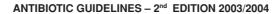
## ANTIBIOTIC GUIDELINES 2<sup>nd</sup> Edition 2003/2004

## MINISTRY OF HEALTH GOVERNMENT OF FIJI

Prepared by the

National Drugs and Therapeutics
Subcommittee

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# Prepared by the Antibiotic Guidelines Sub-Committee of the National Drug and Therapeutics Committee

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We thank the members of the Central/ Eastern Drug and Therapeutics Committee and others for their contribution in the development of the second edition of Antibiotic Guidelines.

The sub-committee would especially like to thank the following individuals: Dr Alan Garvez, Dr Bera Kaitani, Dr Naimun Chand, Dr Jai Narayan, Dr Esala Nainoca and Dr Tuicakau who made specific contributions.

We gratefully acknowledge the financial support of the WHO and the assistance provided by the Chief Pharmacist, Mr. Peter Zinck, in the successful completion of these Antibiotic Guidelines.

The inspiration for the development of the first edition of Antibiotic Guidelines for Fiji came from the Australian Antibiotic Guidelines, published and distributed by the Victorian Medical Postgraduate Foundation Therapeutics Committee.

Last but not least we would like to thank the Ministry of Health for their support and assistance in the preparation of this document.



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These guidelines have been reviewed by the Central/ Eastern Drug and Therapeutics Committee for the Ministry of Health, for usage at its Health Care Institutions.

The aim of the guidelines is to rationalise the usage of antibiotics on our Essential Drug List and to establish consistency in the treatment of various infectious conditions.

The recommendations for the treatment of paediatric infections have been kept separate for easy referral, located at the back of the booklet.

In addition four separate chapters have been included on treatment of anaphylactic reactions, tuberculosis, opportunistic infections in patients with HIV and antibiotic usage during pregnancy and breastfeeding.

Great care has been taken in preparing these guidelines, however they may not be free from errors. It is the responsibility of the prescriber to ensure that the correct dosage is prescribed; if there is any doubt you are advised to check the manufacturer's dosage recommendations.

These are general guidelines, treatment of individual patients may vary depending upon local conditions and experience.

Useful suggestions for the improvement of these guidelines will be appreciated.

Chairman National Drugs & Therapeutics Committee 2003







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## 1. ABOUT ANTIMICROBIAL IN GENERAL

The number of antimicrobials available for the therapy has increased rapidly during the past few years. Brief summaries of antimicrobial agents commonly used in therapy are presented in this chapter.

ANTIBACTERIAL AGENTS
These are grouped based on their site of action on the bacteria, as given below:

## **CELL WALL SYNTHESIS**

#### **BETA-LACTAMS**

- Penicillins
- Cephalosporins Carbapenems
- Monobactams

## **GLYCOPEPTIDES**

Vancomycin

- - Lincosamides (e.g. Clindamycin)
    Macrolides (e.g. Erythromycin)
    Tetracyclines

## RNA POLYMERASE

Rifampicin

## <u>DNA</u>

- Quinolones (e.g. Nalidixic acid, Ciprofloxacin)
- Metronidazole
- Nitrofurantoin

### **FOLATE METABOLISM**

- Trimethoprim Sulphonamides





#### **BETA - LACTAM ANTIBIOTICS**

Penicillins, cephalosporins, monobactams and carbapenems have a  $\beta\mbox{-lactam}$  ring in their molecular structure. These bactericidal antibiotics act primarily on the bacterial cell wall. Although some bacteria produce  $\beta$ -lactamase and therefore have developed resistance, these drugs on the whole remain useful in treating many different types of infections

#### **Penicillins**

Penicillin is active against Streptococci, Neisserriae, Spirochaetes, some anaerobes including *Clostridia* and a few other organisms. In Fiji, about 80 - 90% of the *Staphylococcus aureus* are  $\beta$ -lactamase producers and hence are resistant to penicillin G and aminopenicillins. The prevalence of penicillinase producing Neisseria gonorrhoea is on the increase. There are reports of decreased susceptibility of pneumococci and streptococci to penicillin from other parts of the world. The only serious disadvantage of penicillins is hypersensitivity reaction.

## A. Penicillin formulations available are:

- 1. Penicillin G (Crystalline Penicillin or Benzyl Penicillin) for intravenous (IV) use. Needs to be given frequently (4 – 6 hourly).

  2. Procaine Penicillin – Intramuscular (IM) preparation with a longer duration
- of action. Needs to be administered less frequently i.e. daily.

  3. Benzathine Penicillin given IM provides low levels of penicillin in the
- circulation for 3-4 weeks.
- 4. Penicillin V (Phenoxymethyl Penicillin) an oral preparation, intrinsically less active than Penicillin G

Penicillin is the drug of choice for the treatment of the following infections:

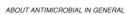
- 1. Streptococcal infections e.g. tonsillopharyngitis
- 2. Infections due to Streptococcus pneumoniae.
- 3. Meningococcal infections e.g. meningitis, septicaemia
- 4. Syphilis
- 5. Clostridial infections, anthrax, diphtheria
- 6. Leptospirosis

#### B. Aminopenicillins

Ampicillin and amoxycillin are destroyed by staphlococcal \( \beta \)-lactamases but have a slightly broader spectrum than penicillins because of their activity against some gram negative bacilli like E.coli, salmonella sp and shigella sp. They also have better activity against H.influenzae and enterococci compared with penicillin.

Although initially sensitive, resistance to these drugs among *E.coli* is now widespread.





Many strains of H.influenzae also produce  $\beta$ -lactamases, which can destroy these drugs.

Amoxycillin is better absorbed than ampicillin and has a longer half life and hence is preferred for oral therapy. These drugs are used in empirical treatment of respiratory infections and in the treatment of susceptible urinary tract infections. They may be used for typhoid fever.

## C. Anti -Staphylococcal Penicillins

These are narrow spectrum penicillins, resistant to Staphylococcal  $\beta$ -lactamases. Methicillin, oxacillin, and cloxacillins fall into this category. Of these only cloxacillin, flucloxacillin and dicloxacillin are clinically useful and are to be used only for proven or suspected staphylococcal infections.

Flucloxacillin, suitable for oral administration, can cause cholestatic jaundice in some patients.

Some staphylococci have developed resistance to this group, by the mechanisms other than lactamase. These methicillin resistant *Staphylococcus aureus* (MRSA) will be resistant to all other lactams (i.e. all penicillin, cephalosporins, monobactams and carbapenems).

### <del>)</del>

## D. Anti - Pseudomonal Penicillins

Newer penicillins with a high grade of activity against gram negative bacteria including pseudomonas, e.g. piperacillin, ticarcillin\*

#### $\underline{\textbf{E.}} \hspace{0.2cm} \beta \underline{\textbf{-lactam and }} \beta \underline{\textbf{-lactamase inhibitor combinations}}$

Eg Clavunate, Sulbactam\*

Augmentin is a preparation containing amoxycillin and clavulanic acid. Clavulanic acid has minimal antibacterial activity but inhibits  $\beta$ -lactamase effectively. This combination is useful in the treatment of  $\beta$ -lactamase producing bacteria. Sulbactam\* is another  $\beta$ -lactam inhibitor used in combination with penicillins. Combinations are more expensive and so should be used only while treating infections with known  $\beta$ -lactamase producers. Amoxycillin/ clavulanic acid combination can cause cholestasis.

Note: Hypersensitivity to any penicillin implies hypersensitivity to all penicillins. 5-10% of patients with Penicillin hypersensitivity, especially those with early manifestations, are also hypersensitive to cephalosporins.

\* NOT ON EDL





#### Cephalosporins and related drugs

The cephalosporins have been traditionally divided into to three "generations" based on their spectrum of activity. In general, cephalosporins are less prone to hypersensitivity reactions, are more stable to staphylococcal penicillinases and have a broader spectrum than penicillins.

However, they are expensive and have very little action on enterococci. **None of them are effective against MRSA**. Cephalosporins also have been shown to select MRSA, vancomycin resistant enterococci and ceftriaxone resistant gram-negative bacilli. Therefore, indications for their use should be limited.

First generation cephalopsorins include among others, cephalexins \* (oral), cephalothin and cefazolin \* (parenteral). The spectrum of activity is similar, being effective against penicillinase producing staphylococci and other Gram-positive cocci (except MRSA and enterococci) and a few gram- negative enteric bacilli. There is no special advantage for any one first generation cephalosporins over another. They are not usually first choice for any infection. They may be used in some patients with penicillin hypersensitivity.

Cephamandole\* (parenteral), cefuroxime axetil\* and cefaclor (oral) are "second generation drugs" which are more stable to some Gram-negative  $\beta$ -lactamase. Their activity against Gram-positive organisms is similar to, or less than, that of the first generation cephalosporins and they have varying degrees of activity against anaerobes. These drugs have a limited role in therapy and are more expensive.

The major activity of the third generation cephalosporins (ceftriaxone, ceftazidime\*, cefotaxime\*) is against gram-negative bacilli. They have some activity on gram-positive cocci and that against anaerobes varies. A major advantage of these agents is their ability to reach the central nervous system. Ceftazidime\* has specific antipseudomonal activity. Ceftriaxone and cefotaxime\* are useful in hospital-acquired and any other gram-negative septicaemia and meningitis.

## $\underline{\textbf{Monobactams}}$ (Aztreonam\*) and $\underline{\textbf{Carbapenem}}$ (Imipenem\*)

Aztreonam is active against gram-negative bacteria including pseudomonas and \_-lactamase producing enterobacteriaceae. Carbapenems have a much broader spectrum, including gram-positive, gram-negative and some anaerobic bacteria.

### **Aminoglycosides**

This group of antibiotics (gentamicin, tobramycin\*, netilmicin\*, amikacin\*, Kanamycin\*, neomycin, streptomycin) act by inhibiting protein synthesis in bacteria. They have good activity against aerobic gram-negative bacilli, including brucella. When given together with penicillin, they have good activity against enterococci. Streptomycin

\* NOT ON EDL









ABOUT ANTIMICROBIAL IN GENERAL is useful against my bacteria. Aminoglycosides are not absorbed when given orally and should be administered parenterally for systemic effects.

Aminoglycosides are ototoxic and nephrotoxic. The therapeutic index is low and blood levels need to be monitored. In spite of this disadvantage, they are used widely for their action on gram-negative bacilli. Gentamicin is the least expensive and has good activity against 90% of gram-negative bacilli isolated in Fiji. It is the aminoglycoside of choice for empirical treatment of severe gram-negative sepsis including nosocomial infections.

Recent studies have shown that aminoglycoside may be given in a single daily dose, having efficacy similar to that of multiple dose and toxic effects may be lower. The recommended dose is  $5-7\,\mathrm{mg/kg/day}$  (based on lean body weight) as a single daily dose. This regimen is now preferred in many places. Monitoring is advisable if treatment is for more than 3 days. However single daily dose is not recommended in pregnant women and endocarditis. In patients with impaired renal function, doses and/or frequency of administration must be modified according to serum levels and the degree of impairment.

Whenever possible it is desirable to determine peak and trough serum gentamicin levels periodically and to adjust the dosage to maintain the desirable serum levels. In general, peak levels of 4 to 10 g/mL and trough levels not exceeding 0.5 to 1 g/mL are sought. Note: The loading dose is usually 2.5 - 7 mg/kg irrespective of renal function and should be given slowly over 20 minutes. Subsequent doses must be adjusted according to renal function.

Bacterial resistance to gentamicin is mainly mediated by three different types of enzymes that destroy the drug. Netilmicin\* and amikamicin\* are destroyed by fewer enzymes and therefore have a broader spectra. These drugs should only be used when the bacteria under question is proven to be resistant to gentamicin. Netilmicin\* may be less nephrotoxic.

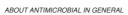
#### **Tetracyclines**

Tetracyclines also act by inhibiting protein synthesis and have broad spectrum of activity. This includes staphylococci, neisseriae, *H.influenzae*, some members of enterobacteriaceae, mycoplasma, clamydiae, rickettsiae and spirochaetes. For chlamydial and rickettsial infections this is the drug of choice.

This group also has action against protozoa like Entamoeba histolytica and plasmodium sp.

The spectrum of activity of different tetracyclines is similar, but they are different in their pharmacokinetics. Most tetracyclines are excreted through the kidneys except doxycycline, which is safer in patients with renal impairment, but caution is required in patients with hepatic disease. Tetracycline should be used with caution in patients with pre-existing hepatic or renal disease, as they can lead to worsening of function. Doxycycline has a longer half-life than tetracycline.

Because of their effect on growing bones and teeth, these drugs are contraindicated in pregnancy, lactating mothers and in children.



#### Chloramphenicol

Also a broad-spectrum antibiotic, it acts by inhibiting protein synthesis. The spectrum includes both aerobes and anaerobes. It can be used topically, orally or parentally. Bioavailability after oral administration is as good as parenteral use and the oral preparation can be used to initiate treatment in emergencies if the injection is not available. Chloramphenicol is not safe in pregnancy and in neonates as it may cause Grey baby syndrome. This drug can also cause bone marrow suppression. Its use as far as possible should be limited to specific indications like typhoid fever, invasive salmonellosis, meningitis, brain abscess and occasionally anaerobic infections.

#### Macrolides

Erythromycin, roxithromycin\*, azithromycin\* and clarithromycin\* act by inhibiting protein synthesis. They have similar antimicrobial spectra but differ in their pharmacokinetics and adverse effects.

They are active against gram-positive organisms, *H.influenzae*, neisseriae, mycoplasma sp, chlamydia and rikettsiae. They also act on toxoplasma, which is a protozoa

Erythromycin is absorbed orally and is distributed well. It does not cross the blood brain barrier. The main adverse reaction is gastric irritation. Some patients develop jaundice. Parenteral preparations can cause phlebitis and occasional cardiac arrhythmias (in high doses).

Its main use is in respiratory infections and as an alternative to penicillin in those hypersensitive to penicillin. It is the drug of choice in neonatal and obstetric chlamydial infection and is used in camphylobacter infection. The newer macrolides have better bioavailability and fewer side effects. Azithromycin\*, in addition to its use similar to that of erythromycin, is also used to treat toxoplasmosis. Clarithromycin\* is used in treating mycobacterium avium complex (MAC) infections and *H.Pylori*.

#### Quinolones

These drugs interfere with transcription of DNA. The first drug to be used in this group, nalidixic acid, had a very narrow spectrum mainly limited to gram-negative bacilli. This drug is used widely in the treatment of UTI, since it concentrates in the urine.

The newer Quinolones, norfloxacin\* and ciprofloxacin have a broader spectrum of activity. Norfloxacin is used in the treatment of urinary and gastrointestinal infections. Ciprofloxacin reaches high levels in the blood and is very effective against enterobacteriaceae, pseudomonas sp, and mycobacteria. Ciprofloxacin is not very effective against Streptococcus pneumoniae. It is therefore useful in treating gram-negative infections like hospital acquired septicaemias and gram-negative pneumonias. It is also useful in treating chloramphenicol resistant Salmonella typhi infections. Bacterial resistance







#### ABOUT ANTIMICROBIAL IN GENERAL

develops rapidly if these agents are widely used.

They are well absorbed when given orally and have a good penetration into cells like macrophages. They do not cross the blood brain-barrier. Unlike many other antibiotics they reach the prostate.

Nalidixic acid\* can cause GI upset and skin reactions. Quinolones have many adverse effects including dizziness, depression, and they can precipitate seizures. They may interact with many other drugs, for example theophylline.

Quinolones/ Ciprofloxacin is not recommended in pregnant women, infants, children and breastfeeding mothers.

#### Rifampicin

Rifampicin is used in the treatment of tuberculosis and infections with *S.aureus*. Rifabutin\* is used in the treatment and prophylaxis of MAC infection. Since resistance emerges rapidly, these drugs should always be used in combination with other antibiotics.

Rifampicin colours urine, tears and other body fluids red. It can accelerate the metabolism of other drugs including oral contraceptives, warfarin, and phenytoin.

#### **Nitroimidazoles**

Metronidazole and tinidazole\* are active against all anaerobic bacteria and protozoa like *T. vaginalis, G. lamblia* and *E. hystolitica*. Metronidazole is well absorbed and can be administered IV, orally or rectally. The rectal preparation produces high levels and can be used to treat serious infections. It crosses the blood brain barrier.

Metronidazole is usually well tolerated. Common minor side effects include nausea, vomiting, metallic taste in mouth and disulfiram like reaction with alcohol.

Tinidazole has longer half-life and therefore can be administered less frequently.

### Glycopeptides

Vancomycin acts by inhibiting peptidoglycan (cell wall) synthesis. All gram-positive organisms are susceptible. However, the drug is reserved for treating Gram-positive infections resistant to  $\beta$ -lactams (MRSA and Ampicillin/ Gentamicin resistant enterococci) and some patients hypersensitive to  $\beta$ -lactams. Its oral use for antibiotic associated diarrhoea should be limited to those caused by <code>Clostridium difficule</code> and unresponsive to metronidazole.



\* NOT ON EDL





#### ABOUT ANTIMICROBIAL IN GENERAL

Vancomycin is given IV slowly over at least over one hour (10mg/ min) to prevent anaphylactoid reaction. Renal toxicity can occur, especially if given with aminoglycosides. Therefore, pay attention to dosage schedules and monitor serum levels and renal function.

## ANTIVIRAL AGENTS

Several agents are now licensed for use; only Lamivudine tablets, Zidovudine (AZT) elixir and tablets, acyclovir injection and eye ointment are available on the Fiji EDL.

Acyclovir is a nucleoside analogue, effective against Herpes Simplex virus types I and II and Varicella zoster virus. Topical application or oral therapy is used for skin, mucous membranes and eye infections. Parenteral use is indicated for herpes encephalitis.

 $\label{eq:famciclovir*} \textit{Famciclovir*}, \textit{penciclovir*} \textit{ and } \textit{valaciclovir*} \textit{ are recent modifications of acyclovir}.$ 

Other Antiviral Drugs	Cinical Use
Amantadine*	Prevention and treatment of influenza A, during
	outbreaks
Ganciclovir*	CMV infection in the immunocompromised
Tribavirin*	Effective against many viruses. Used for
	treatment of RSV infection in small children,
	Lassa fever and hanta virus infection
Zidovudine (AZT)	Treatment & post-exposure prevention of HIV/
Lamivudine	AIDS
Dideoxyinosine* (Didanosine)	
Dideoxycytosine* (Zalcitabine)	
Protease inhibitors:	Treatment & post- exposure prevention of HIV/
Saquinavir*	AIDS
Indinavir*	
Interferon alpha*	Treatment of hepatitis B, hepatitis C
Idoxuridine*	Topical treatment for HSV

At present in Fiji, anti HIV drugs are available on the EDL for post exposure prophylaxis and prevention of vertical transmission only.



\* NOT ON EDL











#### ABOUT ANTIMICROBIAL IN GENERAL

#### **ANTIFUNGAL AGENTS**

## <u>Azoles</u>

There are two main groups.

#### a. For systemic Use: Ketoconazole, fluconazole \* and itraconazole\*

Fluconazole\* and ketoconazole are active against yeasts like candida, crytococci, and histoplasma. These agents are useful in the treatment of systemic infections due to these organisms.

 $\label{eq:flucon} \mbox{Fluconazole*} \mbox{ is well-absorbed following oral administration and has good CNS penetration.}$ 

Ketoconazole can cause hepatic toxicity.

Itraconazole\* has activity against asperillus.

## b. For topical Use: Miconazole\*, clotrimazole\* and econazole.

These are used in the treatment of superficial candidiasis and dermatophytosis.



## Amphotericin B

It is useful against most systemic fungal infections. Its use is associated with many side effects including flu-like reactions, GI effects and nephrotoxicity. It should be given IV slowly over 4-6 hours.

#### Flucytosine\*

This drug is used mainly in combination with amphotericin in the treatment of systemic candida and cryptococcal infections. Adverse effects include bone marrow suppression and hepatotoxicity.

### Griseofulvin

When given orally, it concentrates in keratinised tissues and prevent further invasion by dermatophytes.

#### Terbinafine\*

A new drug effective against dermatophytes when used orally or topically.

## Whitfield Ointment/Lotion (salicylic acid 10%)

Still the mainstay of treatment of dermatophytosis in Fiji.

#### Nystatin

Suspension is useful for oral thrush.

\* NOT ON EDL







## 2. GENERAL GUDELINES FOR USE OF ANTIBIOTICS

#### PRUDENT USE OF ANTIBIOTICS

#### Genera

Antibiotics should be used only where the benefits are scientifically demonstrable and substantial.

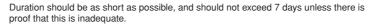
In general, the antimicrobial spectrum of the drug selected should be the narrowest to cover the known or likely pathogen(s).

Single agents should be used unless it has been proven that combination therapy is required to ensure efficacy or reduce the selection of clinically significant resistance.

The dosage should be high enough to ensure efficacy and minimise the risk of resistance selection, and low enough to minimise the risk of dose related toxicity

#### Therapy

Choice of therapy should be based on either cluture and susceptibility tests results (directed therapy), or known common pathogens in the condition and their current resistance patterns (empirical therapy).



#### Prophylaxis

Choice should be based on known or likely target pathogen(s).

Duration should be as short as possible. A single dose of antibiotic is recommended for surgical prophylaxis. Longer-term prophylaxis should be administered only when it has been demonstrated that the benefits outweigh the risk of resistance selection or propagation.

REF: Australian Therapeutic Guidelines - Antibiotic, version 11, 2000

Antibiotics are frequently misused. This misuse not only causes unwanted reactions in the recipient, it also favours proliferation of resistant organisms. Unnecessary wastage of resources is another result.

There are two main situations where antibiotics are used:

- 1. For treatment of proven or suspected bacterial infections.
- 2. For prevention of bacterial infections.





#### 2.1 Treatment Guidelines

- (a) Decide whether an antibiotic is really indicated.
- (b) If an antibiotic is indicated, the choice will depend on:
  - (i) Efficacy

This depends on the spectrum of activity of the drug in relation to known or suspected causative agents. Whenever possible, antibiotic therapy should be guided by in vitro susceptibility tests. Knowledge about local susceptibility patterns of organisms frequently causing infection and previous clinical experience help in deciding on empirical, when this option is essential. Choose antibiotic with narrowest spectrum when applicable.

As far as possible, collect samples for culture before starting empirical therapy AND review the empirical therapy when susceptibility results are available. In some situations, a gram stain (e.g. pus from deep abscesses) or directs antigen testing (e.g. meningitis) can help in initiating early directed therapy

Efficacy also depends on pharmacokinetics factors like absorption, distribution (penetration into tissue etc.) metabolism and excretion of the drug.

- (ii) Safety
   Potential adverse effects and risk of super infection should be considered.
- (iii) Costs.
- (iv) Potential for selection of resistant organisms.

## (c) Route of administration

This is another important consideration. Parenteral preparations are generally more expensive. These are indicated when the patient is seriously ill so oral absorption may be impaired, when large doses are required, when the drug has poor bioavailability when given orally, or the patient is unable to take oral medication. Reassess parenteral therapy daily, and switch to oral therapy as soon as possible.

Topical antibiotic use should be limited to a few indications where its efficacy is proven, such as eye infections. Widespread and inappropriate use of topical antibiotics can cause sensitisation in the patient and favours development of bacterial resistance

- (d) Combinations of antibiotics should be used only when indicated:
  - (v) To extend the spectrum of cover in empirical therapy of suspected mixed infections or in life threatening infections.
  - (vi) To achieve synergistic bactericidal effect e.g. \_ lactam and gentamicin for treatment of enterococcal infection.
  - (vii) To prevent the emergence of resistant organisms e.g. treatment of  $% \left\{ 1\right\} =\left\{ 1\right\} =\left\{$

## GENERAL GUDELINES FOR USE OF ANTIBIOTICS

#### tuberculosis

- (viii) To reduce dosage of a toxic drug e.g. amphotericin and flucytosine in the treatment of fungal infections.
- (e) Duration of therapy varies depending on the nature of infection and the causative organism.
- (f) The following IV antimicobials are given by **slow infusion**:
  Gentamicin loading dose over 20 30 minutes
  Vancomycin over 1 hour (10mg/ minute)

Metronidazole – over 20 minutes Erythromycin – over 1 hour

Acyclovir – over one hour Amphotericin – over 4 – 6 hours

Ciprofloxacin - over 1 hour

Note: In paediatric practice all IV antimicrobials are given by slow infusion to avoid thrombophlebitis.

#### 2.1 Prophylactic use of Antibiotics

Indications for these are very limited and are discussed in detail in appropriate chapters.

## Stategies for antimicrobial use in an institution

To prevent misuse or abuse of drugs, the following steps are recommended:

- 1. Each hospital should have a drug committee to formulate policies suitable for that hospital.
- Antimicrobials available in the hospital should be categorised into a restricted and unrestricted list based on cost, safety and emergence of antimicrobial resistance. The restricted antibiotics in Fiji are:

Ceftriaxone	InjAmphotericin B Inj	Vancomycin Inj
Ciprofloxacin Inj and Tabs	Piperacillin Inj	Erythromycin Inj
Metronidazole Inj	Fusidic Acid Tabs	Acyclovir Inj

## (Note: This list is subject to change in future)

These drugs can be obtained only after a consultant approves. Initial supply will be limited to 48 hours for empirical therapy.

- 3. Audit antimicrobials, by product and by clinical unit on a routine basis (or by individual clinicians when indicated).
- Continuing education for medical and paramedical staff on new drugs and the changing susceptibility patterns.
- Microbiology investigation e.g. blood culture, wound swab, must be taken prior to commencement of an antimicrobial. Prior cleaning of wounds with normal saline should be done when taking wound swabs.









#### 3. CARDIOVASCULAR SYSTEM INFECTIONS

#### **BACTERIAL ENDOCARDITIS**

There are 3 important principles of management:

- 1. Treatment must be given intravenously except in rare cases
- 2. Treatment is prolonged usually 4 -6 weeks.
- 3. Drug regimens should be used in high enough concentrations.

There are several alternative regimens reported in the literature and consultation with a physician or a microbiologist should be sought if necessary. Surgical consultation should be considered especially in cases, which are fulminating, complicated or slow to respond.

If bacterial endocarditis is suspected taking at least 3 bloodcultures from 3 different venipuncture sites is advisable before initiating therapy.

#### 3.1 Empirical Therapy

- Penicillin G 1.8g (3 mega units) IV 4 hourly PLUS
- Cloxacillin 2g IV 4 hourly PLUS
- Gentamicin 80mg IV 8 hourly (maintenance dose adjusted for renal function).

In patients hypersensitive to penicillin, in hospital acquired endocarditis, and if cardiac prosthetic valve in situ use:

- Vancomycin 1g IV 12 hourly, each dose to be infused over at least 1 hour (dose adjusted for renal function), PLUS
- Gentamicin 80mg IV 8 hourly (maintenance dose adjusted for renal function).

These regimens may need to be modified when the organism and its sensitivity pattern is known. If culture is negative, continue with the same regimen, if the patient is responding.

### 3.2 Specific Therapy

#### 3.2.1 Streptococcal Endocarditis

Streptococcus viridans is generally highly sensitive to penicillin G.

- Penicillin G 1.8g (3 mega units) IV 4 hourly for 4 weeks PLUS
- Gentamicin 80mg IV 8 hourly for 2 weeks (maintenance dose adjusted for renal function).

For patients with hypersensitivity to penicillin

- Cephalothin 2g IV 4 6 hourly X 4 weeks PLUS
- Gentamicin 80mg 8 hourly X 2 weeks

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#### CARDIOVASCULAR SYSTEM INFECTIONS

Note: A longer course of 4-6 weeks of both penicillin and gentamicin therapy is advisable if relative resistance to penicillin is suspected. However, currently Streptococcus is highly sensitive to penicillin in Fiji.

Also in suitable patients responding to treatment well after the initial therapy, a home based therapy with ceftriaxone 2g IV daily to complete a course of 4 weeks can be considered.

#### 3.2.2. Enterococcal Endocarditis

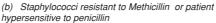
- Penicillin G 1.8g (3 mega units) IV 4 hourly for 6 weeks **PLUS** Gentamicin 80mg IV 8 hourly for 6 weeks (maintenance dose adjusted for renal function).

For resistance or patients hypersensitivity to Penicillin use
• Vancomycin **PLUS** Gentamicin, as for empirical treatment, for 6 weeks.

#### 3.2.3. Staphylococcal Endocarditis

- (a) Staphylococcus aureus sensitive to Methicillin and Cloxacillin
  - Cloxacillin 2g IV 4 hourly, for 6 weeks

Traditionally gentamicin 1mg/kg intravenously (adjusted for renal function) 8 hourly has been used for the first 3 –5 days as there is some evidence that it reduces the duration of bacteraemia. This is often followed by rifampicin though the evidence is lacking.



Vancomycin 1g IV 12 hourly, for 6 weeks (to be infused over at least 1 hour).

## 3.2.4 Gram-Negative Bacterial endocarditis

This is uncommon and early consultation with a microbiologist or physician is recommended.

## 3.2.5. Culture Negative Endocarditis

This maybe due to previous antibiotic use or unusual micro – organisms, such as fastidious streptococci, legionella sp., or fungi including Candida albicans. Unless fungal infection is strongly suspected, patients with culture negative endocarditis should be treated empirically with Pencillin G plus Gentamicin, for a period of 6 weeks.







#### 3.2.6. Prosthetic Valve Endocarditis

Can be due to low virulent members of normal flora that may be resistant to penicillin or cloxacillin

- (a) Empirical therapy
- Vancomycin 1g IV 12 hourly (to be infused over at least one hour) for 6 weeks PLUS
- Gentamicin 80 mg IV 8 hourly for 2 weeks (dose adjusted for renal function)
- (b) Specific therapy as for native valve endocarditis

#### 3.3 Monitoring Antibiotic Therapy.

Particular attention should be given to Therapeutic Drug Monitoring in Endocarditis. Recommended doses are for the commencement of treatment only and may need to be modified according to plasma levels attained.

#### Gentamicin



Gentamicin levels should be monitored, however the optimum method of monitoring levels in endocarditis is unknown. As doses are lower, dosing more frequent and synergy is the objective, the methods of monitoring used for gentamicin in other circumstances are inappropriate. Instead troughs should be measured only and vaules between 0.5 and 1 mg/L will likely maximise synergy and minimise toxicity with 8 hourly dosing. Patients should be monitored for vestibular and auditory ototoxicity

## Vancomycin

Vancomycin peak and troughs concentration should be first measured at 48hrs to 72 hrs, although a steady state may not have been reached at this time. Peak vancomycin levels of 30 to 40mg/L and trough levels of 5 to 15mg/L are recommended. These levels are specific for the management of endocarditis and do not necessarily apply to other circumstances.

Note: Currently only gentamicin levels can be measured in some laboratories in Fiji. Vancomycin levels are currently not measured in Fiji.





#### 4. RESPIRATORY TRACT INFECTIONS

#### 4.1 ACUTE UPPER RESPIRATORY TRACT INFECTIONS

#### 4. 1.1 GENERALISED INFECTIONS

The aetiology is usually viral. Most common symptoms subside in a few days, although cough may persist longer, ie 14 days or more. For rhinorrhoea, topical or oral decongestants provide some relief.

 Oxymetazoline\* OR Xylometazoline\* 2 to 3 drops into each nostril 2 to 3 times daily for not more than 5 days.

Paracetamol plus adequate hydration gives symptomatic relief of fever and discomfort. Patients require appropriate reassurance that usually antibiotics are unhelpful and are often harmful by causing side effects and increasing bacterial resistance.

#### 4.1.2 ACUTE SORE THROAT

Usually viral. Paracetamol, aspirin (in adults) or ibuprofen provide valuable symptomatic relief. Acute tonsillitis caused by *Streptococcus pyogenes* is difficult to distinguish clinically from viral aetiology, but is more likely if patients have all of the following features:

- Fever > 38° C
- Tender cervical lymphadenopathy
- Tonsillar swelling or exudates
- No cough.

If bacterial infection is suspected, use

- Penicillin V 500mg orally 12 hourly for 10 days OR
- Benzathine Penicillin 1.2 mega units IM as a single dose

Although Amoxycillin is widely used penicillin V has a narrow spectrum and covers Streptococcus pyogenes which is the organism of concern, therefore its use is discouraged.

Alternative/ hypersensitive to Penicillin:

- Erythromycin 500mg orally 12 hourly for 10 days OR
- Erythromycin 250mg orally 6 hourly for 10 days

NOTE It is important to complete the course of therapy to prevent rheumatic fever, which is still a common problem in Fiji

\* NOT ON EDL









#### 4.1.3 ACUTE BACTERIAL OTITIS MEDIA

This is either viral or bacterial and usually self-limiting disease. Symptomatic therapy such as pain relief should be provided for the first 2 days.

If symptoms persists or suppurative complications develop then consider antibiotics

Amoxycillin 500mg orally 8 hourly x 7 days

In complicated cases refer to ENT specialist.

#### 4.1.4 OTITIS EXTERNA

Usual organisms include Pseudomonas aeruginosa, Staphylococcus aureus, Proteus and Klebsiella species.

Ear should be kept dry followed by instillation of topical steroid and antibiotic combination drops.

Indications for systemic antibiotics include fever, spread of inflammation to the pinna or folliculitis.

• Flucloxacillin 500mg orally 6 hourly for 5-7 days

In complex cases such as development of mastoiditis, hearing loss, refer to specialist.

#### 4.1.5 ACUTE SINUSITIS

Acute bacterial sinusitis follows upper (usually viral) respiratory tract infections in between 0.5 to 5% of cases. It is caused by Streptococcus pneumoniae or H. influenzae. Consider antibiotic therapy in severe cases displaying at least three of the following features:

- persistent mucopurulent nasal discharge (> 7-10 days)
- facial pain
- tenderness over the sinuses
- poor response to decongestants
  - Amoxycillin 500mg orally 8 hourly for 7 days.





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In patients hypersensitive to Penicillin

- Tetracycline 500mg orally 6 hourly **OR**
- Doxycycline 100mg orally 12 hourly. **OR** Cefaclor SR 375mg orally 12 hourly (for inpatient use)

Duration of antibiotic therapy should be for 7 – 10 days depending upon response.

## 4.1.6 ACUTE EPIGLOTTITIS

Seen mostly in children, may occur in adults. Urgent hospitalisation is required. Commonly caused by *H influenzae* and Streptococci

• Chloramphenicol 750mg – 1g IV 6 hourly for 5 – 10 days.

#### Alternative:

- Ceftriaxone 2g IV as a single dose daily for 5- 10 days (inpatient use only) OR
- Ampicillin 500mg 1g IV 6 hourly for 5-10 days (when sensitivity known).

Early transfer to oral therapy is desirable.

#### 4.2 LOWER RESPIRATORY TRACT INFECTIONS

#### 4.2.1 ACUTE BRONCHITIS

In an immunocompetent adult or child, acute bronchitis is most often viral and does not require antibiotic therapy. Randomised controlled trials show that antibiotic therapy provides no overall benefit to the patient and may cause harm.

If severe and particularly if sputum is voluminous and purulent, with fever, secondary bacterial infections is assumed:

Amoxycillin 500mg orally 8 hourly for 5 -7 days.

### Alternative:

- Doxycycline 100mg orally 12 hourly for 7 days (if available) **OR** Tetracycline 500mg orally 6 hourly for 5-7 days **OR** Erythromycin 500mg orally 6 hourly for 5-7 days

## 4.2.2 ACUTE EXACERBATION OF CHRONIC BRONCHITIS

Acute exacerbation could be either viral or bacterial infection. Common organisms include Strep pneumoniae, H Influenzae, Moraxella catarrhalis. The indication for Antibiotic therapy is increased cough and dyspnoea TOGETHER with increased sputum volume and/or Purulence



Treatment as for Acute Bronchitis above

#### 4.2.3 PNEUMONIA

#### 4.2.3.1 COMMUNITY ACQUIRED

Choice of antibiotic is often empirical. In immunocompetent, otherwise healthy patients it is usually caused by single microorganisms such as Streptococcus pneumoniae, H. influenzae, Mycoplasma pneumoniae, Chlamydia. Staph aureus is not uncommon in Fiji.

Note: In immunocompromised patients such as diabetics or in elderly patient or patients with co-existent illness (e.g. cancer, liver disease, heart failure or renal failure) a broad-spectrum antibiotic cover may be required.

### (a) Mild Disease

- Amoxycillin 500mg orally 8 hourly for 7 10 days OR
- Procaine Penicillin 3.6g (4 mega units) IM daily for 7 10 days.

#### Alternatives:

- Tetracycline 500mg orally 6 hourly for 7 10 days OR
- Doxycycline 100mg orally 12 hourly for 7 –10 days

If hypersensitive to penicillin or Mycoplasma or Chlamydia suspected:

- Erythromycin 500mg orally 6 hourly for 10 14 days **OR**Doxycycline 100mg orally 12 hourly for 10 14 days

(b) Moderate Disease
• Penicillin G 1200mg (2 mega unit) IV 6 hourly for 7 -10 days

In patients hypersensitive to penicillin

Chloramphenicol 1g IV 6 hourly X 7 -10 days

If the clinical response to parenteral therapy is satisfactory, high dose oral therapy may be substituted after a few days e.g. Amoxycillin 0.5 – 1.0g orally 8 hourly or to Chloramphenicol 0.5g – 1g orally 6 hourly for patients on chloramphenicol

## (c) Severe Disease

In Adults, severe pneumonia should be suspected if the following features are present;

- respiratory rate > 30 per minute
- PaO<sub>a</sub> < 60mmHg or SaO<sub>a</sub> < 90% on room air
- PaCO<sub>2</sub> > 50mmHg on room air Chest X-ray evidence of bilateral involvement or involvement







## of multiple lobes

- Increase in size of chest X-ray opacity by 50% or more within 48 hours of admission
- Requirement for mechanical ventilation or inspired oxygen

#### > 35% to maintain $Sa0_2 > 90\%$ Haemodynamic compromise:

- systolic blood pressure < 90mmHg
- diastolic blood pressure < 60mmHg
- recent deterioration in renal function white blood cell count <4 or >30 x  $10^9$ /L

#### Empirical:

- Penicillin G 1200mg (2 mega unit) IV 6 hourly **PLUS**Cloxacillin 2g IV 6 hourly **PLUS**Gentamicin 4 6 mg/kg IV once daily (maintenance dose adjusted according to renal function)

If severe or no response then the following may be added

Erythromycin IV/ orally 500mg 6 hourly

#### If hypersensitive to penicillin

- Ceftriaxone 2g IV daily **PLUS**Erythromycin 0.5 1g IV (slow infusion over 1 hour) 6 hourly

Definitive therapy should be instituted based on bacteriological data.

If Streptococcus anginosus is proven
• Penicillin G 3 - 4 mega (1800 - 2400mg) IV 6 hourly X 21 days

#### If Pseudomonas aeruginosa is proven

- Piperacillin 4g IV 6 hourly PLUS
- Gentamicin 240mg IV once daily (to be adjusted for renal function)

If Staphylococcus. aureus is proven
• Cloxacillin 2g IV 6 hourly for 3 to 4 weeks (oral therapy can be substituted once patient's condition is stabilised)

All severe pneumonia requires consultant advice.

## 4.2.3.2 HOSPITAL ACQUIRED PNEUMONIA

Refers to pneumonia not present at the time of admission and developing in patients after 48 hours of hospitalisation. It is usually due to Gram -ve organisms but Staphylococcus aureus is not uncommon in Fiji.



- Cloxacillin 1g IV 6 hourly for 14-21 days **PLUS**
- Gentamicin 240mg IV once daily for 14-21 days (maintenance dose adjusted for renal function) **PLUS**
- Metronidazole 400mg PO 8 hourly or PR 500mg 12 hourly X 14 21 days

#### 4.2.3.3 NEUTROPENIC PATIENTS/ PATIENTS WITH RESPIRATORY DEVICE OR TRACHEOSTOMY

Gram-negative bacilli including Pseudomonas

- aeruginosa are common causative agents.Piperacillin 3g IV 6 hourly PLUS
  - Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function) PLUS
  - Erythromycin 0.5g 1g IV (infused over one hour) 6 hourly.

Once bacteriological known, modify regimen accordingly. Total duration of therapy is 14 - 21 days.

#### 4.2.3.4 IMMUNOSUPPRESSED PATIENTS

Pneumonia in these patients may be recurrent and due to unusual organisms. A microbiologist or physician should be consulted regarding diagnosis and treatment. REFER to PCP under opportunistic infections in HIV

#### 4.2.3.5 ASPIRATION PNEUMONIA

Minor debris of aspiration pneumonia does not require antibiotic therapy. For severe disease or abscess formation, more prolonged and high dose treatment is indicated.

Streptococcus anginosus, anaerobes and occasionally gram negative bacilli and staphylococci to be considered.

- Penicillin G 1200mg (2mega units) IV 4-6 hourly for 10-14 days **PLUS**
- Metronidazole 400mg orally 8 hourly for 10 14 days.

Alternative for penicillin hypersensitive patients
• Chloramphenicol 500mg – 1g IV / orally 6 hourly.

Note: In addition, flucloxacillin or gentamicin may be required if infection with either staphylococci or aerobic gram negative bacteria suspected or proven. If Streptococcus anginosus is isolated, high dose penicillin will be required and for a longer duration, usually 21 days.









## 4.2.4 LUNG ABSCESS AND EMPHYSEMA

- Penicillin G 2 mega units IV 6 hourly **PLUS** Cloxacillin 2g IV 6 hourly **PLUS** Metronidazole 400mg orally 8 hourly

Gentamicin should be added if aspiration is the likely cause of the abscess. Adequate drainage of empyema is essential. The duration of therapy is usually prolonged.

## 4.2.5 BRONCHIECTASIS WITH INFECTION

## (a) Mild Disease

• Amoxycillin 500mg orally 8 hourly for 7 – 10 days.

#### Alternative:

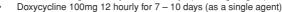
Cefaclor SR 375mg 12 hourly for 7 – 10 days (for inpatient use only).

## (b) Severe Disease

Chloramphenicol 500mg 6 hourly orally /IV for 7 – 10 days

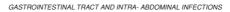
#### Alternative:

- Amoxycillin 500mg orally 8 hourly for 7-10 days **PLUS**Metronidazole 400mg orally 8 hourly for 7-10 days **OR**Cefaclor 375mg 12 hourly for 7 10 days (as a single agent) **OR**Doxycycline 100mg 12 hourly for 7 10 days (as a single agent)









#### 5. GASTROINTESTINAL TRACT AND INTRA- ABDOMINAL INFECTIONS

#### 5.1 CANDIDIAL OESOPHAGITIS

Not generally common in Fiji's population, most likely in immunocompromised patients.

#### Treatment

- Ketoconazole 200mg orally X 14 days OR
- Fluconazole 200mg orally, initially, then 100mg daily for 14 days

## 5.2 DIARRHOEAL DISEASES

Most diarrhoeal diseases are self-limiting and do not require antibiotic therapy. Oral rehydration is all that is required.

Anti-motility agents such as loperamide can be used for symptomatic relief in adults, provided that there is no evidence to suggest invasive disease or obstruction. These agents should NOT be used in children.

Antibiotic therapy is not generally required in the normal host unless there is evidence to suggest invasion with a bacterial pathogen e.g., persistent fever with bloody diarrhoea and/or rigors. However it should be considered for the following and treatment may have to be modified according to the results of culture and sensitivity

## **Bacterial Infections**

**5.2.1** Shigellosis (moderate and severe dysentery only)

Recently in Fiji Shigella has been found to be sensitive to guinolones and cephalosporins only whereas in the past it was more sensitive to chlorampheni col. This may change in future therefore treatment should be guided by sensitivity results.

#### For moderate cases

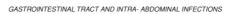
- Chloramphenicol 500mg orally/ IV 6 hourly for 7 days **OR**Cefaclor SR 375mg orally 12 hourly for 7 days **OR**Ciprofloxacin 500mg orally every 12 hours for 7 days (dose adjustment in renal impairment)

#### For severe cases

- Ciprofloxacin IV 200mg (dose adjustment in renal impairment) every 12 hours for 7 Days OR
- Ceftriaxone 2g IV once daily for 7 days







## 5.2.2 Salmonella enteritis

Antibiotic therapy is not generally advisable. Indicated in immunosuppressed or elderly people.

Treat as for typhoid fever (see later) or based on antibiotics sensitivity for 5 days.

#### 5.2.3 Campylobacter enteritis

Antibiotics are unnecessary in most cases. In severe cases;

- Erythromycin 500mg orally 6 hourly for 7 10 days **OR** Ciprofloxacin 500mg 12 hourly for 7 10 days

# **5.2.4** Empirical Therapy As for shigellosis

#### 5.3 PARASITIC INFECTIONS

#### 5.3.1 Intestinal Amoebiasis

Metronidazole 800mg 8 hourly for 6 -10 days

For hepatic involvement, metronidazole should be continued for 14 days.

#### 5.3.2 Giardiasis

Treatment of asymptomatic passage of giardia cysts is unwarranted. For symptomatic patients use;

• Metronidazole 400mg orally 8 hourly for 7 days.

### 5.3.3 Strongyloidiasis

• Thiabendazole 1g orally 12 hourly for 3 days

### 5.4 ANTIBIOTIC ASSOCIATED DIARRHOEA

Cease treatment with any antibiotic likely to be causing the symptoms. Clostridium difficle is responsible for some cases but not all. For mild cases, observe patients after stopping antibiotics

For moderate cases consider:

Metronidazole 400mg orally 8 hourly for 7 - 10 days

For severe and relapsing cases:

- Metronidazole as above and if no response use:
- Vancomycin 125mg orally 6 hourly for 7 -14 days

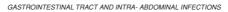
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The emergence of Vancomycin resistant enterococci makes it essential to reserve vancomycin for severe cases unresponsive to Metronidazole Contact Pharmacist-In charge regarding preparation of oral vancomycin.

## 5.5 TYPHOID/ PARATYPHOID FEVER

Check sensitivity pattern with microbiologist. For empirical therapy
Chloramphenicol 500 – 750mg orally 6 hourly for 14 days **OR** 

- Amoxycillin 1g orally 8 hourly for 14 days.

If the organism is found to be resistant to the above, change to:

Ciprofloxacillin 500mg orally 12 hourly for 14 days

#### 5.6 INTRA - ABDOMMINAL INFECTIONS

Example: cholecystitis, cholangitis, diverticulitis, peritonitis and intra – abdominal

Where possible, do culture and sensitivity tests. Surgery may be indicated in addition

Enterobacteriaceae, enterococci and anaerobes are usual pathogens. Strep anginosus (milleri) is sometimes isolated in which case high dose Penicillin G will be required.



- Ampicillin 1g IV 6 hourly for 10 14 days **PLUS**Gentamicin 240mg IV once daily for 10 14 days (maintenance dose adjusted for renal function) **PLUS**
- Metronidazole 500mg suppository PR 12 hourly for 10 14 days OR less severe cases: 400mg orally 8 hourly.

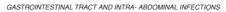
For Streptococcus anginosus

Penicillin G 1.8g (3 megaunit) every 4 hours for 21 days

### 5.7 HELICOBACTER PYLORI INFECTION

All patients with duodenal ulcer, proven Helicobacter pylori associated gastric ulcers or with mucosa associated lymphoma should be treated. Eradication of H pylori in non-ulcer dyspepsia or in asymptomatic patients is not of benefit. Metronidazole resistant H pylori has been noted in many overseas countries but currently in Fiji it is not known to be a problem.





For first line therapy use

- Ranitidine 300mg orally once daily (nocte) X 14 days **PLUS**Amoxycillin 500mg orally every 8 hourly X 14 days **PLUS**Metronidazole 400mg orally every 8 hourly X 14 days

Other regimens with higher efficacy, are more expensive with better compliance include:

- Omeprazole\* 20mg orally 12 hourly X 7 days **PLUS**Clarithromycin\* 500mg orally 12 hourly X 7 days **PLUS**Amoxycillin 1g 12 orally hourly X 7 days **OR**Metronidazole 400mg orally 12 hourly X 7 days

- Omeprazole\* 20mg orally 12 hourly X 14 days **PLUS** Amoxycillin 500mg orally 8 hourly X 14 days **PLUS** Metronidazole 400mg orally 8 hourly X 14 days.

For patients hypersensitive to Penicillin, substitute Amoxycillin with
Tetracycline 500mg every 6 hours

#### 5.8 WORMS (HELMINTHIS)



Hookworm, roundworm, threadworm & whipworm

Mebendazole 100mg orally twice daily x 3 days

Mebendazole should not be used in the  $1^{\rm st}$  trimester of pregnancy or in children under 6 months of age. In such cases Pyrantel can be used.

Pyrantel 20mg/kg up to 750mg orally single dose, repeat after 7 days if heavy infection

\* NOT ON EDL





#### **6. CENTRAL NERVOUS SYSTEM INFECTIONS**

#### **6.1 ACUTE BACTERIAL MENINGITIS**

In **ADULTS**, *Streptococcus pneumoniae* is the mostly likely organism. *Haemophilus influenzae* and *Neisseria meningitidis* are less common. CSF microscopy and culture are vital in directing antibiotic therapy. Therefore a lumbar puncture and blood culture should be performed as soon as possible. Caution is required with lumbar puncture if the patient is in coma, has signs of increased intracranial pressure or has focal neurological signs. A CT scan of the head is preferred before lumbar puncture in such cases if facilities are available.

Bacterial meningitis is a medical emergency and antibiotic therapy should not be delayed if there is difficulty in obtaining a CSF sample. In such cases empirical therapy should be started immediately.

In rural areas or where there is a delay in transferring patient to a major hospital, if meningitis is suspected antibiotics should be started immediately, either with;

- Penicillin G 4 mega unit IV/ IM, 6 hourly OR
- Ceftriaxone 2g IV as a single dose (if available)

In penicillin hypersensitive patients use;

Chloramphenicol 1g IV 6 hourly.

Penicillin, chloramphenicol and ceftriaxone have proven effective in the treatment of meningitis. Chloramphenicol in oral doses achieves good CSF penetration.

Dexamethasone has been found to be useful in children. Recent literature suggests that it has a role in management of bacterial meningitis in adults and is to be given just before the first antibiotic dose.

- Dexamethasone 10mg IV just before the  $1^{\rm st}$  dose of antibiotic followed by 10mg 6 hourly x 4 days

## 6.1.1 Empirical

- Penicilllin G 1.8g (3 mega units) IV 4hourly for 10 days PLUS
- Chloramphenicol 750mg 1g IV 6 hourly for 10 days.

In patients hypersensitive to Penicillin use

- Chloramphenicol alone OR
- Ceftriaxone 4g IV daily in 1 or 2 divided doses

Change to appropriate regimen once the organism and susceptibility result is available. If no organism is identified, continue empirical therapy for a total of 10 days.







## 6.2.2 Specific Therapy

- (a) Pneumococcal/ Neisseria meningitides meningitis
   Penicillin G 1.8g (3 mega units) IV 4 hourly.

In penicillin hypersensitive patients

Ceftriaxone 4g IV daily in 2 divided doses

Pneumococcal meningitis is to be treated for 10 – 14 days. Some very ill patients may require treatment for 21 days. Meningitis due Neisseria meningitidis usually requires only 7 days treatment.

NB: At the end of penicillin therapy, rifampicin 10mg/kg/dose (up to 600mg) orally 12 hourly for 4 doses should be given to eradicate nasopharyngeal carriage in cases of meningococcal meningitis. This treatment should also be given to all close contacts.

(b) Gram negative bacterial meningitis and cryptococcal meningitis, consultation is advisable.

Generally for gram-negative meningitis other than H.influenzae, a combination of ceftriaxone and gentamicin is useful and the treatment is for 21 days



### 6.2 BRAIN ABSCESS

Most brain abscesses are polymicrobial. Usual organisms include anaerobic streptococci and bacteriodes.

Use:

- Penicillin G 2.4g (4 mega units) IV 4 hourly **PLUS** Chloramphenicol 1g IV 6 hourly **PLUS** Metronidazole 500mg IV 8 hourly/ PR 12 hourly

Ceftriaxone 2g IV 12 hourly can be substituted for chloramphenicol if indicated.

For post neuro-surgical brain abscess, use

- Vancomycin 1g IV 12 hourly **PLUS**Chloramphencol 1g IV 6 hourly **OR**Ceftriaxone 2g IV 12 hourly

Add cloxacillin or metronidazole if clinically or bacteriologically indicated.

The duration of treatment depends in clinical response and may be as long as 6-8 weeks. Surgical drainage may be indicated especially if clinically response is poor.





## 6.3 EPIDURAL ABSCESS

This is generally due to Staphylococcus aureus. Urgent surgery is essential.

Treatment should be based on gram stain and culture results of operative material.

For initial therapy use;

Cloxacillin 2g IV 6 hourly PLUS

Gentamicin 240mg IV once daily (maintenance dose to be adjusted depending on renal function)

This condition is often associated with osteomyelitis or disc infection, therefore subsequent management as for treatment of osteomyelitis.

## 6.4 HERPES SIMPLEX ENCEPHALITIS (suspected or proven)

Acyclovir 10mg/kg IV 8 hourly for at least 14 days, each dose to be infused not less than 1 hour (adjust dose according to the renal function).

## 6.5 TOXOPLASMA ENCEPHALAITIS

In AIDS cerebral infection with *Toxoplasma gondii* is common.



See under Opportunistic infections in HIV





## 7. URINARY TRACT INFECTIONS

## 7.1 UNCOMPLICATED LOWER URINARY TRACT INFECTIONS

Culture and sensitivity test are recommended.

Resistance to amoxycillin is as high as about 80% in Fiji. Resistance to trimethoprim is also rising. .

**Empirical Therapy** 

- Trimethoprim 300mg orally daily OR
- Nitrofurantoin 50 100mg orally 6 hourly

In non – pregnant women treatment is for 3-5 days.

For UTI in pregnancy, see section under Female Genital Tract and Obstetric Infections

In men, treatment is for 10-14 days. Note urinary tract infection in younger males is uncommon. All younger males with UTI should be investigated to exclude an underlying abnormality.

Treatment failures are usually due to a resistant organism, an unsuspected underlying abnormality of the urinary tract or re-infection with a similar organism. If relapse occurs, pyelonephritis should be considered and treatment given for 10 to 14 days.



Attempt to define or exclude underlying anatomical or functional abnormality. Treat for 14 days with antibiotics chosen on basis of urine culture and sensitivities. If sufficiently ill to need empirical therapy, treat with:

- Ampicillin 2g IV 6 hourly PLUS
- Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function).

If the use of an aminoglycoside is undesirable. e.g. in the elderly, in the presence of significant renal failure or following a previous adverse reaction, use  $\frac{1}{2}$ 

Ceftriaxone 1g IV daily

If Pseudomonas is isolated use;

Ciprofloxacin 500mg orally 12 hourly









#### URINARY TRACT INFECTIONS

Change to oral therapy based on culture and sensitivity report once patient is clinically well and afebrile for a few days.

If hypersensitive to penicillin, gentamicin can be used alone.

Follow up of urine culture is desirable at the conclusion of therapy.

## 7.3 RECURRENT URINARY TRACT INFECTIONS (PROPHYLAXIS)

Recurrent infections occur either as relapse of a previously treated infection or because of re-infection. Treat the acute infection as above for a duration of 10-14days.

Prophylactic therapy as follows (depending on sensitivity results):

Nitrofurantoin 50 – 100mg orally at night **OR**Nalidixic\* acid 1g orally at night **OR**Trimethoprim 150mg orally at night.

Prophylactic antibiotic therapy to be continued for 3 -6 months, longer if required

#### 7.4 CATHETER - ASSOCIATED URINARY INFECTIONS



Treat only if symptomatic. If treatment is needed, antimicrobial should be selected on the basis of the most recent urine culture results.

Where possible, removing the catheter may be all that is required. If a catheter needs to be changed in the presence of proven urinary tract infection, the procedure should be preceded by a single dose of an appropriate antibitotic administered by the IM or IV route.

### 7.5 ASYMPTOMATIC BACTERIURIA

Decision for treatment will depend on individual circumstances. Treatment often indicated in:

- Young children
- Pregnant women
- Men less than 60 years (should be investigated for chronic prostatitis)
- Patients with abnormal urinary tracts
- Patients undergoing genitourinary instrumentation

Choose antibiotic based on sensitivity results









## 7.6 CHRONIC BACTERIAL PROSTATITIS

Should be guided by culture and sensitivity tests.

- Co-trimoxazole (Trimethoprim 80mg & Sulphamethoxazole 400mg) 2 tablets orally 12 hourly OR
- Trimethoprim 300mg orally daily for 12 weeks.

If culture is negative consider:

- Doxycycline 100mg orally 12 hourly for 3 weeks followed by 100mg daily for 3 weeks **OR**
- Erythromycin 500mg orally 6 hourly for 6 weeks.

In resistant cases:

Ciprofloxacin 500mg orally 12 hourly for 4 weeks may be necessary.

Treatment of this condition is difficult and over treatment with antibiotics should be avoided.

#### 7.7 ACUTE EPIDIDYMO-ORCHITIS

- Mild to moderate

  (a) Sexual transmission:
  - Doxycycline 100mg orally 12 hourly for 10 14 days
  - (b) Non- Sexual transmission:
    - Co -Trimoxazole (trimethoprim 80mg and Sulphamethoxazole 400mg) 2 tablets orally 12 hourly X 14 days

- - Gentamicin 4 6mg/ kg/ IV once daily (adjusted according to renal function.

7.8 CANDIDURIA The presence of Candida in urine is common, particularly in association with indwelling catheters, and does not necessarily indicate renal tract infection. Antifungal therapy is not usually indicated and should not be initiated without consultation





#### 8. SEPTICAEMIA AND GENERALISED INFECTIONS

Patients with clinical features of septicemia require urgent empirical therapy. Microbiological specimens, especially blood cultures, should be obtained before the therapy is commenced. At least 2 blood culture specimens from two different sites should be obtained. Once the causative organisms have been identified and the sensitivities known, the antibiotic therapy may need modification. Duration of treatment varies from 10 - 21 days depending on severity, source of infection and organism involved.

## 8.1 EMPIRICAL THERAPY

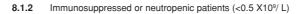
Source of Infection Unknown

- 8.1.1 Staphylococcus aureus is the most common cause of septicaemia in Fiji.
  - Cloxacillin 2g IV 4 hourly **PLUS**
  - Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function)

For patients hypersensitive to penicillin replace Cloxacillin with

Cephalothin 2g IV 6 hourly

**Note**: A small percentage of patients with penicillin hypersensitivity will also be hypersensitive to cephalothin (cephalosporins)



The following regimen aimed primarily at enterobacteriaceae and *Pseudomonas aeruginosa:* 

- Piperacillin 3g IV 4 hourly PLUS
- Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function).

See also section 2.7 below - Pseudomonas infections

If Staphylococcus aureus is suspected then ADD:

Cloxacillin 2g IV 4 hourly.



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#### **EMPIRICAL THERAPY**

Source of Infection is Predictable

8.2.1 Source Of Infection: Urinary Tract
Septicaemia usually due to *E.coli* or other enterobacteriaceae

• Ampicillin 2g IV 6 hourly PLUS

Contamicin 24000 IV appendably (maintenance does adjust

- Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function)

In patients hypersensitive to Penicillin, Gentamicin can be used alone. If aminoglycosides are contraindicated, substitute with

Ceftriaxone 2g IV daily

**8.2.2 Source Of Infection: Biliary or Gastro-intestinal Tract**Likely pathogens include Enterobacteriaceae, anaerobes including Bacteroides fragillis and less often, enterococcus.

- Ampicillin 2g IV 6 hourly PLUS
- Gentamicin 240mg once daily (maintenance dose adjusted for renal function) PLUS
- Metronidazole 500mg IV 12 hourly (infuse over 20 minutes) or suppository 500mg 12 hourly.

If hypersensitive to penicillin, consider:
• Vancomycin 1g IV 12 hourly (to be infused over at least one hour)

If gentamicin is contra-indicated substitute with:

Ceftriaxone 1-2g IV daily.

#### 8.2.3 Source Of Infection: Female Genital Tract

Usual pathogens include anaerobic bacteria, enterococcus, streptococcus and occasionally staphylococcus

- Ampicillin 2g IV 6 hourly **PLUS** Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function) PLUS
- Metronidazole 500mg IV 12 hourly (infused over 20minutes) or suppository 500mg 12 hourly.

In Penicillin hypersensitive patients, substitute Ampicillin with

Cephalothin 2g IV 6hourly

Note: A small percentage of patients with penicillin hypersensitivity will also be hypersensitive to cephalothin (cephalosporins)





#### 8.2.4 Source Of Infection: Skin

#### (a) Carbuncle

- Staphylococcus aureus is the most likely pathogen.
  Cloxacillin 2g IV 4 hourly PLUS
  Gentamicin 240mg IV once daily (maintenance adjusted for renal function)

#### (b) Cellulitis

Streptococcus pyogenes and Staphylococcus aureus are likely pathogens

Cloxacillin 2g IV 4 hourly

If inadequate response, consider adding

Penicillin G 3 mega units 6 hourly

If hypersensitivity to penicillin:

- Chloramphenicol 1g IV 6 hourly, OR
- Cephalothin 2g IV 4 6 hourly

Note: A small percentage of patients with penicillin hypersensitivity will also be hypersensitive to cephalothin (cephalosporins)

(c) Decubitus and ischaemic ulcers, diabetic foot infection

Pathogens may include Staphylococcus aureus, enterobacteriaceae, \_haemolytic streptococcus and anaerobes Surgical debridement of necrotic tissue is important

- Cloxacillin 2g IV 4 6 hourly **PLUS** Gentamicin 240g IV once daily (maintenance dose adjusted for renal function) PLUS
- Metronidazole 500mg IV 12 hourly (infused over 20 minutes) **or** suppository 500mg 12 hourly

## 8.2.5 Source Of Infection: Intravenous Cannulae including Central Venous Line

Infected cannula should be removed and along with any pus, submitted for culture Staphylococcus aureus, coagulase negative staphylococci, aerobic gram-negative bacilli and Candida sp are likely pathogens

- Cloxacillin 2g IV 4 hourly PLUS
- Gentamicin 240mg IV once daily (maintenance dose adjusted for renal function)



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#### SEPTICAEMIA AND GENERALISED INFECTIONS

Patients hypersensitive to penicillin:

Vancomycin 1g IV 12 hourly (each dose infused over 1 hour) can be substituted for cloxacillin

These regimens do not cover candida. Contact microbiologist if infection with candida is suspected Once sensitivity is available, adjust antibiotic accordingly

#### 8.2.6 Source Of Infection: Lung

Organisms include Streptococcus pneumoniae, gram-negative bacteria, Staphylococci, Haemophilus influenzae

- Cloxacillin 2g IV 4 hourly **PLUS** Gentamicin 240 mg IV once daily (maintenance dose adjusted for renal function)

If the response is inadequate ADD

Penicillin G 3mega units IV 6 hourly

Modify treatment once organisms and its sensitivity are known.

If aminoglycosides are contraindicated consider **ADDING** 

Ceftriaxone 2g IV daily

In patients hypersensitive to penicillin:

- Erythromycin 500mg 1g IV (infused over one hour) 6 hourly,
- Gentamicin 240 mg IV once daily (maintenance dose adjusted for renal function)

Streptococcus anginosus is not an uncommon cause. If suspected, add

Penicillin G 3 mega units 4 hourly. Prolonged treatment is indicated, usually for about 21 days

#### 8.2.7 Infections Due to Specific Organisms

In culture positive cases with known sensitivity, when patient is not seriously ill, monotherapy may be adequate, e.g. Streptococcal and Staphylococcal

#### Pseudomonas aeruginosa

- Piperacillin 3g IV 4 hourly **PLUS**Gentamicin 240 mg IV once daily (maintenance dose adjusted for renal function)









If hypersensitive to penicillin, or in cases of inadequate response use:
 Ciprofloxacin 500mg IV 12 hourly as a single drug

Duration of therapy for  $\it Pseudomonas~aeruginosa$  infections varies from 2 -6 weeks, depending on the primary site of infection

Neisseria meningitidis
Penicillin G 1.8g (3 mega units) IV 4 hourly

Alternative;
Ceftriaxone 2g IV daily as a single dose daily

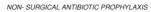
- Enterococcus: difficult to treat, consider:
   Ampicillin 2g IV 6 hourly PLUS
   Gentamicin 240 mg IV once daily (maintenance dose adjusted for renal function)

If patient fails to respond, contact microbiologist.





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# 9. NON- SURGICAL ANTIBIOTIC PROPHYLAXIS

#### 9.1 INFANT IMMUNISATION SCHEDULE

- BCG Bacillus Calmette -Guerin vaccine
  OPV Oral Polio vaccine
  HBV Hepatitis B vaccine
  DPT Diphtheria Pertussis Tetanus vaccine
  HIB Haemophilus influenzae type B vaccine
  TT Tetanus Toxoid

#### INFANTS

Age	Vaccine	Dose and Mode of Administration
Birth	BCG 1 OPV 1 HBV 1	0.1mL – intradermal 2 drops - oral 10micrograms – intramuscular
2 Months	OPV 2 HBV 2 DPT 1 HIB 1	2 drops – oral 10 micrograms – intramuscular 10 micrograms – intramuscular 10 micrograms – intramuscular
3 months	OPV 3 DPT 2 HIB 2	2 drops - oral 10 micrograms – intramuscular 10micrograms – intramuscular
4 months	OPV 4 DPT 3 HIB 3	2 drops – oral 10 micrograms – intramuscular 10 micrograms - intramuscular
5 months	HBV 3	10 micrograms – intramascular
9 months	Measles/ Rubella	0.5mL - intramuscular







#### SCHOOL CHILDREN

Age	Vaccine	Dose and Mode of Administration
At School Entry	BCG 2 TT - booster	0.1mL – intradermal 0.5mL – intramuscular
At School Leaving	TT booster Rubella (for girls)	0.5mL – intramuscular 0.5mL – intramuscular

# NONIMMUNISED CHILDREN

Age	Vaccine	Dose and Mode of Administration
1. As soon as found	TT 1 OPV 1 BCG	0.5mL – intramuscular 2 drops – oral 0.5mL – intradermal
2. Four weeks after	TT 2	0.5mL – intramuscular
TT 1 / OPV 1	OPV2	2 drops – orally
3. Four weeks after	TT 3	0.5mL – intramuscular
TT 2/ OPV 2	OPV 3	2 drops - oral

Three doses of Hepatitis B vaccine can be given but it is recommended that the immunological status of the child is first determined.

# ANTENATAL WOMEN

# A. First Pregnancy

Dose	When to give	Expected duration of protection
TT1	At first contact or as early as possible in pregnancy	None
TT 2	At least 4 weeks after TT 1	1 – 3 years
TT 3	At least 6 months after TT 2	5 years
TT 4	At least one year after TT 3 or during subsequent pregnancy	10 years
TT 5	At least one year after TT 4 or during subsequent pregnancy	All childbearing years

It is envisaged that the first three doses can be administered to a woman in her first pregnancy. The third dose, if need be can given during MCH clinic for the baby.

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## 9.2 HEPATITIS B

#### Pre-exposure

Indications:

- All newborns
   Those at risk of contracting infection e.g. health care workers
   Sexual partners of HbsAg positive individuals

In these cases, a full course of Hepatitis B vaccine should be administered. The vaccine is given at 0, 1 and 6 months (except for infants – refer to immunisation schedule above). For dosage, refer to manufacturer's instructions.

## Post - Exposure

Following needle stick injury or mucosal/ broken skin contamination by body fluids, the following applies:

Test the exposed person and the donor of contaminating blood (patient) for HbsAg and Anti-HBsAg

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EXPOSED PERSON	STATUS	ACTION
Full course of Hepatitis B vaccine with seroconversion within past 5 years	Protected	No prophylaxis needed
HBsAg positive	Already infected	No prophylaxis needed
Anti – HBsAg positive	Protected	No prophylaxis needed
HBsAg and Anti-HbsAg negative	Susceptible	Vaccinate. Add immunoglobulin if: Patient is HbsAg positive Status of the patient is unknown





#### B. Neonates of mothers who are HbsAg positive

All such neonates should receive immunoglobulin in addition to vaccination. The immunoglobulin should be given within 48 hours of birth.

In cases where immunoglobulin is indicated:

A single dose should be given intramuscularly, within 7 days but preferably within 48 hours of exposure. The adult dose is 400 international units.

Dose for neonates: Check manufacturer's instructions.

A full course of Hepatitis B vaccine should be given, starting at the time as the immunoglobulin. For dosage, refer to the manufacturer's instructions

#### 9.3 HIV - POST NEEDLESTICK INJURY PROPHYLAXIS

If the source of the inoculated blood or body fluids is HIV antibody positive, post exposure prophylaxis, with antiretrovirals, should be commenced as soon as possible and the recipient followed up for at least 6 months.

If the source patient is also at a high risk of HIV then prophylaxis antiretroviral is warranted

Antiretroviral therapy should be commenced as soon as possible following the injury or preferably within hours.

Currently only TWO antiretroviral drugs are available on Fiji EDL

Zidovudine 300mg orally X 4 weeks **PLUS** 

- Lamuvudine 150mg orally 12 hourly X 4 weeks

#### 9.4 MENINGOCOCCAL AND H. INFLUENZAE MENINGITIS

Despite prophylaxis disease can still occur.

Indications:

- 1. Close contacts of the index case.
- 2. In epidemics: classroom and other institutional contacts.
- 3. Contacts who are young children (less than 4 years old).
- 4. Index case before discharge from hospital to eradicate nasopharyngeal

Note: for Haemophilus influenzae meningitis, the immunisation status of the contacts should be taken into consideration.

#### a) H. influenzae contacts

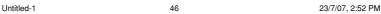
Adult:

Rifampicin 600mg orally daily X 4 days

Children (over 4 weeks old):

Rifampicin 10mg/kg/ dose (max 600mg / dose) orally for 4 days 46











## b) Meningococcal Contacts

Adult:

- Rifampicin 600mg orally 12 hourly for 2 days OR
- Ceftriaxone 250mg IM as a single dose OR
- Ciprofloxacin 500mg orally as a single dose.

Children (over 4 weeks old):

- Rifampicin 10mg/ kg /dose (max 600mg /dose) orally 12 hourly for 2 days OR
- Ceftriaxone 125mg IM as a single dose.

#### 9.5 PREVENTION OF RECURRENCE OF RHEUMATIC FEVER

Continuous antimicrobial prophylaxis against *Streptococcus pyogenes* infection is recommended for patients with a well documented history of rheumatic fever. Intramuscular administration is preferred, especially in remote areas, as it is more effective and usually leads to better adherence.

#### No cardiac involvement:

- Benzathine Penicillin 1.2 mega units every 3 4 weeks for 5 years or up to 18 years OR
- Penicillin V 250 orally 12 hourly, duration same as above.

Patients hypersensitive to Penicillin:

Erythromycin 250mg orally 12 hourly.

#### With cardiac involvement:

Drug prophylaxis same as above but until at least 40 years of age and preferably for life, in cases of severe carditis or moderate to severe residual valve disease (including those who have had valve surgery)

If there is evidence of initial cardiac involvement with subsequent resolution, then antibiotic prophylaxis is recommended for up to the age of 25 years.

Note: Compliance with oral prophylaxis is usually less optimal hence injection is preferred.

#### 9.6 PREVENTION OF INFECTIVE ENDOCARDITIS

Prophylaxis for endocarditis is an accepted but an unproven practice. The recommendations listed below are based upon current international practice. The need for prophylaxis depends upon the cardiac condition of the patient and the procedure being performed. Prophylaxis should be considered in all patients with structural heart disease undergoing surgical procedures.









#### 9.6.1 For all patients At Low Risk (i.e. without prosthetic valves or a previous attack of infective endocarditis) undergoing dental procedures, oral surgery or upper respiratory tract surgery.

#### Under Local Anaesthetic

If the patient is not receiving long-term penicillin:

• Amoxycillin 3g orally, 1 hour before the procedure, then 1.5g 6 hours after the initial dose.

If the patient is hypersensitive to penicillin or receiving long term penicillin:

Erythromycin 1g orally, 2 hours before procedure, then 500mg 6 hours after the procedure.

<u>Under General Anaesthetic</u>
• Ampicillin 2g IV just before the procedure commences or IM 30 minutes before the procedure commences **followed by** Ampicillin 500mg I/IM **or** Amoxycillin 500mg orally 6 hours later.

For patients hypersensitive to penicillin or receiving long term penicillin:

- Vancomycin 1g IV (slowly over 1 hour), the infusion ending just before the procedure commences.
- 9.6.2 Low Risk patients having gastrointestinal, genitourinary procedures or other major surgery

Prophylaxis same as for section (6.3) below

- 9.6.3 For Patients at High Risk (i.e. patients with prosthetic valves or those who have had a previous attack of infective endocarditis) undergoing any type of dental or surgical procedures
  - Ampicillin 2g IV just before the procedure commences or IM 30 minutes before the procedure commences, followed by Ampicillin 500mg IV/IM or Amoxycillin 500mg orally 6 hours later PLUS
  - Gentamicin 1.5mg/kg IV just before the procedure commences **OR** IM 30 minutes before the procedure commences.

Patients hypersensitive to Penicillin

- Vancomycin 1g IV (slowly over 1 hour), the infusion ending just before the procedure commences PLUS
- Gentamicin 1.5mg/kg IV just before the procedure commences







## 9.7 POST SPLENECTOMY PROPHYLAXIS

Vaccine preparations available in Fiji include pneumococcal and HiB vaccines. Pneumoccal vaccine is recommended every 5 years, whereas HiB vaccine is recommended once only if the individual has not already been immunised. For elective splenectomy, vaccination should be given two weeks before surgery.

Antibiotic prophylaxis should be assessed for each patient individually. Those at the highest risk include children, during the 1st two years following splenectomy and those associated with severe underlying immunosuppression.

Penicillin V 250mg orally 12 hourly

For patients hypersensitive to Penicillin
• Erythromycin 250mg orally daily.

Duration of prophylaxis remains unclear but it is recommended that patients should be covered for at least 2 years. However, the risk of bacterial sepsis remains life long.







#### 10. MALARIA

#### 10.1 PROPHYLAXIS FOR VISITORS TO MALARIOUS AREAS

- Personal protection against being bitten by mosquitoes is very important even when one is on drug prophylaxis. Keep body well covered, use mosquito nets and use repellants.
- Chemoprophylaxis. The choice of drug resistance, side effects of drug, patient criteria e.g. age, pregnancy, renal disease etc.

Prophylaxis is relative not absolute. Breakthroughs can occur with any of the recommended drugs. The following is based on WHO recommendations for 1997.

#### Duration of prophylaxis:

Prophylactic drugs should be started at least one week before travel to endemic area. This allows time for the drug to reach adequate tissue levels and also helps to see how well the drug is tolerated. The drugs should be continued for the duration of stay and for a minimum of 4 weeks, preferably 6 weeks, after leaving the malarious area.

#### Return from malaria infested area:

Any illness within a year of return, especially within three months, might be malaria. Travellers should be warned to consult a doctor and say that they have been exposed to malaria if they fall ill within this period.

The following is only recommended and is not an inflexible directive. Some of the drugs mentioned are not available on the Fiji EDL.

Prophylaxis in areas of Chloroquine resistant P.falciparum. e.g. Papua New Guinea, Solomon Islands, Vanuatu.

## <u>Adults</u>

- Mefloquine 250mg orally once weekly **OR**
- Chloroquine 300mg (base) orally once weekly and proguanil 200mg orally once daily **OR**
- Doxycycline 100mg orally once daily

Caution: Mefloquine may be associated with neuropsychiatric side effects. It may have antagonistic effects on anticonvulsants and is not safe in pregnancy. It should be avoided in the first trimester of pregnancy.

It is contraindicated in patients with neuropsychiatric disorder, epilepsy or cardiac conduction defects.

## Doxycycline is also to be avoided in pregnancy





# Children

Babies and children MUST be given malaria prophylaxis even if breastfed. Tablets may be broken and are easily crushed (mix with food e.g. jam). Chloroquine is available as syrup (50mg in 5 ml) and this formulation is recommended for babies and young children.

Either mefloquine alone or chloroquine + proguanil are recommended. Dosage as shown in tables below:

**MEFLOQUINE\*** (Not recommended for children under 15 kg or 2 years of age.)

	Weight	Age	Dose (weekly)
Mefloquine* 250 mg tablet	15 – 19 kg 20 – 30 kg 31 – 45 kg >45 kg	2-5 years 6-8 years 9-11 years > 11 years	<ul><li>tablet</li><li>tablet</li><li>tablet</li><li>1 tablet</li></ul>

# CHLOROQUINE Plus PROGUANIL



Both these drugs are well tolerated by children and appropriate for those under 2 years. The dose /kg given is the minimum recommended to give adequate protection against malaria.

- Chloroquine 5 mg/kg (up to 300mg) once weekly plus Proguanil 3mg/kg (up to 200 mg) once daily

Chloroquine and Proguanil* Adult dosage: Chloroquine 300mg weekly Proguanil 200mg daily	Age	Weight	Fraction of Adult Dose*
	Under 6 weeks		1/8
	6 weeks - 1 year	4 – 9 kg	1/4
	1 - 5 years	10 – 19 kg	1/2
	6 – 12 years	20 – 39 kg	3/4
	Over 12 years	> 40 kg	Adult dose







# 10.2 TREATMENT OF INFECTED PERSONS

If the infective species is not known or if the infection is mixed the treatment should be as for P.falciparum.

#### P.FALCIPARUM MALARIA

As *P. falciparum* is often resistant to Chloroquine, it should not be used for treatment.

#### <u>Adults</u>

#### Oral therapy

If no vomiting or impairment of consciousness and patient is able to swallow:

- Quinine Sulphate 600mg orally 8 hourly for 7 days, **PLUS EITHER** Fansidar\* (pyrimethamine 25mg/ sulphadoxine 500mg) 3 tablets as single dose on day three OR
- Tetracycline 250mg 6 hourly for 7 days.

#### Alternative:

Mefloquine 20mg/kg as a single dose (maximum 1.5g) or preferably should be given as 2 divided doses 6 -8 hours apart



# Intravenous Therapy

If patient is seriously ill or cannot tolerate oral medication:

• Quinine dihydrochloride

Loading dose: 20mg /kg quinine dihydrochloride (maximum 1.4g of quinine salt) infused over 4 hours in 500ml of normal saline. Following this it is advisable to give 500ml 5 % dextrose before the next dose of quinine.

Maintenance dose to start 4 – 8 hours after loading dose is completed: 10mg/kg (maximum 700mg) in 500ml normal saline (infused over 4 hours) 8 hourly. Give 5% dextrose in between doses if necessary.

This is continued until the patient can swallow tablets to complete the 7-day course.

A loading dose of 20mg/kg of quinine should not be given if the patient has received quinine, quinidine or mefloquine during the previous 24 hours. Start on the maintenance dose of 10mg/kg. Dosage may have to be reduced in the presence of hepatic and renal dysfunction.

Quinine can produce hypoglycemia and over hydration may produce pulmonary oedema.

Quinine is followed by EITHER:

• Fansidar\* ( Pyrimethamine 25mg/ sulphadoxine 500mg) 3 tablets as a single

\* NOT ON EDL

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dose on day 3  $\mathbf{OR}$ 

Tetracycline 250 mg 6 hourly for 7 days (Start as early as possible)

#### Children

Dosage should not exceed adult dosage.

# Oral Therapy

Quinine sulphate 10mg/kg orally every 8 hours for 7 days **PLUS**Fansidar\* (pyrimethamine 25mg/ sulphadoxine 500mg) as a single dose on day 3 as shown below:

AGE	DOSE
6 weeks – 1 year	- tablet
1 – 4 years	- tablet
5 – 6 years	1 tablet
7 – 9 years	1- tablets
10 – 14 years	2 tablets
> 14 years	3 tablets

Alternative (not in children <2 years or <15 kg):

 Mefloquine 20mg/kg of Mefloquine base as a dose or preferably as 2 divided doses 6 -8 hours apart.

#### Intravenous Therapy:

- Quinine dihydrochloride 10mg/kg every 8 hours in normal saline as described under adults (with volume reduced as appropriate), until patient can swallow, followed by quinine tablets (dosage as above) to complete a seven day course PLUS
- seven day course PLUS

   Fansidar\* as above on day 3

IV dextrose solution is required as discussed under adult.

Adjust volume as necessary.

## Pregnancy

Malaria is dangerous to the mother in the last trimester and to the foetus throughout the pregnancy. Adult doses of oral and intravenous quinine can safely be given to pregnant women.

# BENIGN MALARIA (VIVAX MALARIA)

Treatment should be aimed at eradicating blood and liver stages (radical cure). Chloroquine is used for the former and primaquine for the latter.

\* NOT ON EDL

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## <u>Adult</u>

- Chloroquine 600mg (of base) orally followed by 300mg (of base) in a single dose 6-8 hours later then 300mg daily for 2 days **FOLLOWED BY**Primaquine 15mg orally daily for 21 days after the chloroquine

# Children

- Chloroquine initial dose of 10mg/kg (max 600mg) of base orally followed by a single dose of 5mg /kg (max 300mg) 6 8 hours later and then a single dose of 5mg/kg (max 300mg) daily for 2 days **FOLLOWED BY**Primaquine 250micograms/ kg (max 15 mg) daily for 21 days

#### **Pregnancy**

Chloroquine (as for adults). Primaquine postponed until after delivery since it is harmful to the foetus

Chloroquine 600 mg (of base) each week to be continued during pregnancy.







## 11. FEMALE GENITAL TRACT AND OBSTETRIC INFECTIONS

#### 11.1 PELVIC INFLAMMATORY DISEASE

The common organisms are gonococci, chlamydia and a mixture of aerobic and anaerobic organisms.

- (a) Mild: for outpatient management
  - Doxycycline 100mg orally 12 hourly X 10 days **PLUS** Metronidazole 400mg orally 8 hourly X 7 days **PLUS** Amoxycilllin 500mg orally 8 hourly X 7 days
- (b) Severe enough for admission

  - Ampicillin 1g IV 6 hourly **PLUS** Gentamicin 240mg IV daily (maintenance dose adjusted for renal function) PLUS
  - Metronidazole 400mg orally 8 hourly or 500mg PR 8 hourly

Change to oral when infection is resolving:

- Doxycycline 100mg orally 12 hourly X 10 days **PLUS** Metronidazole 400mg orally 8 hourly X 7 days.

Add other antibiotics if indicated. Drainage may be required.

## 11.2 BARTHOLIN'S ABSCESS

Surgical drainage is required.

Ampicillin 1g IV as a single dose, followed by oral Amoxycillin if indicated.

## 11.3 SEXUAL ASSAULT

For empirical therapy against gonorrhoea (and may cover Syphilis). This regimen may not prevent other STIs. It is important that the patient is followed up with clinical examination and serological tests. Antibiotic therapy is indicated where f ollow up maybe difficult

### Adult

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Augmentin (Amoxycillin 500mg and Clavulanic acid 125 mg) 1 tablet











## orally PLUS

Amoxycillin 2.5g as single dose orally

If Augmentin is not available, then use

- Amoxycillin 3g as a single dose **PLUS** Probenecid 500mg as a single dose

#### Children

• Ceftriaxone 125mg IM as a single dose.

#### 11.4 INFECTIONS IN OBSTETRIC PRACTICE

# 11.4.1 LOWER URINARY TRACT

If asymptomatic delay treatment until culture and sensitivity result is known. If empirical treatment is indicated:

Nitrofurantoin 50mg orally 6 hourly OR

Cephalexin\* 250mg orally 6 hourly

Change to appropriate antibiotic based upon culture and sensitivity result.

Treatment is usually for 10 - 14 days.

### 11.4.2 PYELONEPHRITIS

Urine and blood culture are indicated.

- Ampicillin 1 g IV 6 hourly **PLUS**Gentamicin 240mg IV daily (maintenance dose adjusted for renal function) OR
- Cephalothin 1 g IV 6 hourly as a single agent

Gentamicin may have adverse effects on the foetus. It should only be used where the benefits to the mother outweigh the risks to the foetus. It should be used for the minimum necessary period.

Once patient has become afebrile for 72 hours, switch to oral therapy based on culture sensitivity results. Total duration of therapy is 10- 14 days.

#### 11.4.3 BREAST ABSCESS

Cloxacillin 1g IV 6 hourly X 48 hours, followed by oral treatment as indicated.

If hypersensitive to penicillin consider:

- Cephalothin 1 g IV 6 hourly OR
- Erythromycin 500mg orally 6 hourly.





Note: A small percentage of patients hypersensitive to penicillin may also be hypersensitive to Cephalothin (Cephalosporin)

# 11.4.4 PREMATURE RUPTURE OF MEMBRANES

#### (a) Pre-term

- Ampicillin 1g IV 6 hourly **PLUS**Gentamicin 240mg IV daily (maintenance dose adjusted for renal failure) **Followed by**Erythromycin 500mg orally 6 hourly **PLUS**Metronidazole 400mg orally 8 hourly

Duration of therapy is usually for one week.

# (b) At Term

If the rupture has been for more than 18 hours, antibiotic has to be started

- Ampicillin 1g IV 6 hourly Gentancian 240mg IV once daily (maintenance dose adjusted for renal function)

The patient is usually induced, and antibiotic coverage includes pre-labour, during labour and post labour for a total of 72 hours.





#### 12. SEXUALLY TRANSMITTED INFECTIONS

#### 12.1 GONOCOCCAL INFECTIONS

Diagnosis should be confirmed by culture where possible and antibiotic sensitivity tests performed

All patients should also have serological tests for syphilis and HIV after confidential counselling.

Follow up cultures should be obtained 5 days following completion of treatment of all gonococcal infections.

### 12.1.1 <u>Uncomplicated urethral, endocervical, rectal or pharyngeal infections</u>

All these patients are treated for gonococcal and chlamydial infections.

- Augmentin (Amoxycillin 500 mg with Clavulanic acid 125 mg)
  - 1 tablet single dose PLUS
- Amoxycillin 2.5 g orally as a single dose PLUS
- Probenecid 1g as a single dose

#### **PLUS EITHER**

- Doxycycline 100 mg orally 12 hourly for 7 days OR
- Tetracycline 500 mg orally 6 hourly for 7 days.

Procaine penicillin 4 mega units IM as a single dose may be used in place of Augmentin and Amoxycillin.

#### Alternatives:

- Co-trimoxazole (sulphamethoxazole 400 mg with trimethoprim 80 mg) 5 tablets orally 12 hourly for 5 days OR
- Ceftriaxone 250 mg IM as a single dose

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# EITHER OF THE ABOVE WITH

- Doxycycline 100 mg 12 hourly for 7 days **OR**
- Tetracycline 500 mg orally 6 hourly for 7 days.

# 12.1.2 <u>Pregnant women</u>

# As above, but

 Erythromycin 500mg orally 6 hourly for 7 days should be substituted for doxycycline or tetracycline.

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## 12.1.3 Sex partners

Persons exposed to gonorrhoea within the preceding 30 days should be examined, appropriate samples cultured and treated presumptively as above.

#### 12.1.4 <u>Disseminated Gonococcal Infection</u>

Patients should be hospitalised.

- Ceftriaxone 1g IM/ IV once daily until 2 days after symptoms resolve **followed by**
- Augmentin (amoxycillin 500 mg with clavulanic acid 125 mg) 1 tablet orally 8 hourly to complete 7 days

  OR if infection is proven to be penicillin sensitive:

  Ampicillin 1g IV 6 hourly followed by oral Amoxycillin 500 mg
- orally 8 hourly to complete 7 days.

# 12.1.5 <u>Infants with Gonococcal Ophthalmia</u>

Exclude disseminated gonococcal infection by careful physical examination; aspiration of joints, blood and CSF cultures, where

All patients must have careful ophthalmological examination and systemic therapy is essential.

- (a) Penicillin sensitive N. gonorrhoeae
- Penicillin G 50,000 units/kg/dose IV 12 hourly (6 hourly for infants more than 1 week old) for 7 days.

If meningitis or disseminated infection is present, the duration should be increased to 14 days.

- (b) Penicillinase producing N. gonorrhoeae
- Ceftriaxone 25 50 mg/kg/day IV or IM as a single daily dose for 7  $\,$

Irrigation of eyes with saline is a useful adjunctive therapy. Topical antibiotics alone are insufficient and are unnecessary when appropriate systemic therapy is given.

Treat mother and sexual partner.



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#### 12.1.6 Gonococcal Ophthalmia in older children and adults

Gonococcal Ophthalmia in adults and children more than 20 kg with non-disseminated disease.

Ceftriaxone 1g IM as a single dose.

Irrigation of eyes with saline is a useful adjunctive therapy.

Topical antibiotics alone are insufficient, and are unnecessary when appropriate systemic therapy is given.

## 12.1.7 <u>Infection in children following child abuse</u>

- Ceftriaxone 125mg IM as a single dose (or 250mg if child >45kg)
   PLUS
- Doxycycline 100mg orally 12 hourly for 7 days or
- for children <9 years old: use Erythromycin

#### 12.2 CHLAMYDIAL INFECTIONS

The decision to treat chlamydial infection alone is based on proven absence of gonorrhoea and presence of chlamydia on testing. Antigen testing is done on endocervical swabs, male uretheral swabs or urine, and conjunctival swabs.

Special swabs are available in Suva only at present.

Doxycycline 100mg orally 12 hourly for 7 days

Where doxycycline is contraindicated (e.g. children, infants, pregnancy):

Erythromycin 500mg orally 6 hourly for 7 days.

Test of cure is not necessary since resistance of *C. trachomatis* to the above regimens has not been observed.

#### Sex Partners

Sex partners should be treated for *C. trachomatis* if their contact was within 30 days of onset of symptoms in index case.

#### 12.3 SYPHILIS

Irrespective of clinical presentation anyone suspected of having a STI should have a serological test for syphilis. VDRL test may be negative during primary syphilis therefore treatment should be given to persons presenting with clinical features of primary syphilis e.g. genital ulcers.

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#### SEXUALLY TRANSMITTED INFECTIONS

All patients with syphilis should be counselled regarding the risks of HIV and be encouraged to be tested for HIV.

#### 12.3.1 Early syphilis – Primary, secondary or early latent syphilis of less than 1 year's duration.

Benzathine penicillin 2.4 mega units IM as a single dose.

For penicillin hypersensitivity in non-pregnant patients: Doxycycline 100mg orally 12 hourly for 2 weeks **OR** Erythromycin 500mg orally 6 hourly for 2 weeks.

Pregnancy: see (3.5) below

Follow up:

Patient should be re-examined clinically and serologically at 3 months and 6 months.

- Patients should have a CSF examination and repeat treatment if:

  (i) VDRL titres have not declined four fold by 3 months in primary or secondary syphilis or by 6 months in early latent syphilis **OR**(ii) Signs or symptoms persist and re-infection has been ruled out.

### 12.3.2 HIV infected patient with syphilis

Initial treatment as in (3.1).

Follow up is as follows:

Repeat VDRL test at 1,2,3,6 and 12 months. Any patient with a four-fold increase in titre at any time should have a CSF examination and treated accordingly.

### <u>Late latent syphilis (more than 1 year's duration) Gummas and Cardiovascular syphilis</u> 12.3.3

Benzathine Penicillin 2.3 mega units IM as a single dose per week for 3 consecutive weeks (total dose 7.2 million units).

Penicillin hypersensitivity in non-pregnant patients:

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Doxycycline 100 mg orally 12 hourly for 4 weeks



Follow up:

Repeat VDRL at 6 and 12 months.

- Patient should be evaluated for neurosyphilis if:

  Neurological signs or symptoms are present
  Treatment failure (serologically or clinically)

#### 12.3.4 Neurosyphilis

- Penicillin G 1.2 2.4 g (2-4 mega units) IV every 4 hours for 10 -14 days followed by
- Benzathine Penicillin 2.4 mega units IM weekly for 3 doses.

For those hypersensitive to penicillin, contact Microbiologist or Physician.

Follow up:

If CSF pleocytosis is present initially, repeat CSF examination (VDRL and cell count) every 6 months until the cell count is normal. Re-treat if no decrease in cell count at 6 months or if it is not normal by 2 years.

# 12.3.5 Syphilis in pregnancy

Pregnant women should be screened early in pregnancy, at first booking to antenatal clinic.

Screening should be repeated in the third trimester and preferably again at delivery.

Patients should be treated with the penicillin regimen appropriate for the stage of syphilis except for early syphilis, which should be treated as follows:

- Benzathine penicillin 2.4 million units IM given as one dose per week for 3 weeks OR
- Procaine penicillin 1.2 mega units daily for 10-14 days (especially after 20 weeks gestation).

Following treatment, monthly VDRL testing should be done for duration of pregnancy.

A repeat course is indicated:

- (a) if the sexual partner was not treated simultaneously
- (b) if the titre is not falling within 6 weeks.





Those hypersensitive to penicillin:

Penicillin therapy after desensitisation is preferred.

For less serious allergy:
 Cephalothin 1 g IV 6 hourly (initially as inpatient), for 10 days.

**Note**: A small precentage of patients hypersensitive to penicillin are also hypersensitive to cephalothin (cephalosporins).

# 12.3.6 Infants of VDRL positive mothers

Infants should be treated if any one of the following is present:

#### In mothers:

- (1) Syphilis untreated or inadequately treated during pregnancy
  (2) Syphilis during pregnancy treated with non-penicillin regimen
  (3) Syphilis during pregnancy treated with an appropriate regimen but the expected decrease in VDRL antibody titres did not occur after therapy
- (4) Syphilis treated less than one month before delivery
- (4) Syphilis treated esset filan one month before delivery
   (5) Undocumented syphilis treatment
   (6) Syphilis treated before or during pregnancy but with insufficient serological follow up during pregnancy to assess the response to treatment and current infection status.

## In infants:

- (1) Any evidence of active disease (revealed by physical or X-ray findings)
- (2) VDRL titres more than in the mother
- (3) Baby is positive in FTA-Abs IgM test.

#### If there is no evidence (physical or X-ray) of congenital syphilis but mother or infant has one of the above-mentioned features:

- Procaine Penicillin 50,000 units/kg IM daily for 10-14 days OR
- Benzathine Penicillin 50,000 units/kg IM once a week for 3 weeks

Follow up: Repeat VDRL every 3 months till VDRL is negative: Repeat treatment if it remains positive.

## 12.3.7 Congenital syphilis

When baby has any of the following features it is considered to have congenital syphilis:

- (1) Clinical signs of syphilis
- (2) X-ray changes



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## 3) Neurological manifestations

Do CSF VDRL test, cell count, protein estimation in such babies.

#### If any of the following abnormalities exist indicating neurosyphilis i.e.:

- a) CSF VDRL is reactive
  b) Leucocyte count is >5/mm³
  c) Protein >400 mg/L.

#### Then treat as follows:

Penicillin G 50,000 units/kg IV 12 hourly for 7 days followed by 50,000 units/kg 8 hourly for a total of 14 days

# If CSF findings are negative (congenital syphilis without neurosyphilis) give the same treatment but for 10 days only.

#### Follow up:

If CSF was positive, repeat VDRL at 6 months. If CSF was negative, follow up as in (3.6).

If no response to treatment, re-treat or consider neurosyphilis.

# 12.3.8 Older infants and children

If congenital syphilis or neurological involvement:

Penicillin G 50,000 units/kg IV every 4 - 6 hours for 10-14 days.

Definite acquired syphilis and no neurological involvement:

Benzathine penicillin 50,000 units/kg IM (max 2.4 mega units/dose) as a single dose.

Evaluate for sexual abuse.

# 12.3.9 Sex partner

Trace and treat contacts wherever possible as appropriate for the stage of the disease.

#### HERPES SIMPLEX VIRUS INFECTION (HSV) 12.4

#### Primary Genital Herpes

Acyclovir\* 200mg orally 5 times daily for 10 days.

# Sex Partners:

Evaluate sex partner(s) with genital lesions and treat.





Patient and partner(s) should be counselled about the natural history of the disease with emphasis on potential for recurrences. Advise abstinence from sexual activity while lesions are present.

#### **TRICHOMONIASIS** 12.5

- Metronidazole 2 g orally as a single dose. Recurrent attacks:
- Metronidazole 400 mg orally 8 hourly for 7 days.

Trichomoniasis during pregnancy: For severe infections in the first trimester.

Econazole pessaries 150mg at night for 3 days.

Metronidazole is contraindicated in the first trimester of pregnancy and its safety in the last two trimesters of pregnancy is not proven.

For patients with severe symptoms and resistant to econazole the following may be considered after the first trimester:

Metronidazole 2g orally as a single dose.

Sex Partners:

Require evaluation and treatment.

### 12.6

<u>VULVOVAGINAL CANDIDIASIS</u> (This is not strictly a STI but is covered in this section for convenience)

Econazole nitrate (150 mg vaginal Pessary) 1 pessary at bed time for 3

Sex partners: Treatment not necessary unless candidial balanitis is present, when local therapy is sufficient.

## **BACTERIAL VAGINOSIS**

(non specific vaginitis, Gardnerella vaginosis)
(This is not a proven STI but is covered in this section for convenience.)

Metronidazole 400 mg orally 8 hourly for 7 days.

Alternative:

Amoxycillin 500 mg orally 8 hourly for 7 days (preferred if treatment required in pregnancy but less effective than metronidazole).

Sex Partner: Treatment of sex partner is not necessary.







#### 12.8 **GENITAL WARTS**

Genital warts are caused by certain types of Human papilloma virus (HPV). The goal of treatment is removal of exophytic warts and relieving signs and symptoms and not eradication of HPV.

3.1 External Genital/ Perianal Warts

Podophyllin\* (20-25%) in compound tincture of benzoin applied to the wart/s.

Note: Burning, pain and inflammation may occur in up to 50% of patients treated with podophyllin.

Total area of wart treated per session should be less than 10cm<sup>2</sup> Avoid surrounding skin as far as possible. Thoroughly wash off after 4 hours.

Repeat applications at weekly intervals.

Contraindicated in pregnancy and breastfeeding mothers.

# 12.8.2 Vaginal warts

Podophyllin\* (20-25%) in compound tincture of benzoin.

Treatment area should be dry before speculum is removed.

Treat less than 2cm<sup>2</sup> per session.
Repeat application at weekly intervals.

Contraindicated in pregnancy and breastfeeding mothers.

#### 12.8.3 **Urethral meatus**

Podophyllin regimen

#### 12.8.4 **Anal Warts**

Surgical removal.

#### 12.8.5 **Oral Warts**

Surgical removal or electrocautery.

#### 12.9 CHANCROID (Causative agent H. ducreyi)

Considered in the differential diagnosis of any patient with a painful genital ulcer. Nearly 50% of cases have painful inguinal adenopathy.

• Erythromycin base 500 mg orally 6 hourly for 7 days.

\* NOT ON EDL





#### Alternatives:

Co-trimoxazole (Sulplhamethoxazole 400 mg with trimethoprim 80 mg) 2 tablets orally 12 hourly for 7 days OR
Augmentin (amoxycillin 500 mg with clavulanic acid 125 mg) 1 tablet orally 8 hourly for 7 days.
Sex Partners:
Sex partners, within the 3 weeks preceding onset of symptoms in patients, whether symptomatic or not, should be examined and treated with the above recommended regimen.

Follow up: Symptomatic improvement is seen within 3 days and resolution of lesions within 7 days. Observe patients until ulcer is completely healed.

#### LYMPHOGRANULOMA VENEREUM (LGV) 12.10

Caused by LGV serovars of C. trachomatis. Inguinal adenopathy is the most common clinical manifestation.

Doxycycline 100 mg orally 12 hourly for 21 days.

# Alternative:

Erythromycin 500 mg orally 6 hourly for 21 days.

#### Sex partners:

Should be treated unless shown to be free from infection.





#### 13. SURGICAL ANTIBIOTIC PROPHYLAXIS

The primary objective of antibiotic prophylaxis is to prevent postoperative wound infection and to a lesser degree postoperative respiratory tract and urinary tract infections. Surgical procedures which do not traverse areas with normal flora, other than the prepared skin, do not routinely require prophylaxis.

Prophylaxis should be considered where:

- there is significant risk of infection e.g. colonic resection
- 2. infection although uncommon would have severe consequences e.g. that associated with prosthetic implants.

Sufficient concentration of drugs should be available in the tissues at the time of exposure. Therefore, give IM antibiotics 30 minutes to one hour earlier, or IV at induction, rectal 2 - 4 hours and oral 6 - 12 hours

Antimicrobial prophylaxis should not continue for more than one day in most cases. This is to avoid suppression of normal flora and superinfection. Giving more than 1 or 2 doses post-operatively is not advised except where specifically recommended such as;

- when there is a delay in starting the operation
- ii) if the operation is prolonged.

Broad-spectrum drugs are usually chosen. All pre-existing infections should be treated before surgery.

#### 13.1 **ORTHOPAEDICS**

### 13.1.1 Clean procedure

Cloxacillin 2 g IV or Cephalothin 2 g IV at the time of induction for procedures involving insertion of prosthetic or transplant material and internal fixation of fractures of large bones.

Allow 5 minutes to elapse between administration of antibiotics and application of tourniquet.

13.1.2 Compound fractures will be classified, as "dirty" and a course of antibiotics should be given as treatment should be given as follows:

- Gentamicin 240mg IV daily PLUS EITHER Cephalothin 1g IV 6 hourly OR
- Ampicillin 500mg IV 6 hourly







### SURGICAL ANTIBIOTIC PROPHYLAXIS

Duration of therapy may vary. If there is no sign of infection antibiotics may be required only for 24 hours.

Consider penicillin G if wound is suspected to be contaminated with *Clostridium* perfringens.

Add metronidazole if there is perineal involvement.

Tetanus toxoid should be given if patient is not immune (last immunisation more than 5 years ago).

#### 13.2 GENTO-URINARY SURGERY

Patients should be shown to have sterile urine pre-operatively. If there is evidence of urinary tract infection the patient should be treated based on the culture and sensitivity results.

If there is clinical evidence of UTI and culture/susceptibility results are unavailable or immediate operation is required, commence

- Ampicillin 1g IV PLU
- Gentamicin 2-3mg/kg IV at the time of induction

### 13.3 CAESARIAN SECTION

- **13.3.1** All patients who have had vaginal examination(s) and/or ruptured membranes:
  - Ampicillin 1- 2g IV at induction (after the cord has been clamped)
- 13.3.2 If any sign of infection is present: full 5-7 days of antibiotic therapy recommended
  - Gentamicin 240mg IV daily (maintenance dose adjusted for renal impairment) PLUS
  - Ampicillin 1g IV 6 hourly

Other antibiotics may be required as indicated by culture and sensitivity results or if there is no clinical response.

### 13.4 <u>ELECTIVE GYNAECOLOGICAL SURGERY</u>

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When the vaginal vault is to be opened, i.e. at hysterectomy.

Metronidazole 400mg orally 4 hours before surgery.



#### ELECTIVE GASTRODUODENAL AND BILIARY SURGERY, 13.5 APPENDICECTOMY AND COLORECTAL SURGERY

- Ampicillin 1-2g IV **PLUS** Gentamicin 2-3mg/kg IV at induction **PLUS**
- Metronidazole suppository 1g rectally 2 hours before surgery as a single dose or 500mg IV (over 20 minutes) to finish at the time of induction.

In low risk surgery, metronidazole is not necessary.

In major colorectal surgery, a second dose of Metronidazole may be required at 4 -6 hours post-induction.

 $\begin{tabular}{ll} Appendicectomy in situations of gangrene, perforation or abscess should be \\ \end{tabular}$ followed by a therapeutic course with ampicillin, gentamicin and metronidazole for approximately 7 days.

#### 13.6 **NEUROLOGICAL SURGERY**

Use of prophylaxis is indicated in prolonged procdures, re-explorations and microsurgery.

- Cephalothin 2g IV **OR**Cloxacillin 2g IV at the time of induction

#### 13.7 OROPHARYNGEAL AND THORACIC SURGERY

Cephalothin 2g IV at time of induction.

Add metronidazole if there is a risk of anaerobic infection.

#### 13.8 **VASCULAR SURGERY**

For arterial reconstructive surgery involving the abdominal aorta and/or the lower limb particularly if a groin incision is involved or if there is implantation of prosthetic material.

- Gentamicin 2mg/kg IV at the time of induction PLUS Cephalothin 2 g IV at induction as a single agent **OR** Cloxacillin 2 g IV at the time of induction

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#### LOWER LIMB AMPUTATION 13.9

- Penicillin G 1.2 g (2 mega units) IV 6 hourly to start at induction and continue for 48 hours **PLUS**Metronidazole 1g rectally 2 hours before surgery **followed by** 400mg orally 8 hourly for 48 hours.

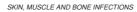
In amputations for diabetic sepsis **add**:
 Gentamicin 2mg/kg IV at induction **followed by** 80 mg IV 8 hourly for 48 hours (dose adjusted for renal function)

# PROPHYLAXIS FOR PATIENTS AT RISK FOR INFECTIVE ENDOCARDITIS 13.10

Refer to Section on Non-Surgical prophylaxis, chapter 9.







#### 14. SKIN, MUSCLE AND BONE INFECTIONS

#### 14.1 SKIN AND SOFT TISSUE INFECTIONS

Mild infections require only good skin hygiene.

Antiseptics such as chlorhexidine or povidone iodine may be useful in some cases.

#### 14.1.1 Impetigo

Mild – above measures are adequate More severe which may be due to Streptococcus or Staphylococcus, consider

- Penicillin V 500 mg orally 6 hourly for 10 days OR
- Procaine Penicillin 1.2 mega units daily for 10 days OR
- Flucloxacillin 500 mg orally 6 hourly for 10 days

For patient hypersensitive to penicillin

Erythromycin 500 mg orally 6 hourly for 10 days

## 14.1.2 Follicullitis, Boils and Carbuncles

Usually caused by Staphylococcus aureus and/or Streptococcus pyogenes

If mild and patient is not diabetic or immunosuppressed, antibiotics are not usually required. If the lesions are small and few in number they may be managed by local antiseptics and hot compresses, with drainage if appropriate.

In other cases:

• Flucloxacillin 500 mg orally 6 hourly for 7 days.

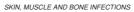
If hypersensitive to penicillin:

• Erythromycin 500 mg orally 6 hourly for 7 days.

If boils are persistent seek expert advice.

### 14.1.3 <u>Cellulitis</u>

Usually due to *Streptococcus pyogenes* and *Staphylococcus aureus* but could be due to variety of organisms. Therefore, do pus and blood cultures if possiblebefore starting therapy, especially in diabetics and immunocompromised patients. Depending upon severity the duration will range from 10 – 14 days.



Mild Infection in patients who are not diabetic or immunosuppressed:

- Penicillin V 500 mg orally 6 hourly for 7 days **OR**Procaine Penicillin 1.2 megaunits IM daily for 7 days

Patients hypersensitive to penicillin:

#### Mild Infection:

Erythromycin 500 mg orally 6 hourly for 7 days

If not responding, add:
• Flucloxacillin 500 mg orally 6 hourly for 7 days.

Severe cases including diabetics and immunosuppressed patients:

Cloxacillin 2g IV 6 hourly initially (until infection subsides), followed by Flucloxacillin 500mg orally 6 hourly to complete a course of 10 - 14 days.

If not responding, change antibiotic according to culture results and /or consult physician or microbiologist.

Patients hypersensitive to penicillin

Cephalothin 2 g IV 6 hourly for 10 -14 days

Note: A small percentage of patients hypersensitive to penicillin will also be hypersensitive to cephalothin (cephalosporins)

#### 14.1.4 Abscess

In mild infections IV cloxacillin is not necessary. A full course of oral flucloxacillin is still indicated.

Incision and drainage should be done if and when indicated. Staphylococcus aureus is the single most important cause. However, do culture and

Cloxacillin 2 g IV 6 hourly generally for 48 hours, or until infection subsides, followed by Flucloxacillin 500 mg 6 hourly to complete a course of 7-14 days

#### 14.1.5 <u>Bites</u>

Bites can get infected with a variety of aerobic and anaerobic organisms.

Wounds require thorough cleaning, debridement and rest. The decision to use antibiotics will depend on the severity of the bite and evidence of infection.

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#### SKIN, MUSCLE AND BONE INFECTIONS

For small superficial wounds seen early, antibiotics may not be necessary. If obviously infected, a wound swab should be taken and the wound debrided and its closure delayed. These require broad-spectrum antibiotic cover.

Tetanus Toxoid should be administered if the patient is not immune (immunisation more than 5 years ago). See immunisation schedule

#### If antibiotic necessary:

Less severe wounds:

- Procaine Penicillin 1.2 mega units IM for 5 days OR
  Procaine Penicillin 1.2 mega units IM as a single dose, followed by Penicillin V 500 mg orally 6 hourly for 5 days.

#### Severe, multiple or infected wounds:

- Metronidazole 400 mg orally 8 hourly **PLUS** Cefaclor 375 mg SR 12 hourly for 7-14 days OR
- Penicillin G 600 mg (1 mega unit) IV 6 hourly for 5-10 days **PLUS** Gentamicin 240 mg IV once daily for 5-10 days (adjust maintenance dose according to renal function).

#### 14.1.6 <u>Surgical or Traumatic wound infections (restricted to skin and soft</u> tissues)

Antibiotics may not be necessary in mild infections. Where there is collection, drain the pus. Take sample for culture and sensitivity before starting antibiotics.

Flucloxacillin 500 mg orally 6 hourly for 10- - 14 days

#### For severe infection:

- Gentamicin 240 mg IV once daily (adjust maintenance dose according to renal function)

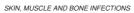
## Alternatively, as a single agent:

Cephalothin 2 g IV 6 hourly.

Antibiotics may need to be changed based on culture and sensitivity results.

#### 14.1.7 <u>Diabetic Foot Infections</u>

Diabetic foot infection may involve the skin and soft tissue as well as underlying muscle and bone, and should always be regarded as serious. They are often caused by mixed infection with aerobes and anaerobes,



gram-positive and gram-negative organisms. Surgical debridement is often necessary. Surgical advice should be sought (this may not be necessary in mild cases)

Proper dressings and wound care are also extremely important.

Adjust drug dosage according to renal function (for drugs excreted via the kidneys), as renal impairment is common in diabetic patients.

The duration of treatment depends upon response.

Mild to Moderate infections:

- Metronidazole 400 mg 8 hourly **PLUS** Flucloxacillin 500 mg orally 6 hourly

#### Severe Infections:

- Metronidazole 400 mg orally 8 hourly PLUS
- Cloxacillin 2 g IV 6 hourly PLUS
- Gentamicin 240 mg IV once daily (adjust maintenance dose according to renal function)

Consider PR Metronidazole 500mg 12 hourly or 500mg IV 8 hourly if patient is too sick to take oral medications.

#### Alternative:

- Metronidazole 400 mg orally 8 hourly PLUS Cephalothin 2 g IV 4 hourly PLUS
- Gentamicin 240 mg IV once daily (adjust maintenance dose according to renal function)

Change to oral therapy when the infection is under control Depending on the organism subsequently isolated, other antibiotics may be indicated. The duration of treatment will depend upon the response.

#### 14.1.8 Necrotising Cellulitis or Fasciitis

Urgent surgical debridement in addition to broad-spectrum antibiotics to cover enterobacteriacea, anaerobes, streptococcus species and Staphylococcus aureus.

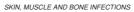
- Penicillin G 2.4 g (4 mega units) IV 4 hourly **PLUS** Gentamicin 240 mg IV once daily (adjust maintenance dose according to renal function) PLUS
- Metronidazole 1 g rectally 12 hourly.











#### Alternative:

- Cephalothin 2g IV 4 hourly PLUS
- Metronidazole 1g rectally 12 hourly

#### 14.1.9 Myositis/ Myonecrosis

Gas gangrene is usually caused by *C perfringens*. It is a surgical emergency.

Penicillin G 2.4 g (4 mega units) IV 4 hourly

If hypersensitive to penicillin:
• Metronidazole 500 mg IV 8 hourly (infused over 20 minutes)

#### 14.1.10 Burns

For minor burns use sterilised gauze dressing impregnated with white soft paraffin, supplied by pharmacy.

For severe burns or if there is evidence of infection:

Topical Silverzine cream (1 % silver Sulphadiazine + 0.2 % Chlorhexidine) It is active against most gram-positive organisms and gram-negative bacteria and yeasts. It can be used with or without a light dressing. It does not penetrate eschar

Systemic antibiotic treatment should not be used routinely but only to treat infections based on culture results.

A single dose of an antibiotic maybe given before surgical debridment, in accordance with the latest microbiology results.

## 14.1.11 Pyomyositis

Staphylococcus aureus is the most common organism

Cloxacillin 2 g IV 6 hourly

Duration of therapy depends upon severity and may vary from 14 –21 days. Oral therapy should be instituted as soon as it is appropriate.

If hypersensitive to penicillin:

Cephalothin 2 g IV 6 hourly

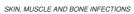
Note: a small percentage of patients hypersensitive to penicillin will also be hypersensitive to cephalothin (cephalosporins)

Surgical drainage may be necessary. Antibiotics may need to be adjusted according to culture results.

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#### 14.1.12 Mastitis

Acute mastitis is usually associated with lactation and is frequently due to Staphylococcus aureus. Milk stasis is to be avoided and therefore suckling and manual expression are important.

In the absence of systemic symptoms these measures are adequate.

If systemic symptoms develop, start antibiotics

Flucloxacillin 500 mg orally 6 hourly

In severe cases IV therapy may be required.

Cloxacillin 2 g IV 6 hourly.

In patients hypersensitive to Penicillin

Cephalothin 2g IV 6 hourly Duration of therapy will depend upon response.

Note: a small percentage of patients hypersensitive to penicillin will also be hypersensitive to cephalothin (cephalosporins)

#### 14.2 **BONE INFECTIONS - OSTEOMYELITIS**

The usual causative agent is Staphylococcus aureus, or occasionally streptococcus sp.

Intravenous treatment should be given until the patient has been afebrile for several days, when appropriate oral treatment can be substituted. The duration of treatment should be at least 4 and usually 6 weeks. Do culture and sensitivity.

## Empirical therapy:

Cloxacillin 2 g IV 6 hourly, followed by Flucloxacillin 1 g orally 6 hourly, for a total course of 6 weeks or longer if necessary, especially in chronic

If oral flucloxacillin causes nausea and vomiting the dose can be reduced to
• Flucloaxillin 500 mg orally 6 hourly **PLUS**• Probenecid 500 mg 12 hourly

In severe infections consider adding:

Gentamicin 240 mg IV once daily for the first 3-5 days (adjust maintenance dose according to renal function)

In patients hypersensitive to Penicillin
 Cephalothin 2g IV 6 hourly





- In Methicillin resistant Staphylococcus aureus use;
   Vancomycin 1g IV 12 hourly (infuse over 1 hour) followed by
   Rifampicin 600mg orally daily
   The duration of treatment is as above.

#### SEPTIC ARTHRITIS 14.3

Empirical therapy:
Cloxacillin 2 g IV 6 hourly for at least 2 weeks or until the patient has been afebrile for a few days, **followed by** Flucloxacillin 1 g orally 6 hourly

Duration of treatment and the use of Probenecid is the same as that stated

under oeteomyelitis
Urgent surgical drainage /lavage and microbiological examination of pus
maybe required. Change antibiotics if necessary.







#### EYE INFECTIONS

#### 15. EYE INFECTIONS

Note: In eye an infection there is no place for local steroids, unless specifically recommended by an ophthalmologist.

#### 15.1 ACUTE BACTERIAL CONJUNCTIVITIS

Red and sticky eyes with pus discharging in varying degrees.

- Chloramphenicol 0.5% eye drops 1 drop in both eyes during daytime for 1 week. Chloramphenicol 1% eye ointment applied at night OR
- Polyantibiotic eye drops hourly during day time and Polyantibiotic eye ointment applied at night.
- Gentamicin 0.3% eye drops to be used if no response on above regimen or organism sensitivity demands.
- Tetracycline 1% eye ointment to be used at least 4 times a day or preferably every 2 hours according to the severity of the infection.

#### 15.2 TRACHOMA

Conjunctivitis caused by Chlamydia trachomatis

- Tetracycline 1% eye ointment twice a day in both eyes for minimum of 6 weeks. Repeat if necessary for another 6 weeks after an interval of 6 months OR
- Erythromycin 500mg orally 8 hourly for 14 days.

## 15.3 EYE LID INFECTIONS

Stye or tarsal cysts.

Painless tarsal cyst – no antibiotic therapy is necessary.

If infected:

• Use same regimen as in acute conjunctivitis.

#### 15.4 CORNEAL INFECTION

Corneal ulcer with or without pus in anterior chamber.

- Gentamicin 0.3% eye drops to be reviewed after organism sensitivity is known. Gentamicin to be used half hourly in affected eye with active ulceration. Treatment to be supplemented with subconjunctival injection if pus is present in anterior chamber. (This injection should be given preferably by an Ophthalmologist).
- If Gentamicin not available, polyantibiotic or Chloramphenicol eye drops to be used.









#### EYE INFECTIONS

Frequency is reduced with clinical improvement

Atropine 1% eye ointment must be applied 3 times daily along with antibiotics in active corneal ulceration.

#### **DENDRITTIC CORNEAL ULCERATION** 15.5

Aetiology is herpes simplex virus

Acyclovir 3% eye ointment 3 hourly during day time and 4 hourly at night. Continue for at least 3 days after complete healing.

#### Alternative

Idoxuridine\* 0.1% eye drops hourly during daytime and 2 hourly at

Antibiotic eye drops have no place here.

#### **INJURIES** 15.6

#### (a) Non perforating eye injuries where cornea is not involved Non-Infected:

Chloramphenicol 0.5% eye drops at least 4 times daily during day (to prevent infection) for up to 7 days.

#### Infected:

If infected as suggested by sticky discharge, use the regimen as for acute conjunctivitis.

 (b) Cornea involved, without infection
 Chloramphenicol 1% eye ointment stat (applied once only), and eye pad daily until cornea is healed. Healing should take place within a few days.

# (c) Cornea involved and infection present (suggested by corneal opacification around injury, redness and discharge)

Treated as for corneal ulcer

#### 15.7 **ORBITAL CELLULITIS**

This is a serious infection requiring specialised Ophthalmology care and referral to an ophthalmologist.

Cloxacillin 2 g IV 6 hourly PLUS

Gentamicin 240 mg IV once daily (maintenance dose adjusted

- according to renal function)

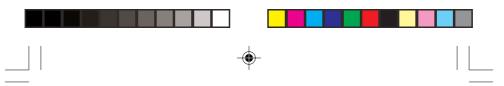
If response is inadequate ADD

Ampicillin 2 g IV 6 hourly

\* NOT ON EDL





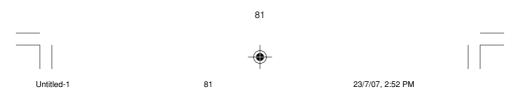


EYE INFECTIONS

## 15.8 OPTHALMIA NEONATORUM

See Sexually Transmitted Infections







#### **16. DENTAL INFECTIONS**

#### 16.1 **ACUTE DENTO-ALVEOLAR INFECTIONS** (Abscesses and Periodontal infections)

#### (a) Abscess without Cellulitis

- Penicillin 500 mg orally 6 hourly for 5 days **OR**
- Amoxycillin 500 mg orally 8 hourly for 5 days.

#### If hypersensitive to Penicillin

Erythromycin 500 mg orally 6 hourly for 5 days

#### (b) Abscess with Cellulitis

- Penicillin 1g stat and 500 mg 6 hourly for 7 days OR
- Amoxycillin 500 mg orally 8 hourly for 7 days PLUS EITHER OF THESE WITH
- Metronidazole 200 400 mg orally 8 hourly for 7 days.

#### If hypersensitive to Penicillin:

- Erythromycin 500 mg orally 6 hourly for 7 days PLUS
- Metronidazole 200 400 mg orally 8 hourly for 7 days.

#### Severe Infection:

- Penicillin G 600 mg (1 mega unit) IV 4 6 hourly **OR** Ampicillin 1 g IV 4 6 hourly **PLUS (either of these with)**
- Metronidazole 400 mg 8 hourly for 7 days.

#### 16.2 **PERICORONITIS**

#### (Infection of gums around teeth)

- Metronidazole 200 mg orally 8 hourly for 5 days OR
- Penicillin V 250 mg orally 6 hourly for 5 days OR
- Amoxycillin 500 mg 8 hourly for 5 days PLUS
- Chlorhexidine 0.2% aqueous solution mouthwash

#### **ACUTE NECROTISING ULCERATIVE GINGIVITIS** 16.3

- Metronidazole 400 mg orally 8 hourly for 7 days Peroxide\* or Perborate\* mouth rinse 3 times a day hold it for 2 minutes **PLUS**
- Chlorhexidine 0.2% aqueous solution mouthwash 3 times a day.

Scaling, oral hygiene and appropriate pain relief are also important.

\* NOT ON EDL









#### 16.4 **ORAL ULCERS**

- Antiseptic oral rinse e.g. Chlorhexidine 0.2% aqueous solution mouthwash 3 times daily for 7-14 days **PLUS**
- Topical use of analgesic/ anaesthetic e.g. Choline salycilate\* **PLUS** Triamcinolone 0.1% in Orabase\* 2 3 times daily for 7 14 days.

#### 16.5 **CANDIDIASIS**

- Nyastatin suspension 100,000 U (1mL) 4 times a day (swirl around the mouth and swallow) for 7 14 days  $\bf OR$  Nyastatin Lozenges\* 1 lozenge (dissolves slowly in mouth) 4 times a
- day for 7 14 days.

For severe infections needing systemic therapy, consider amphotericin, ketoconazole or fluconazole\*. These patients require referral.

#### 16.6 **FACIAL FRACTURES**

- Penicillin V 500 mg orally 6 hourly for 7 days **OR**
- Amoxycillin 500 mg 8 hourly for 7 days

If patient cannot swallow tablets or capsules, suspension may be used.

- If parenteral therapy is considered necessary:
   Procaine penicillin 1.2 mega units IM daily for 5 days **OR**
- Ampicillin 1g IV 6 hourly for 5 days

If hypersensitive to Penicillin:

Erythromycin 500mg 6 hourly orally (or IV if severe) 6 hourly for 7 days.

Gentamicin 240mg IV once daily for 7 days as may be required.

Antiseptic mouthwash e.g. Chlorhexidine 0.2% aqueous solution for 4-6weeks and analgesics as required.

#### **DENTO-ALVEOLAR SURGERY** 16.7 (Tooth extraction, cyst removal etc)

Indications for antibiotic therapy

- (i) Signs of infection e.g. fever (i) Excessive trauma during surgery
- Penicillin V 500 mg orally 6 hourly for 5 days OR
- Amoxycillin 500 mg orally 8 hourly for 5 days

\* NOT ON EDL

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If hypersensitive to Penicillin:
• Erythromycin 500 mg orally 6 hourly for 5 days.

Analgesics, anti-inflammatory and decongestant therapy as required.

Patients with valvular heart disease require prophylaxis – see Non Surgical Prophylaxis – chapter 9

#### ALVEOLITIS (DRY SOCKET) 16.8

Antibiotics are required only if there are systemic signs of infection. Various dental dressings, antiseptics mouthwash as required.







#### General Information – Prescribing Antimicrobials in Children

- 1. In general, children over 40kg should receive adult dose.
- 2. In all cases the maximum dose should not exceed adult dosage.
- Special care should be taken in neonates, especially in the first few weeks of life.
   Dosage intervals may need to be extended owing to poor drug metabolism and excretion.
- All intravenous antimicrobials should be given by slow infusion to avoid thrombophlebitis.
- Gentamicin levels should be monitored whenever possible; renal function however needs to be monitored in all children on gentamicin

#### CARDIOVASCULAR SYSTEM INFECTIONS

#### **BACTERIAL ENDOCARDITIS**

There are 3 important principles of management:

- 1. Treatment must be given intravenously for at least 2 weeks
- 2. Treatment is prolonged usually 4-6 weeks
- 3. Drug regimens must be used in high enough concentrations.

There are several alternative regimens reported in the literature and consultation with a physician or microbiologist should be sought if necessary. Surgical consultation should be considered especially in cases, which are fulminating, complicated or slow respond

Take blood for culture before initiating therapy. *S aureus* is not an uncommon cause of endocarditis in Fiji, hence cloxacillin is included in the empirical regimen. Other common organisms include *viridans* group streptococci and occasionally gram-negative bacilli.

#### Empirical treatment:

- Penicillin G 100,000 units/kg/dose (max 3 mega units/dose) 4 -6 hourly for 6 weeks PLUS
- Cloxacillin 50mg/kg/dose (max 2g/dose) 4 6 hourly for 6 weeks **PLUS**
- Gentamicin 2 2.5mg/kg/dose (max 80 mg/dose) 8 hourly IV for 2 weeks (dose adjusted for renal function)







Monitor gentamicin levels.

If hypersensitive to Penicillin:

• Vancomycin 10 – 15mg/kg/dose (max 50 mg/dose) IV (infused over 1 hour) 8 hourly

Blood levels should be monitored if possible

Drug therapy may need to be altered based on culture and sensitivity results. If the blood culture is negative, continue with empirical therapy if the patient is responding.

#### PREVENTION OF INFECTIVE ENDOCARDITIS

All children with structural heart disease (congenital or rheumatic) require antibiotic prophylaxis.

- (a) Low risk patients (i.e. without prosthetic valves or previous attack of endocarditis) undergoing dental treatment, oral surgery or surgical procedures of the respiratory tract.
  - Ampicillin 50mg/kg (max 1g) IV just before procedure OR Amoxycillin 50mg/kg (max 2 g) orally 1 hour before procedure,
- (b) Low risk patients (as above) having gastrointestinal, genitourinary procedures and other major surgery
  Prophylaxis as for this section (c) below.

- (c) High risk patients (i.e. prosthetic valve, previous attack of the infective endocarditis ) undergoing any type of dental or surgical procedures
  - Ampicillin 50mg/kg (max 1g IV just before procedure followed by Ampicillin 25mg/kg (max 500mg) IV/IM or Amoxycillin 25mg/kg (500mg) orally 6 hours later **PLUS**
  - Gentamicin 2-2.5 mg/kg IV just before the procedure.

If hypersensitive to penicillin:

Vancomycin 20mg/kg IV (max 500mg) (infused over 1 hour) started 60minutes before procedure. May repeat once after 12 hours.









#### RESPIRATORY INFECTIONS

#### **ACUTE SORE THROAT**

Streptococcal sore throat is diagnosed by the presence of tender enlarged lymph nodes in the neck and white exudates in throat or  $\_$  –haemolytic streptococcus group A on throat swab culture.

- Benzathine Penicillin:  $25\ 000-50\ 000$  units /kg/ dose (max 1.2 mega
- units) as a single IM dose **OR** Penicillin V 7.5 15mg/ kg/ dose (max 500 mg/ dose) orally twice daily for 10 days **OR** Amoxycillin 10 – 20mg/kg/dose (max 500 mg/ dose) orally 8 hourly for
- 10 days.

If hypersensitive to penicillin:

Erythromycin 10 - 12.5mg/kg/dose (max 500mg/dose) orally 6 hourly for 10 days.

#### **ACUTE BACTERIAL OTITIS MEDIA & SINUSITIS**

Common organisms are *Streptococcus pneumoniae, Haemophilus influenzae & Staphylococcus aureus.* Diagnostic criteria: pus draining from ear for less than 2 weeks or red, bulging immobile eardrum

> Amoxycillin 10 - 20mg/kg/dose (max 500mg) orally 8 hourly for 7 - 10 days.

The following shows Amoxycillin oral dosage according to age/weight:

62.5 mg 8 hourly < 2 months (< 5kg): 2 months up to 12 months (6 – 9kg): 12 months up to 5 years (10 – 19 kg): 125 mg 8 hourly 250 mg 8 hourly OR

Co – trimoxazole syrup (40mg Trimethoprim & 200mg Sulphamethoxazole/ 5ml). Dosage for co-trimoxazole is calculated as Trimethoprim 2.5mg/ kg/ dose orally 12 hourly for 7 -10 days.

The following shows Co-trimoxazole syrup dosage according to age/weight:

2.5mLs 12 hourly 1-2 months (< 5kg): 2 months to 12 months (6-9 kg): 5.0mLs 12 hourly 12 months to 5 years (10 – 19 kg): 7.5mLs 12 hourly

#### Infants < 2 months ideally should be admitted and treated as follows (Note: avoid Co-trimoxazole infants < 1 month)

- Ampicillin 25 50mg/kg/dose IV 6 hourly for 7 10days PLUS
- Gentamicin 2 2.5 mg/kg/dose IV 12 hourly for 5 7 days







If hypersensitive to penicillin:

Erythromycin 10 – 12.5mg/kg/dose (max 500mg/dose) orally 8 hourly for 7 days

#### **ACUTE EPIGLOTITTIS**

This is paediatric emergency. Stridor of epiglotittis is usually of lower pitch than that heard in croup and there is expiratory element which resembles a snore. These children can also present with aphonia and drooling. The most common microorganism is *Haemophilus influenzae*.

Do not attempt to examine the throat as this can precipitate a laryngeal spasm

causing acute obstructive episode. Refer for admission.

• Chloramphenicol 50mg/kg (max 2g) IV as a loading dose followed by 25mg/kg/dose (max 1g/dose) IV 6 hourly for 5 days, or longer if septicaemia is present.

#### **COMMUNITY ACQUIRED PNEUMONIA**

Infants under 2 months of age are considered to have severe pneumonia – see Moderate to Severe disease below.

#### Mild Disease

 $\frac{Infant\ age:\ 2months-1\ year\ (6-9\ kg)}{Clinically-cough\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ tachypnoea\ (respiratory\ rate\ 50\ breaths\ per\ minute\ or\ and\ per\ noea\ (respiratory\ rate\ per\ noea\ per\ noe$ more). No chest indrawing.

- Procaine Penicillin 400,000 units/dose IM daily for 5 days OR
- Amoxycillin syrup (125mg/5mL) 10–20 mg/kg/dose (max 125mg) orally 8 hourly X 5 days **OR**
- Co-trimoxazole syrup (40mg Trimethoprim and 200mg Sulphamethoxazole in 5ml), dosage for Co-trimoxazole is calculated as trimethoprim 2.5mg/kg/dose orally 12 hourly for 5 days

Review child after two days or sooner if child is not improving.

#### Children aged 1 to 5 years

Cough, no chest indrawing. Fast breathing is 40 breaths per minute or more.

- Procaine Penicillin 25,000 50,000 units/kg (max 1 mega unit) IM daily for 5 days **OR**
- Amoxycillin syrup (125mg/5mLs) 10 -20 mg/kg/dose (max 250mg /dose) orally 8 hourly for 5 days OR
- Co-trimoxazole syrup (40mg Trimethoprim & 200mg







Sulphamethoxazole in 5mLs), dosage for co-trimoxazole is calculated as trimethoprim 2.5mg/kg/dose orally 12 hourly for 5 days.

Review as above.

#### Children over age 5 years

- Procaine Penicillin 25,000 50,000 units/kg (max 2 mega unit) IM daily for 5 days OR
- Amoxycillin syrup 10 20mg/kg/dose (max 500 /dose) 8 hourly orally for 5 days

Review as above.

#### Moderate to severe disease

Clinical finding of chest indrawing in child with cough, fever. Before starting antibiotics blood or any other samples required should be taken for culture

#### Infants less than 2 months

All pneumonia in infants less than two months should regarded as severe and therefore child should be admitted and treated.

Severely malnourished children should receive double antibiotic therapy to cover gram positive and gram-negative organisms, until culture and sensitivity results are available.

#### Regimen 1:

- Penicillin G 50,000 units/kg/dose IV X 7 days 6 hourly in infants over three weeks old 8 hourly IV in infants 1-3 weeks old 12 hourly in infants < 7 days old PLUS
- Gentamicin 2 2.5mg/kg/dose IV 12 hourly in infants < 2months.

## Regimen 2

- Ampicillin 50mg/kg/dose IV 6 hourly PLUS
- Gentamicin 2.5mg/kg/dose IV 12 hourly in infants < 2 months

If there is any suggestion of Staphylococcal infection:

- Cloxacillin 25mg/kg/dose IV 6 hourly **PLUS** Gentamicin 2 2.5mg/kg/dose IV as above

The total duration of therapy is 7 - 10 days













#### Children aged 2 months and over

- Penicillin G 50,000units/kg/dose (max 1 mega unit / dose) IV 6 hourly for 3 days **followed by** Procaine penicillin 50,000 units/kg/dose (max 2 mega units) IM daily to complete at least 7 days OR
- Ampicillin 25mg/kg/dose (max 1g /dose) IV 6 hourly for 3 days followed by Amoxycillin 20mg/kg/dose (max 500mg/ dose) orally 8 hourly to complete at least 7 days OR
- Chloramphenicol 12.5mg 25mg /kg/dose (max 1g /dose) IV 6 hourly for at least 3 days **followed by** Chloramphenicol 12.5 mg 25mg/kg/dose (max 500mg/dose) orally 6 hourly to complete at least 7 days.

Where Staphylococcal infection is suspected;
• Cloxacillin 25 – 50 mg /kg/dose (max 500mg/dose) IV 6 hourly for 7 days

#### **ATYPICAL PNEUMONIA (Non-Viral)**

Mycoplamsa pneumoniae can cause pneumonia in children. Onset is more gradual than pneumonia due to Streptococcus pneumoniae or H. influenzae.

Erythromycin 10 -12.5mg/kg/dose (max 500mg/dose) IV 6 - 8 hourly (by slow infusion in Normal Saline) OR orally (depending on clinical condition) for 7 - 10 days



## **LUNG ABSCESS AND/OR EMPHYSEMA**

Usually staphylococcal in aetiology. Other organisms including anaerobes may be involved; consider the addition of other antibiotics if necessary e.g. Metronidazole or Chloramphenicol. Empyma must be drained adequately. Surgical drainage of lung abscess will depend on the size.

- Cloxacillin 50-mg/kg/dose (max 2g/dose) IV 6 hourly for 2 weeks followed
  - Flucloxacillin 25mg/kg/dose (max 500mg /dose) orally 6 hourly. Treatment is usually for 10 -14 days, or longer if required depending upon the response.

#### PLUS

Gentamicin 2 –2.5mg/kg/dose (max 80mg/dose) IV 8 hourly for 5 – 7 days (to start at the same time as cloxacillin)

Clinician may decide to add Rifampicin  $10-15 mg/kg \ (max \ 600 mg)$  orally daily for 10 –14 days together with Cloxacillin (never alone)







#### **GASTROINTESTINAL INFECTIONS**

#### **ACUTE GASTROENTERITIS**

 $\label{eq:continuous} Oral\ rehydration\ the rapy-(ORS-Oral;\ Rehydration\ Solution)\ is\ still\ the\ mainstay\ of$ treatment to correct dehydration and replaces losses.

The indications for antibitotic treatment in acute diarrhoea are cholera and bloody diarrhoea. Regimens shown below are for empirical therapy where immediate treatment is required. Antibiotic therapy should be modified if necessary according to culture and sensitivity results

#### **Bloody Diarrhoea**

Rule out non-bacterial causes of bloody stools and collect stool sample for culture and sensitivity testing before starting antibiotic therapy. Antibiotic therapy is generally not recommended in uncomplicated diarrhoea or mild cases.

#### Regimens include:

- Co-trimoxazole (Trimethoprim 40mg and Sulphamethoxazole 200mg/5mLs) calculated as Trimethoprim 2.5/kg/dose, 12 hourly for 5 days **OR**Chloramphenicol 12.5 mg/kg/dose (max 500mg/dose) orally 6 hourly for 5
- Ceftriaxone 50mg/kg daily IV X 3 days in dysentery with seizures (neurotoxin)

## Cholera

Co-trimoxazole suspension (Trimethoprim 40mg and Sulphamethoxazole 200mg/5mLs) calculated as Trimethoprim 2.5mg/kg/dose, 12 hourly for 3-4 days.

#### **CHRONIC DIARRHOEA**

Chronic diarrhoea also requires investigation for non-infective causes

#### Giardiasis:

Metronidazole 6mg/kg/dose (max 400mg/dose) orally 8 hourly for 7 – 10 days

## Amoebic Dysentery:

Metronidazole 10mg/kg/dose (max 400mg/dose) orally 8 hourly for 7 -10 days

#### Strongyloidosis:

Thiabendazole 25mg/kg/dose (max 1g/dose) orally 12 hourly for 3 days.











#### TYPHOID /PARATYPHOID FEVER

Chloramphenicol 25mg/kg/dose (max 750mg/dose) IV or orally 6 hourly for 10 -14 days

# CENTRAL NERVOUS SYSTEM INFECTIONS

#### **ACUTE BACTERIAL MENINGITIS**

Treatment of meningitis in children includes:

- Antibiotic therapy
- 2) Dexamethasone
- Therapy to eradicate nasopharyngeal carriage of *H. influenzae* and meningococcus, both for index case and contacts see Non Surgical Prophylaxis chapter 13

#### Infants two months and above

Usually blood cultures are taken and a CSF specimen is taken for culture and bacterial antigens before institution of antibiotics, unless there is a contraindication to lumbar puncture, for example a very sick child and/or raised intracranial pressure



- Penicillin G 100,000 units/kg/dose (max 4 mega units/dose) IV 6 hourly **PLUS**
- Chloramphenicol 25mg/kg/dose (max 1g /dose) IV 6 hourly

Duration of treatment varies from 10 to 14 days depending on causative organism and clinical state of the child.

#### Added Drugs:

Dexamethasone 0.15mg/kg/dose IV 6 hourly for 4 days (first dose to be given before antibiotics are given).

If there is no clinical response or bacteria resistant to above drugs:

Ceftriaxone 100mg/kg (max 4g/dose) IV as a single dose daily for 10-14days







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When the brain abscess is suspected then consider addition of the following drugs, and treat for 6 weeks. CT scan and surgical drainage may be indicated.

- Metronidazole 15mg/kg stat then 7.5mg/kg/dose IV 8 hourly  $\mathbf{OR/PLUS}$
- Cloxacillin 50mg/kg/dose (max 2g/dose) IV 6 hourly

- In infants under the age of 2 months
   Ceftriaxone 100mg/kg/dose IV in a single daily dose PLUS
- Ampicillin 50-75mg/kg/dose IV 6 hourly

Treatment should be for 21 days.

#### **HERPES SIMPLEX ENCEPHALITIS** (SUSPECTED OR PROVEN)

Acyclovir 500mg/m²/dose IV 8 hourly for 14 days, each dose to be infused over not less than one hour (dosage adjusted according to renal function)

[Body Surface Area in m2:

Multiply height (in cm) by weight (in kg), and then divide the answer by 3600. Find the square root of this figure.]



#### **URINARY TRACT INFECTIONS**

Appropriate collection of urine is essential before starting antibiotics. Adjust antibiotics if necessary when culture and sensitivity results are available. Treat for 7 – 10 days. All children with proven UTI should undergo investigations to determine whether there is some abnormality in the urinary system.

- Trimethoprim suspension (50mg/5mLs) 2.5mg/kg/dose, 12 hourly (maximum 300mg/day) for 7 -10 days **OR**
- Amoxycillin 10 20 mg/kg/dose (max 500mg/dose) orally 8 hourly for 7 10

Serious cases of UTI (temperature equal to or more than 39° C), dehydrated, not tolerating oral medication, not feeding well and less than 3 months of age of age need admission.

- Ampicillin 25 50 mg/kg/dose (max 1g/dose) IV 6 hourly PLUS
- Gentamicin 2mg/kg/dose (max 80mg/dose) IV 8 to 12 hourly OR
- Ceftriaxone 50mg/kg/dose dose (max 1g/day) IV daily as a single agent



Children with vesico-uretic reflux will need to be on prophylactic antibiotics:

- Trimethoprim suspension (50mg/5mLs) 1-2 mg/kg as a single dose daily at night OR
- Co-trimoxazole suspension (Trimethoprim 40mg and Sulphamethoxazole 200mg/5mLs) calculated as Trimethoprim 1-2mg/kg/dose at night (avoid in children under 1 month old) **OR**
- Nitrofurantoin 1-2mg/kg/dose (max 50mg/dose) at night (avoid in children under 3 months old) OR
- Nalidixic acid 12.5mg/kg/dose at night (avoid in children under 3 months old)

#### SKIN, MUSCLE AND BONE INFECTIONS

#### **SKIN AND SOFT TISSUE INFECTIONS**

Mild disease requires only good skin hygiene. Topical antibiotics are not required.

#### Impetigo

Can resolve with topical agents e.g. Chlorhexidine or Povidone iodine. If oral antibiotics are required, they need to cover the causative agents, usually *Staphylococcus aureus*, occasionally *Streptococcus pyogenes*.

Flucloxacillin 12.5mg/kg/dose (max 500mg/dose) orally 6 hourly for 5-7 days.
 Do culture & sensitivity of pus and change antibiotic if necessary.

If hypersensitive to penicillin:

 Erythromycin 10 – 12.5mg/kg/dose (max 500mg/dose) orally 6 – 8 hourly for 5 – 7 days

#### Folliculitis, Boils and Carbuncles

Usually caused by Staphylococcus aureus and /or Streptococcus pyogenes.

If small and few apply local antiseptics and hot compresses with drainage where appropriate.

If lesions are extensive or if there are systemic signs:

- Cloxacillin 12.5mg/kg/dose (max 500mg/dose) IV 6 hourly until there is a clinical response, followed by
- Flucloxacillin 12.5mg/kg/dose (max 500mg/dose) orally 6 hourly for a total of 7 days











In some cases oral flucloxacillin may be all that is required.

If hypersensitive to penicillin:

Erythromycin 10 – 12.5mg/kg/dose (max 500mg/dose) orally 6 -8 hourly for 7 days.

#### Cellulitis

Usually due to *Streptococcus pyogenes* and *Staphylococcus aureus* but could be due to variety of organisms. Therefore do culture before starting therapy

- due to variety of organisms. Therefore, do culture before starting therapy.
   Cloxacillin 25 50mg/kg/dose (max 1g/dose) IV 6 hourly until there is a clinical response, followed by
- Flucloxacillin 12.5 mg/kg/dose (max 500mg/dose) orally 6 hourly to complete a total of 5-10 days.

Consider the following if flucloxacillin is not available, or addition of Penicillin if patient is not responding to cloxacillin:

- Penicillin G 50, 000 units/kg/dose IV 6 hourly until there is clinical response, followed by
- Amoxycillin 10 -20 mg/kg/dose (max 500mg/dose) orally 8 hourly to complete a total of 5-10 days.

Mild Cases;

Consider oral amoxycillin or flucloxacillin; dosage as above.

#### Erysipelas

- Penicillin G 50,000 100,000 units/kg/dose (max 4 mega units/dose) 6 hourly until there is clinical response, followed by
- Procaine Penicillin 50,000 units/kg IM daily to complete 7 days

#### Abscess

Incision and drainage should be done if and when indicated. *Staphylococcus aureus* is the single most important cause. However, do culture.

If abscess is mild and there are no signs of systemic infections:

Flucloxacillin 12.5mg/kg/dose (max 500mg/dose) orally 6 hourly for 7 days

In moderate to severe cases:

- Cloxacillin 25-50mg/kg dose (max 1g/dose) IV 6 hourly until there is clinical response followed by
- Flucloxacillin 12.5mg/kg/dose (max 500mg/dose) orally 6 hourly for 7 days









#### Supparative Wound Infections (Surgical or Traumatic)

Local measures such as surgical drainage and irrigation with normal saline usually suffice. If there is surrounding cellulitis and/or systemic symptoms are present - take wound swab and treat with:

Flucloxacillin 12.5mg /kg/dose (max 500mg / dose) orally 6 hourly

- If gram-negative organisms are suspected add:
  Gentamicin 2mg/kg/dose (max 80mg/dose) IV 8 –12 hourly **OR**Cephalothin 25 –50mg /kg/dose (max 2g/dose) IV 6 hourly **OR**Ceftriaxone 50mg/kg/dose (max 2g/dose) IV as a single dose daily.

Duration depends on response to therapy, approximately 7 days or may be longer.

Antibiotics may need to be changed based on culture and sensitivity results.

#### **ACUTE OSTEOMYELITIS**

Advice from orthopaedic surgeon is to be sought in all cases of osteomyelitis (acute and chronic)

- Cloxacillin 50 mg/kg/dose (max 2g/dose) IV 4 6 hourly for 2 -4 weeks followed by
- Flucloxacillin 25mg/kg/dose (max 1g/dose) orally 6 hourly for a total of 4 -6 weeks PLUS
- Gentamicin 2mg/kg/dose (max 80mg/dose) IV 8 hourly for 1 week

#### Alternative:

Ceftriaxone 50mg/kg/dose (max 2g/dose) IV as a single dose daily for 4-6weeks

#### In children under 5 years

In this age group *H.influenzae* is not an uncommon causative organism. If cultured, to cloxacillin therapy (as above) add:

• Ampicillin 25 – 50 mg/kg/dose IV 6 hourly for 2 -4 weeks, followed by

- Amoxycillin 25 50mg/kg/dose orally 8 hourly to complete a total of 4 6 weeks









## SEPTIC ARTHRITIS

Aspiration and culture of synovial fluid and surgical consultations is essential. Treatment should be altered according to culture and sensitivity results.

- Empirical therapy:
  Cloxacillin 50mg/kg/dose (max 2g/dose) IV 6 hourly for at least 2 -4 weeks followed by

  "" 65 mg/kg/dose (max 1g/dose) orally 6 hourly for a total of
- Flucloxacillin 25 50 mg/kg/dose (max 1g/dose) orally 6 hourly for a total of 4-6 weeks

#### In neonates and Infants ADD:

Gentamicin 2mg/kg/dose IV 12 hourly for 1 week initially

#### Alternative:

Ceftriaxone 50mg/kg/dose (max 2g /dose) IV as a single dose daily for 4 – 6 weeks

## ORBITAL CELLULITIS

The most common organisms are *streptococci*, *staphylococci*, *H influenzae*. Other organisms are possible, hence the use of gentamicin

Cloxacillin 50mg/kg/dose (max 2g/dose) IV 6 hourly **PLUS**Gentamicin 2mg/kg/dose (max: 80mg/dose) IV 8 – 12 hourly for 7 – 10 days.







#### 18. SPECIAL INFECTIONS

#### MRSA INFECTION/COLONISATION

Infections due to methicillin resistant staphylococci (MRSA) are increasing in Fiji. All patients or staff from whom MRSA is isolated will not necessarily require systemic antibiotic therapy

The drug of choice in treating serious MRSA infection is vancomycin, however, it is expensive, is not readily available and is not efficient in eradicating the organisms from carriage sites such as the pharynx, nares and gut.

For treating serious infections (septicaemia, endocarditis etc) vancomycin injection is recommended, to be given according to the dosage regimens in the individual chapters.

If vancomycin is not available, and in conditions such as diabetic foot ulcers due to MRSA, a combination of rifampicin with another anti-staphylococcal drug like fusidic acid can be used. Since resistance to rifampicin develops during treatment, this drug should never be given alone. The choice of these drugs should be based on sensitivity test results. Consult microbiologist for advice.

- Rifampicin 300- 600mg orally 12 hourly PLUS
- Fusidic Acid 500mg orally 8 12 hourly for 2 weeks

Some patients may not be able to tolerate this regimen and other combinations may have to be tried.

For clearance of MRSA carriage topical application of mupirocin ointment may be useful.

#### **LEPTOSPIROSIS**

- Penicillin G 1.2g (2 mega units) IV 6 hourly for 7 days OR
- Tetracycline 500mg orally 6 hourly for 7 days OR
- Erythromycin 500mg orally 6 hourly for 7 days

Note: Chloramphenicol is not effective.





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#### 19. TREATMENT OF TUBERCULOSIS

The WHO declared a global emergency in 1993 in recognition of its growing importance in Public Health problems. A new framework for effective TB control was developed and a global strategy called DOTS (Directly Observed treatment Short course) was introduced. This is the method of treatment of TB and wherever possible, should be the treatment of

Diagnosis of TB wherever possible should be based on smear examination, culture, imaging techniques and if indicated tissue biopsy.

Treatment of TB consists of multi-drug regimen administered to the patient for a period of 6-9 months, or longer in exceptional cases. Choice of regimen and its length is determined by the category of the disease. Vast majority of TB cases present as pulmonary TB. Ensuring adherence with treatment and performing appropriate contact tracing and screening, with institution of preventive treatment if appropriate is vital.

Prior to treatment a baseline renal function and liver function is important. Visual acuity is to be assessed if Ethambutol is used.

#### Essential Anti-Tuberculosis Drugs

There are three main properties of antituberculosis drugs: bactericidal activity, sterilizing activity and ability to prevent resistance. Rifampicin and isoniazid are the most powerful bactericidal drugs, active against all populations of TB bacilli. Rifampicin is the most potent sterilizing drug available. Pyrazinamide and streptomycin are also bactericidal against certain population of TB bacilli. Streptomycin is bactericidal against rapidly multiplying TB bacilli. Ethambutol is used in association with more powerful drugs to prevent the emergence of resistant bacilli, as it is bacteriostatic.

Rifampicin, Isoniazid and Pyrazinamide are hepatotoxic. Streptomycin and ethambutol are excreted by the kidney.

Combination preparations available in Fiji include rifampicin and isoniazid (Rifampinah), which comes in two strengths: 300/150mg and 150/100mg

In adults pyridoxine is usually given with isoniazid to mimimise neurological side effects.

#### 19.1 Standard Short Course Therapy

- Isoniazid 300mg orally daily X 6 months **PLUS**Rifampicin 600mg (<50 kg: 450mg) orally, daily X 6 months **PLUS**
- Pyrazinamide 2g (< 50kg : 1.5g) orally, daily X 2 months Ethambutol 15mg / kg orally, daily X 2 months

Streptomycin 15mg/kg IM daily can be added in patients intolerant to any of the drugs mentioned above.

Pyridoxine 25mg orally daily is given as a routine with the above drugs.



#### TREATMENT OF TUBERCULOSIS

To maximize adherence and drug levels, all medication should ideally be given together in a single daily dose, 30 minutes before breakfast. Standard short course therapy usually renders a patient non-infectious within 2 weeks of starting therapy and produces a cure in approximately 98% of cases. The usual cause of treatment failure is poor adherence and in such cases directly observed therapy should be given or treatment regimen changed, and TB specialist contacted. Resistance to current anti-TB drugs has not been reported in Fiji.

#### 19.2 Treatment of Extra-pulmonary TB

This is more difficult to diagnose than pulmonary disease, often requiring invasive procedures to obtain diagnostic specimens. Patients with this form of disease are not infectious unless they also have pulmonary involvement.

Therapeutic regimen is the same as for pulmonary TB but of longer duration and is often combined with surgery.

Corticosteroids are essential in the treatment of tuberculous pericarditis and may be of benefit in the first few weeks of treatment of tuberculosis meningitis.

#### 19.3 Treatment Regimens in Special Situations

Pregnancy and Breastfeeding
Most anti-TB drugs are safe for use in pregnancy except for
streptomycin, which can cause ototoxicity in the foetus. All anti-TB drugs are compatible with breastfeeding which should not be stopped during treatment. The baby should be given prophylaxis with isoniazid for at least 3 months beyond the time the mother is considered non-infectious. BCG vaccination of the newborn should be postponed until the end of isoniazid prophylaxis.

#### Oral Contraceptives 19.3.2

Rifampicin interacts with oral contraceptives and another form of contraception is advisable. Rifampicin interacts with other drugs as well, see under chapter 1.

#### 19.3.3 <u>Liver Disorders</u>

Liver function should be checked before commencement of therapy. In the presence of pre-existing liver disease or if aspartate transaminase (AST)/ alanine transaminase (ALT) levels are more than twice the normal, regular monitoring of liver function test is essential. If patient develops jaundice or AST and ALT levels rise above 5 times the normal, anti-TB drugs should be stopped. However, Streptomycin and Ethambutol can be continued.

#### Renal Failure

Isoniazid, Rifampicin and Pyrazinamide are safe. Streptomycin and Ethambutol usage requires dose adjustment and monitoring of renal



function.

#### 19.3.5 HIV Infection

Standard short course therapy may be prolonged to 9 months in HIV infected individuals. Adverse drug reactions and interactions are common.

## 19.4 Chemoprophylaxis

In cases of latent infection, full-blown TB can develop later. Treatment at this stage will prevent development of this disease later. Preventive treatment should be considered in:

- recent tuberculin converters, especially in contacts of TB cases, children and adults less than 35 years of age with strongly positive tuberculin reaction.
- immunosuppressed patients with evidence of previously untreated inactive TB.
- children born to mothers suffering from TB
- Isoniazid 300mg daily at least for 6 months PLUS
- Pyridoxine 25mg orally once daily

#### 19.5 TB in Children

When a child contracts TB it means there are infectious adults around. The vaccine BCG is given at birth to all babies to protect them from severe manifestation of the disease i.e. TB meningitis, milliary TB.

Diagnosis is dependant on history, chest X-ray findings, and positive mantoux tests. Gastric washing for smear has not been found to be useful in our setting.

## 19.5.1 Pulmonary TB

Initial Phase - combination of 4 drugs

- Rifampicin 10 15mg/ kg (max 600 mg) orally once daily X 2 months
- Isoniazid 10mg/ kg (max 300 mg) orally once daily X 2 months Isoniazid 10mg/ kg (max 300 mg) orally once daily X 2 months Pyrazinamide 20 35mg / kg (max 3g) orally once daily X 2 months Streptomycin 20 30mg / kg (max 1g) IM daily X 2 months

#### Continuation Phase - 2 drugs combination

- Isoniazid doses as above X 4 6 months
- Rifampicin doses as above X 4 6 months





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#### TREATMENT OF TUBERCULOSIS

19.5.2 TB Meningitis

Initial Phase as above together with Prednisone 2mg/ kg/ day X 4 weeks

Continuation Phase – 2 drugs as above X 9 months

Prophylaxis is indicated for the following:

-Koch's reaction following BCG

-Baby exposed to infectious mother

-Recrudescense in immunocompromised patient

- Isoniazid 10mg/ kg (max 300 mg) daily orally X 6 12 months





# OPPORTUNISTIC INFECTIONS IN HIV 20. OPPORTUNISTIC INFECTIONS IN HIV

#### 20.1 Pneumocystis carinii Pneumonia

Pneumocystis carinii pneumonia is the commonest pulmonary infection in AIDS.

Presentation is often with indolent onset of fever, non-productive cough, progressive dyspnea and bilateral crackles

The gold standard for therapy at present is with Co-trimoxazole (trimethoprim 80 mg and Sulphamethoxazole 400mg), which is effective in approximately 90% of patients. However the incidence of side effects is higher.

 Co-Trimoxazole (Timethoprim component 15-20 mg/kg per day orally / IV in 2-4 divided doses) X 21 days

Prophylaxis (after first episode of infection):

 Co-trimoxazole (Trimethoprim 160mg / Sulphamethoxazole 800 mg) orally daily or two times daily for 2 days of a week

#### 20.2 CEREBRAL TOXOPLASMOSIS

Toxoplasmosis is the most common cause of secondary CNS infection in patients with AIDS. The standard treatment is combination therapy with Pyrimethamine\* and Sulphadiazine\*.

## (a) Primary Therapy

- Sulphadiazine\* 1 1.5g orally / IV 6 hourly PLUS
- Pyrimethamine\* 200mg orally initially then 50 –100 mg orally daily,

## with or without

Folinic Acid\* 7.5mg orally daily.

Calcium Folinate is usually added to reduce bone marrow suppression, and hence the white cell and platelet count must be monitored closely.

In patients hypersensitive to Sulphonamides substitute sulphadiazine with

Clindamycin\* 600mg orally/ IV 6 hourly

Duration of therapy is for 3-6 weeks depending on clinical response.

\* NOT ON EDL







## (b) Prophylaxis

Relapse is common, so maintenance therapy is necessary while the patient is immunosuppressed;

- Co-trimoxazole (Trimethoprim 160mg / Sulphamethoxazole 800 mg) orally once daily PLUS
- Pyrimethamine\* 500mg orally daily with or without
- Folinic Acid\* 7.5mg orally once daily

#### OR

- Clindamycin\* 600mg orally 8 hourly PLUS
- Pyrimethamine\* AND Folinic Acid\* as above.

#### 20.3 CRYPTOCOCCAL MENINGITIS

Therapy should be initiated immediately in any patient with evidence of cryptococcal infection. The diagnosis depends upon the presence of budding yeasts in the CSF and detection of cryptococcal antigen in the blood and CSF.

#### (a) Initial Therapy:

 Amphotericin B 0.3 – 0.5 mg/ kg/ day IV once daily, increasing to 1mg/ kg/day. The duration of treatment is achieved when total cumulative dose reaches 3g

#### with or without

• 5-flucytosine\* 100 mg/ kg/ day IV or orally 6 hourly X 14 days

## OR (Mild Disease)

Fluconazole 200 – 400mg /day IV or orally once daily X 8 –10 weeks (as a single agent)

#### (b) Maintenance Therapy

Fluconazole 200mg / day

Primary prophylaxis is not recommended

## 20.4 HERPES SIMPLEX

Therapy (primary infection, 1st episode):

 Acyclovir\* 200 mg – 400mg orally 5 times daily X 5 – 10 days OR until clinical resolution.

\* NOT ON EDL







#### OPPORTUNISTIC INFECTIONS IN HIV

#### Severe Disease

Acyclovir 5-10mg/kg IV every 8 hours X 7-10 days OR until clinical resolution (dose and frequency reduced in renal failure)

Long-term suppressive therapy for frequent recurrent episodes

Acyclovir\* 200mg orally three times daily

#### 20.5 HERPES ZOSTER

## Therapy

Acyclovir\* 800 mg orally 5 times per day at least for 7 days (until lesions crust)

#### Disseminated

Acyclovir 30-36 mg/kg/day IV at least 7 days

## 20.6 FUNGAL SKIN INFECTIONS

#### 20.6.1 Superficial Skin Infection

- Whitfield's ointment applied two times daily X 2 weeks
- Ketoconazole\* cream or ointment

#### Extensive disease

- Griseofulvin 250-500mg orally once daily **OR** Ketoconazole 200mg orally once daily

Duration of therapy is usually 1 – 3 months

#### 20.6.2 Candidiasis

#### Oropharyngeal (Oral Thrush)

Nystatin 500,000 units gargle 4 times per day until symptoms resolve (10 – 14 days)

## Vaginitis

Econazole intravaginal suppository 200mg X 3 days

#### Oesophagitis

Fluconazole 200 – 400mg orally X 2 – 3 weeks

Consider maintenance therapy with Fluconazole 100 – 200mg orally for recurrent oesophagitis.

\* NOT ON EDL









OPPORTUNISTIC INFECTIONS IN HIV

#### 20.7 HIV RELATED DIARRHOEA

Salmonella / Shigella Enterocolitis (may cause bacteremia or focal extraintestinal infection)

Ciprofloxacin 500mg orally 12 hourly X 14 days

# Campylobacter colitis (this may cause bacteremia) • Erythromycin 500mg orally 6 hourly X 2 weeks

Cryptosporodiasis
Paramomycin\* 500mg orally every 8 hours X 2 weeks

## TREATMENT OF *Mycobacterium tuberculosis*Refer to chapter 19 under Tuberculosis. 20.8





\* NOT ON EDL





# TREATMENT OF ANAPHYLAXIS 21. TREATMENT OF ANAPHYLAXIS

Allergic reactions may be triggered by a variety of factors including drugs (eg Penicillin), foods (eg shellfish), insect stings and chemicals. There is a wide spectrum of severity ranging from a harmless skin rash (urticaria), to potentially fatal airway obstruction (laryngeal oedema) and full blown anaphylaxis (hypotension, bronchospasm). Anaphylaxis is mich more common in adults than children.

The mainstays of treatment are oxygen, adrenaline and intravenous fluid. Steroids may prevent relapse and antihistamine provide some relief of uticaria itch but these drugs do notthing for the life threatening features of acute severe anaphylaxis.

#### In ADULTS:

#### 1. Airway and Breathing

Administer high flow oxygen via face mask. Administer a bronchodilator. Give Salbutamol  $5mg\ via\ nebulizer.$ 

#### 2. Adrenaline

If severe (hypotension or sever bronchospasm or stridor or hypoxia):

Give adrenaline 0.5mg intramuscularly and repeat in 5 minutes if required

#### OR

If less severe (systolic blood pressure greater than 90mmHg, mild bronchospasm, no stridor and no hypoxia):

Give adrenaline 100mcg intravenously each minute until symptoms subside

#### 3. Intravenous Fluids

Give 0.9% saline 250ml intravenous bolus and repeat if necessary.

NOTE: Large volumes of intravenous fluid may be necessary to maintain adequate blood pressure in severe anaphylaxis.

#### 4. Corticosteroids

Give hydrocortisone succinate 200mg intravenously THEN 100mg six hourly

OR

Give dexamethasone 8mg intramuscularly

OR

Give prednisone 50mg orally daily

NOTE All patients with significant anaphylaxis should be observed for 24 hours as relapse may occur.





#### 5. Anti-histamines

Give promethazine 25mg intramuscularly followed by either 25mg intramuscularly three times daily or 20mg orally

#### 6. Other Issues

- 1. Proper documentation of all cases.
- 2. Patient and relatives should be educated to avoid subsequent episodes.
- 3. Patient may be provided with a medic alert bracelet.
- 4. Some patients may require adrenaline syringes for home use.

#### In CHILDREN:

#### 1. Airway and Breathing

Adminster high flow oxygen via a face mask. Bronchodilators reduce bronchospasm.

Give salbutamol 2.5mg via nebuliser in children 5 years of age or under, for children older than 5 years give salbutamol 5mg via nebuliser

#### 2. Adrenaline

Give adrenaline 10mcg/kg IV over 1 minute and repeat in 5 minutes if required.

NOTE: If intravenous acess is not available then adrenaline may be given via the intramuscular route:

Give adrenaline 10mcg/kg intramuscularly

## 3. Intravenous Fluids

Give 0.9% saline 10ml/kg bolus intravenously and repeat as necessary

## 4. Corticosteroids

Give Hydrocortisone succinate 4mg / kg intravenously

OR

Give Dexamethasone 0.2mg/ kg intramuscularly

OR

Give Prednisolone 1mg/kg orally

 $\ensuremath{\mathsf{NOTE}}\xspace$  : All patients with significant anaphylaxis should be observed for 24 hours as relapse may occur.

#### 5. Anti-histamines

Consider use of promethazine

#### 6. Other Issues

- 1. Proper documentation of all cases.
- 2. Patient and relatives should be educated to avoid subsequent episodes.
- 3. Patient may be provided with a medic alert bracelet.
- 4. Some patients may require adrenaline syringes for home use





# ANTIBIOTIC USE DURING PREGNANCY AND BREASTFEEDING 22. ANTIBIOTIC USE DURING PREGNANCY AND BREASTFEEDING

#### PREGNANCY

The nature of adverse effects of drug use during pregnancy depends upon the time of exposure, teratogenicity being a major risk to drug exposure during the first trimester, while in second and third trimesters fetal growth and functional development may be affected.

Categorization of drugs in pregnancy\* By Australian Drug Evaluation Committee (ADEC)

#### Category A

Drugs which have been taken by a large number of pregnant women and women of childbearing age without any proven increase in the frequency of malformations or other direct or indirect harmful effects on the fetus having being observed.

#### Category B1

Drugs which have been only taken by a limited number of pregnant women and women of child bearing age, without an increase in the frequency of malformation or other direct or indirect harmful effects on human fetus having been observed. Studies in animals have not shown evidence of an increased occurrence of fetal damage.

#### Category B2

Drugs which have been taken by only a limited number of pregnant women and women of childbearing age, without an increase in the frequency of malformation or other direct or indirect harmful effects on the human fetus having been observed. Studies in animals are inadequate or maybe lacking, but available data show no evidence of an increased occurrence of fetal damage.

#### Category B3

Drugs which have been taken by only a limited number of pregnant women and women of childbearing age, without an increase in the frequency of malformation or other direct or indirect harmful effects on the human fetus having been observed. Studies in animals have shown evidence of an increased occurrence of fetal damage, the significance of which is considered uncertain in humans.

#### Category C

Drugs, which owing to their pharmacological effects, have caused or maybe suspected of causing, harmful effects on the human fetus or neonate without causing malformations. These effects maybe reversible. Accompanying texts should be consulted for further details.







#### Category D

Drugs, which have caused, are suspected to have caused or maybe expected to cause, an increased incidence on human fetal malformations or irreversible damage. These drugs may also have adverse pharmacological effects. Accompanying texts should be consulted for further details.

ANTIBIOTIC USE DURING PREGNANCY AND BREASTFEEDING

Category X

Drugs which have such a high risk of causing permanent damage to the fetus that they should not be used in pregnancy or when there is a possibility of pregnancy.

#### BREASTFEEDING

There are two important issues to consider when prescribing drugs/ antibiotics during breast-feeding; firstly the likely exposure of the drug to the infant (who is an innocent bystander) and secondly the likely effect the drug may have on milk supply. A risk benefit analysis is warranted.

Simple advice such as feeding the infant just before the next dose or alternatively taking the medication just after breastfeeding thus avoiding likely peak milk concentrations can be given.

Antimicrobial	Use in breastfeeding	Use in Pregnancy
Acyclovir	Safe to use	B3
Amoxycillin	Safe to use, may cause loose bowel action	
	in infant.	A
Amoxycillin +	Safe to use, may cause loose bowel action	
clavulanic acid	in infant.	B1
Amphotericin	Safe to use	B3
Ampicillin	Safe to use, may cause loose bowel action	
	in infant.	A
Benzathine penicillin	Safe to use, may cause loose bowel action	
	in infant.	A
Benzyl penicillin	Safe to use, may cause loose bowel action	
	in infant.	A
Cefaclor	Safe to use, may cause loose bowel action	
	in infant.	B1
Cefriaxone	Safe to use, may cause loose bowel action	
	in infant.	B1
Cephalothin	Safe to use, may cause loose bowel action	
	in infant.	A
Chloramphenicol	Safe to use, may cause loose bowel action	
	in infant.	A
Chloroquine (prohpylaxis)	Safe to use	Α
Chloroquine (treatment)	Contact specialist, risk benefit ratio in	
	favour of use.	D







#### ANTIBIOTIC USE DURING PREGNANCY AND BREASTFEEDING

Antimicrobial	Use in breastfeeding	Use in Pregnancy
Ciprofloxacin	Use alternatives when possible, short	
	courses maybe acceptable in some	
	circumstances.	B3
Cotrimoxazole	Avoid use in ill, stressed, pre-term infants	C, Contraindicated
	or infants with hyperbilirubinaemia or	in late pregnancy
	G6PD deficiency.	
Dapsone	Neonates with G6PD deficiency are	
	susceptible to dapsone haemolysis. Contact	
	specialist.	B2 <b>D.</b> safe to use during
Doxycyline	Theoretical risk, no case reported. Short	the 1st 18 weeks of
	courses of 7 – 10 days.	pregnancy
Ferments	0-4	
Econazole	Safe to use	A
Erythromycin	Safe to use, may cause loose bowel action	
Ethambutol	in infant. Safe to use	A A
Flucloxacillin	Safe to use Safe to use, may cause loose bowel action	A
Flucioxaciiliri	in infant.	B1
Gentamicin	Safe to use	
Gentamicin	Sale to use	D, reserve for severe or life threatening infections,
		fetal nephrotoxicity and
		ototoxicity have been
		reported.
Griseofulvin	Avoid use	B3
Isoniazid	Safe to use	Α
Ketoconazole	Maybe used, very small amounts excreted	
	in breast milk.	B3
Lamivudine	Discourage breastfeeding, risk of postnatal	
	transmission.	B3
Mebendazole	Maybe used, poorly absorbed by mother	B3
Metronidazole	Safe to use, may cause bitterness in milk.	
	Dose preferably twice daily after breastfeeding.	B2
Nitrofurantoin	Use with caution, may cause haemolysis in	
	G6PD deficiency	A
Nystatin	Safe to use	A
Phenoxymethylpenicillin	Safe to use, may cause loose bowel actions	
	in infant	A
Piperacillin	Safe to use, may cause loose bowel actions	
	in infant	B1
Primaquine	Avoid use	<b>D</b> , avoid use in 3 <sup>rd</sup>
		trimester, may casue
		neonatal hamolysis &
		methaemoglobinaemia
Procaine penicillin	Safe to use, may cause loose bowel actions	
	in infant	Α
Pyrantel	Safe to use	B2, avoid use in 1st trimester.









Antimicrobial	Use in breastfeeding	Use in Pregnancy
Pyrazinamide	Amount too small to be harmful	B2
Quinine		D
Rifampicin	Maybe used	С
Sulphonamides		С
Streptomycin	Safe to use	D
Tetracycline	Theoretical risk, no case reported.	D, safe to use during
	Short courses of 7 – 10 days.	the 1st 18 weeks of
		pregnancy
Trimethoprim	Safe to use	B3
Vancomycin	Safe to use	B2
Zidovudine	Discourage breastfeeding, risk	
	of postnatal transmission.	B3

ANTIBIOTIC USE DURING PREGNANCY AND BREASTFEEDING







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As the final user of this volume, you can give us valuable advice about the contents , layout and usability of this book. Your comments will be highly appreciated and considered during the preparation of  $\,$  the next edition.

Please send comments to :

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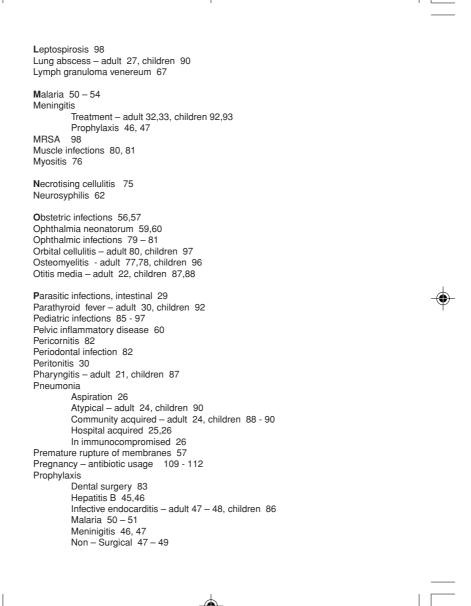
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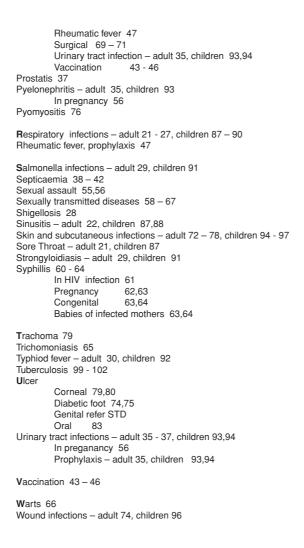




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