapid methods with good sensitivities and specificities.¹³¹

As false positive results may occur especially from anatomical sites where other bacteria may be present, they should not be used in unapproved sites and in medicolegal and sexual abuse cases.¹³² Antigen detection tests should be used with caution in low prevalence populations because of the poor positive predictive value. They may be used to exclude infection because of the high negative predictive value, but a positive result should be confirmed, if possible, by culture. In medico-legal cases, if culture is not available, DIF is the preferred alternative as it can be stored and reassessed.

Methods of specimen collection

For culture, epithelial cells containing viable organisms are required. Success is dependent upon an adequate sampling technique and proper specimen transportation. In male children, the procedure for obtaining urethral material for culture or antigen detection is invasive and painful as the swab has to be inserted into the urethra. The use of a first catch urine specimen to detect chlamydial antigen using DIF or EIA has recently been shown to be sensitive and specific in a high risk adult population¹³³⁻¹³⁷ and could be applied to children. If confirmed, this will be a non-invasive way of diagnosing urethral infection.

In lymphogranuloma venereum, bubo aspirate or swab from the ulcer base or rectal lesion should be taken for diagnostic tests, preferably culture. Aspiration of bubo should be performed using a large bore needle through adjacent intact skin to prevent fistula formation. Biopsy of granulation tissue can also be stained for chlamydial antigen.

Serology

The micro-immunofluorescence method to detect chlamydial antibodies has limited application. It is of use in complicated infections especially when the patient has had antichlamydial suppressive therapy. Serological tests should be interpreted with caution because of cross reaction between the serovars as well as the different chlamydial species. The presence of IgM, or more than a four fold increase in IgG, or high IgG titres (> 1:256) may indicate that the disease is related to chlamydia.

Surrogate markers for chlamydial infection

If chlamydial diagnostic tests are not available, surrogate markers using the presence of pus cells in the absence of gram-negative intra-cellular diplococci and other bacteria in conjunctival and urethral smears can be used for the presumptive diagnosis of chlamydial ophthalmia neonatorum⁴⁸ and non-gonococcal urethritis including those of chlamydial origin.¹³⁸

A criterion of five to ten or more pus cells per high power field ($\times 1000$) in urethral material¹³⁸⁻¹⁴⁰ is diagnostic of non-gonococcal

urethritis. An early morning smear after the patient has held urine will detect additional cases of asymptomatic urethritis.¹⁴¹ First catch urine¹⁴² and endocervical pus counts¹⁴³ may correlate with chlamydial urethritis and mucopurulent cervicitis respectively but are not supported by other studies.^{144 145}

Other techniques

DNA probes and the polymerase chain reaction (PCR) are still research tools. The DNA probe, although specific, is not sensitive. PCR is extremely sensitive but prone to false positive results if laboratory contamination is not meticulously avoided.

Prevention, treatment and follow up regimes

Antenatal screening

Screening and treatment of pregnant women for chlamydial infection where prevalence is high and in those with casual sexual partners would be the best method of preventing perinatal transmission to babies. It is estimated that it is cost effective to screen women when the prevalence of chlamydial infection is above 7%.¹⁴⁶

Neonatal prophylaxis

The use of ocular prophylaxis by instillation of silver nitrate, erythromycin or tetracycline drops or ointment after birth to prevent neonatal ophthalmia is controversial. The presence of gonococcal or chlamydial ophthalmia neonatorum is a marker that other sites such as the nasopharynx and rectum, as well as the mother and her sexual partner(s), are infected. Furthermore 12-30% of untreated mothers will develop pelvic inflammatory disease.^{49 83 147} While it has been shown that the use of 1% silver nitrate (Crede's prophylaxis) drops or tetracycline ointment led to a three fold decrease in the incidence of ophthalmia in Kenya, both treatments did not prevent chlamydial infection of other sites; the onset of pneumonitis between 1-2 months was described in some of the babies.^{86 148} Thus ocular prophylaxis incompletely prevents ocular infections, has no effect on other infected sites and the opportunity for contact tracing is lost. In countries where the prevalence of neonatal gonococcal ophthalmia is high with poor health care facilities and limited resources, ocular prophylaxis may be justified to prevent blindness in the babies. This is at the expense of the health of the parents and control of infection. Ocular prophylaxis should strongly be discouraged in developed countries with good health care facilities. Most states in the USA are still required by law to provide ocular prophylaxis, whereas this practice has long stopped in many European countries including the United Kingdom and Denmark. If prophylaxis is to be used, the aim is to prevent blindness from gonococcal ophthalmia. It might, arguably, be an added advantage if it will also cover chlamydial ophthalmia, but current prophylaxis for gonococcal ophthalmia have

not been effective for chlamydia.^{86 149} Although the cornea may be involved in chlamydial ophthalmia, the risk of blindness is negligible. Furthermore, antibiotics effective against chlamydia are generally poor treatment for gonorrhoea because of the development of drug resistance. If ocular prophylaxis is deemed necessary, 1% silver nitrate eye drops, 1% tetracycline or 0.5% erythromycin eye ointment can be used; silver nitrate is cheaper and as effective but may cause chemical conjunctivitis. It should be instilled immediately into the eyes after birth.

Treatment in neonates

In infants with chlamydial ophthalmia, treatment is with erythromycin stearate or ethylsuccinate suspension 50 mg/kg/day in four divided doses before feeds for two weeks. If pneumonitis is present, treatment should continue for three weeks. Once systemic treatment is started, local eye treatment with ocular tetracycline is unnecessary but may speed up healing.

Treatment in children

Tetracycline should not be given to children under 12 years of age. Erythromycin is the antibiotic of choice. The dosage will depend on the weight or age of patients.

Treatment in adolescents

The treatment of choice for uncomplicated infection is tetracycline 500 mg 6 hourly or doxycycline 200mg daily either as a single dose or 100mg 12 hourly for 7 days.²³ For pregnant women and patients who are allergic to tetracycline, erythromycin stearate, base or ethylsuccinate, 500mg 6 hourly, is given for 7 days. For complicated infections such as pelvic inflammatory disease or epididymitis, treatment is for 3 weeks. Azithromycin, an azalide macrolide has been shown to be effective in a single dose of 1 g¹⁵⁰ but requires more assessment. It is, at present, not recommended for pregnant women and studies on its efficacy and safety in children are under way. A single dose regime will be the treatment of choice in the future as it will lead to better compliance.

Contact tracing, partner notification and other considerations

Chlamydial infection is a marker for other sexually transmitted diseases, hence screening for other common infections such as gonorrhoea is important.

It is essential to look for the source of infection. Checking for chlamydia and other concomitant sexually transmitted diseases in other family members in "non-sexual" cases, perpetrators in sexual abuse cases and sexual partners in cases acquired from voluntary sexual intercourse, is an integral part of management. In cases of neonatal ophthalmia, the parents and their other sexual partners should be screened. Advice on the use of condoms and nonoxynol-9 spermicide may further reduce transmission both of chlamydia and other sexually transmitted diseases.

In cases of suspected child sexual abuse, other professionals, such as the medical social worker, child psychiatrist and paediatrician may need to be involved.

Follow-up

Follow-up of patients is necessary to check compliance and to confirm that sexual partners have been screened. The necessity for a test of cure is controversial as treatment is highly effective. However, it will test for reinfection as well as reassure patients or parents of infected babies that the infection has been successfully treated. If a test of cure or for reinfection is to be performed, it should preferably take place at least two weeks after treatment, particularly if antigen detection tests are used as they can detect dead organisms. Their use as a follow-up test should be used with caution: a negative test is useful for excluding infection but a positive test should be further evaluated.

Future developments

Resources should be provided to implement screening programmes in high risk groups, now that antigen detection methods are available. The further development of cheap, rapid, sensitive, specific, non-invasive user-friendly techniques for diagnosing chlamydial infection is desirable for screening both high and low risk populations. In areas with a high prevalence of infection screening of pregnant women and adolescents, those with casual partners and babies with ophthalmia neonatorum or afebrile pneumonia should be implemented. In countries where LGV is common, prospective antenatal studies to ascertain the prevalence, transmission risk and disease syndromes due to the LGV serovars, if any, in the neonate, would further define its role in children.

New antimicrobials effective against *C trachomatis* are needed which give prolonged high serum levels from a single dose. An alternative to erythromycin with less side effects is desirable for pregnant women. A simple, safe and cheap antibiotic may lead to the possibility of an eradication programme by mass treatment in high prevalence populations. Molecular research on chlamydial immune response with a view to developing a vaccine would be ideal but public acceptance of such a vaccine may be difficult.

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