
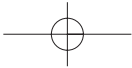
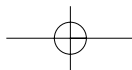
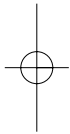


**NATIONAL ANTIBIOTIC
GUIDELINE
2008**





MESSAGE FROM THE DIRECTOR GENERAL OF HEALTH, MALAYSIA

From the 2007 audit on utilisation of 13 antibiotic injections in 15 major hospitals, it was found that the most used antibiotic was the cephalosporin group. Of particular concern was the consistent increase in the use of Cefoperazone-Sulbactam combination by nearly 30% each year for the past 2 consecutive years although we know that this antibiotic should only be reserved for treating multi-resistant organisms. Similarly, the use of 3 other major groups of antibiotics namely the Carbapenems, Quinolones and Vancomycin showed steady increases by 50%, 38% and 30% respectively as compared to 2005. This increase in the trend of use cannot be taken lightly and measures must be taken to ensure that they are prescribed appropriately. In terms of expenditure, it was noted that hospitals spent between 5-15 percent of their annual drug budget on antibiotics alone.

Strategies such as good infection control practices, conduct of multidisciplinary antibiotic rounds, establishment of national antimicrobial guideline, surveillance programmes, audits, continuous training and education amongst health personnel are necessary and vital to promote and ensure the quality use of antibiotics. Inappropriate use of antibiotics as we all know is a major factor contributing to the development of resistance. Information on the trends and pattern of use is essential towards formulating control measures on antibiotic prescribing.

This revised National Antibiotic Guideline, I am sure, will be a useful and important guide for prescribers towards making appropriate antibiotic choices but local sensitivity patterns, particularly in tertiary hospitals, should also be taken into consideration where necessary. If local guidelines are developed, then the Hospital Infection Control and Antibiotic Committee must initiate regular audits to check for any non-compliance and misuse.

I would like to congratulate all specialists including heads of discipline and pharmacists who have contributed to the publication of this guideline. Special thanks also go to the external reviewers for their input and comments. Lastly, I must commend the editorial committee for successfully putting everything together to make it as comprehensive as possible. I am sure this is not an easy task. The next important step is to ensure that all relevant healthcare personnel gain access to this publication for easy reference.

Thank you



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INTRODUCTION TO THE GUIDELINES

Global and National Threat

The World Health Organization (WHO) in its document on Containment of Antimicrobial Resistance urges governments and the medical profession throughout the world to take active and concrete measures to address this threat. The rates of multiresistant organisms have increased significantly and, in a relatively short period of time in many countries. Methicillin Resistant *Staphylococcus aureus* (MRSA) and Extended Spectrum Beta-lactamase (ESBL) producing organisms like *Klebsiella pneumoniae* are now major adversaries in many of our local hospitals especially in the critical care settings. Broad spectrum antibiotics like the carbapenems, which once were very effective for most gram negative organisms are now experiencing up to 20% resistance in *Pseudomonas aeruginosa*.

What is driving Antibiotic Resistance?

The belief that antibiotic use or misuse is a major driving force for antibiotic resistance is now an established and recognised fact. It is thus imperative for all healthcare practitioners to play their role in combating this threat so as to preserve the effectiveness and the relevance of current antibiotics in our practice. Rational antibiotic use must be viewed as a skill that all medical practitioners must acquire so as to ensure effective, safe and appropriate patient care. Appropriate treatment in our current approach is not only about using an antibiotic that the organism is sensitive to but also includes the use of one that will have minimal collateral damage to the ambient bacterial flora.

National Antibiotic Guideline 2008

The last national antibiotic guideline for the Ministry of Health was published in 1997; an which was a collaborative effort with the Academy of Medicine. With new clinical information and challenges over the last decade, it is certainly time for developing a new document to provide guidance in the use of antimicrobials in common infections encountered in the Ministry of Health clinical facilities.

This document is a collaborative effort involving a large number of specialists from within the Ministry of Health; spanning all major clinical disciplines and bringing together the expertise and experience of many senior clinicians from all regions of the country. The recommendations are based on *current clinical evidence* similar to the approach taken in the production of clinical practice guidelines, the *current list of antimicrobials in the ministry drug formulary*, the *pattern of antimicrobial resistance seen in the country* as well as the *current practice within Ministry of Health hospitals*.

Nonetheless because of the large spectrum of clinical infections; some of which involved several disciplines, consensus decision-making involving the relevant stakeholders was pursued whenever differences of opinion occurred. While the editorial committee aimed to address all common infections in the numerous clinical settings within the ministry, they also took due cognizance of the need to keep the document concise for the purpose of producing a pocket handbook. Hence, the editorial committee decided to include only the more common and critical infections for mention. Less common infections and those seen only in specialised areas, regrettably, had to be omitted. Most portions of the document are formatted in a standardised manner so as to provide uniformity and to make it more reader friendly.

NATIONAL ANTIBIOTIC GUIDELINE 2008

Antibiotic choices are classified into preferred and alternative recommendations based on clinical evidence of effectiveness, adverse effects, potential of collateral damage as well as cost and access. References have been inserted whenever possible.

This document aims to guide clinicians in their empirical choice of antimicrobial agents; balancing the need to get the right choice from the outset and the necessity to contain antimicrobial misuse so as to preserve future treatment options especially in the current era of growing antimicrobial resistance. Nonetheless, this document merely acts as a guide and each case must still be accessed according to its own merits.

Appreciation

On behalf of the editorial committee and the secretariat, I would like to thank the numerous contributors from all clinical disciplines, all heads of discipline, infectious diseases specialists, microbiologists and pharmacists who have directly or indirectly assisted in this document. I would also like to thank our external reviewers for their invaluable input. Their commitment and patience in this endeavor is much appreciated. We would also like to convey our gratitude to Tan Sri Datuk Dr Hj. Mohd Ismail Merican, the Director-General of Health for all his support and advice.

Dr Christopher K.C. Lee
Chairman
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Ministry of Health
14th December 2007

PRINCIPLES OF ANTIBIOTIC THERAPY AND RATIONAL ANTIBIOTIC PRESCRIBING

Infections remain a common cause of presentation to the outpatient department and inpatient admissions to the hospital. Antibiotics are widely being prescribed to treat infections, both in the community and hospital setting. Selection of appropriate anti-infective therapy can be challenging to the clinician. Consequently, understanding the basic principles of anti-infective therapy is important to ensure optimal outcome and to reduce selective pressure on antibiotics, which may be associated with the development of antibiotic resistance. The overuse and misuse of antibiotics have contributed to increased bacterial resistance to antibiotics, among other contributory factors. Antibiotics are frequently prescribed for indications in which their use is not warranted, or an inappropriate or suboptimal antibiotic is prescribed. The available evidence suggests that, when antibiotic use is warranted, choosing the therapy most likely to achieve clinical cure and treating for the shortest length of time to achieve clinical and microbiological efficacy will result in a lower incidence of retreatment and lower incidence of antibiotic resistance. The rational use of medicines has been defined by the WHO as requiring that patients receive medications appropriate to their clinical needs, in doses that meet their own requirements, for an adequate time, and at the lowest cost to them and their community.

A thorough clinical assessment of the patient is imperative to ascertain the underlying disease process, and if it is an infection, to predict the pathogens associated with the infection and select an antibiotic that will target the likely organisms. Where appropriate and clinically indicated, the initial assessment should be supported by relevant laboratory investigations to establish a definitive microbiological diagnosis and to determine the susceptibility of the organism to various antibiotics. The routine use of antibiotics to treat fever is inappropriate, as not all fever is caused by infection and antibiotics are only indicated for bacterial infections. Antibiotics should not be prescribed when bacterial infections are unlikely, such as for common cold, coughs and bronchitis, as irrational antibiotic prescribing is documented as one of the main factors that encourage emergence of antibiotic-resistant pathogens.

When choosing an antibiotic for empirical treatment of an infection, the following factors are important to assist and guide the decision making process:

Is there an indication for an antimicrobial agent?

Indications for an antibiotic include the unambiguous demonstration or the strong suspicion that the etiologic agent is bacterial. This should be based on the signs and symptoms of infection, as well as on other factors, including the age of the patient, the patient's medical history, and the presence or absence of comorbidities.

What are the most common organisms causing the infection and the local antibiotic susceptibility pattern?

Knowledge of the likely organisms causing a particular infection and the local susceptibility profile are useful to select the antibiotic. For example, erysipelas is caused primarily by *Streptococcus pyogenes* which is usually sensitive to penicillins and macrolides, while impetigo may be caused by *Streptococcus pyogenes* or *Staphylococcus aureus*, both sensitive to penicillase-resistant penicillins such as cloxacillin.

NATIONAL ANTIBIOTIC GUIDELINE 2008

What is the antibiotic spectrum of the chosen empirical agent?

The antibiotic spectrum refers to the range of microorganisms an antibiotic is usually effective against and is an important consideration for empiric therapy. Decision on choice of antibiotic based on the spectrum of coverage should be made based on severity of illness, pathogen probabilities (whether gram-positive or gram-negative bacteria), local resistance patterns, comorbid conditions and recent antibiotic exposure. The definitive choice of antibiotics should be made after review of culture and susceptibility results and therapy should be tailored accordingly.

What are the known pharmacokinetics and pharmacodynamics that are associated with a particular antibiotic?

Knowledge of the pharmacokinetics and pharmacodynamic principles assist the clinician in predicting the clinical and microbiologic success of antibiotic treatment. Concentration-dependent bacterial killing is a feature of antibiotics such as aminoglycosides and fluoroquinolones, higher concentrations resulting in more rapid killing. Time-dependent bacterial killing is associated with beta-lactam antibiotics, greater degree of bacterial killing occurring when the time of exposure is above the minimal inhibitory concentration of the pathogen.

What host factors might affect antibiotic selection and dosing?

Host factors, such as patient age and underlying disease, are important considerations in selecting appropriate antibiotic therapy for suspected bacterial infections. Host factors influence the types of bacteria likely to be pathogenic and organ failures may impact on dosing regimens and predispose to adverse drug reactions.

What is the cost-effectiveness of the antibiotic selection?

Choosing inappropriate therapy is associated with increased costs, including the cost of the antibiotic and increases in overall costs of medical care because of treatment failures and adverse events. Using an optimal course of antibiotics can have economic as well as clinical advantages, including a faster return to normal daily routine and earlier return to work.

What are the antibiotic adverse reactions?

Antibiotic prescribing may be associated with potential side effects that may affect the relative risks and benefits of therapy. All antibiotics have potential side effects, and it is important for the clinician to be aware of how these might affect the patient.

What is the optimal duration of treatment?

There are very few infections for which the duration of treatment has been precisely defined. This reflects the fact that the end-points for assessing treatment are largely clinical rather than microbiological. Clinical features that are driven by the inflammatory response usually subside after microbial elimination. Clinicians should assess the time frame for discontinuing antibiotics after careful review of the clinical response, guided by microbiological clearance of the pathogen whenever appropriate.

NATIONAL ANTIBIOTIC GUIDELINE 2008

In conclusion, antibiotic prescribing should be made after careful consideration of the underlying infective process, the likely etiologic agents, local susceptibility pattern, known spectrum of a chosen antibiotic, host factors and comorbidities. Rational antibiotic prescribing can minimize development of antibiotic resistance and reduce costs of healthcare.

What is de-escalation therapy and when is it warranted?

De-escalation of antibiotic therapy refers to short-term, broad-spectrum antibiotic coverage followed by changes to more narrow focused regimens that are driven by culture and other laboratory results. This limited use does not expose the patient to the potential adverse effects of untreated serious infections or to the complications associated with long-term broad-spectrum antibiotic use, which are primarily the emergence of resistant organisms or new infections. This approach is particularly pertinent when dealing with life-threatening conditions especially infections in the critical care patients, immunocompromised patients and patients with risk factors for hospital acquired infections; where delay in initiating the appropriate antibiotic therapy may result in mortality. Broad-spectrum initial therapy does not appear to result in the emergence of antibiotic resistance as long as the duration of use was limited. The choice of the initial antibiotic regimen should be based on the local microbiological surveillance data.

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SECTION A: ADULTS



CARDIOVASCULAR INFECTIONS

A. INFECTIVE ENDOCARDITIS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|---|-------------|--|
| | Preferred | Alternative | |
| Empirical Treatment | | | |
| | Benzylpenicillin 24 mega units/24h IV either continuously or in 4-6 equally divided doses PLUS Gentamicin ¹ 3mg/kg IV/IM q24h If there is a strong possibility of staphylococcal infection, e.g. IV drug abuse, infected haemodialysis lines or pacemaker infection: Cloxacillin 12g/24h IV in 4-6 divided doses PLUS Gentamicin ¹ 1mg/kg IM/IV q8h | | Treatment can be modified once the blood result is known |

6

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| Viridans Streptococci & Streptococcus Bovis It is recommended MIC estimation is done for these isolates to facilitate management | | | |
| Native Valves MIC: ≤ 0.12µg/mL <i>Penicillin-Susceptible Viridans Streptococci & Streptococcus Bovis</i> | Benzylpenicillin 12-18 mega units/24h IV either continuously or in 4-6 equally divided doses for 4 weeks | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV/IM q24h for 4 weeks OR Benzylpenicillin 12-18 mega units/24h IV either continuously or in 4-6 equally divided doses for 2 weeks PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 2 weeks OR 3 rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV/IM q24h for 2 weeks PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 2 weeks | 4-weeks regimen preferred for patients > 65 years or patients with impaired renal or 8 th cranial nerve function 2-weeks regimen not intended for patients with <ul style="list-style-type: none"> ◆ known cardiac or extracardiac abscess ◆ creatinine clearance <20ml/min ◆ impaired 8th nerve function |

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|----------|
| | Preferred | Alternative | |
| Native Valves MIC: > 0.12µg/mL- ≤ 0.5µg/mL <i>Penicillin-Relatively Resistant Viridans Streptococci & Streptococcus Bovis</i> | Benzylpenicillin 24 mega units/24h IV either continuously or in 4-6 equally divided doses for 4 weeks PLUS Gentamicin ¹ 3mg/kg IM/IV q24h for 2 weeks | <i>3rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV/IM q24h for 4 weeks</i> PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 2 weeks <u>If unable to tolerate Penicillin/Ceftriaxone:</u> Vancomycin ¹ 15mg/kg IV q12h for 4 weeks, not to exceed 2g/24h (unless serum levels are monitored) | |
| Native Valves MIC > 0.5µg/mL <i>Penicillin-resistant Viridans Streptococci & Streptococcus Bovis</i> | Treat as enterococcal endocarditis - see below ** | | |
| Prosthetic Valves MIC < 0.12µg/mL <i>Penicillin-Susceptible Viridans Streptococci & Streptococcus Bovis</i> | Benzylpenicillin 24 mega units/24h IV either continuously or in 4-6 equally divided doses for 6 weeks PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 2 weeks | <i>3rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV/IM q24h for 6 weeks</i> PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 2 weeks <u>If unable to tolerate Penicillin/Ceftriaxone:</u> Vancomycin ¹ 15mg/kg IV q12h for 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) | |

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| Prosthetic Valves MIC > 0.12µg/mL <i>Penicillin-relatively resistant or fully resistant Viridans Streptococci & Streptococcus Bovis</i> | Benzylpenicillin 24 mega units/24h IV either continuously or in in 4-6 equally divided doses for 6 weeks PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 6 weeks | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV/IM q24h for 6 weeks PLUS Gentamicin ¹ 3mg/kg IV/IM q24h for 6 weeks If unable to tolerate Penicillin/ Ceftriaxone: Vancomycin ¹ 15mg/kg IV q12h for 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) | |
| ** Enterococcus (<i>It is recommended that all these isolates are tested for high level resistance (HLR) to Gentamicin</i>) | | | |
| Native and Prosthetic Valves <i>Enterococcal Endocarditis sensitive to Gentamicin</i> | Ampicillin 2g IV q4h for 4-6 weeks PLUS *Gentamicin ¹ 1mg/kg IM/IV q8h for 4-6 weeks | Benzylpenicillin 18-30 mega units/24h IV in 4-6 equally divided doses for 4-6 weeks PLUS *Gentamicin ¹ 1mg/kg IM/IV q8h for 4-6 weeks If unable to tolerate Penicillin: Vancomycin ¹ 15mg/kg IV q12h for 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) PLUS Gentamicin ¹ 1mg/kg IM/IV q8h for 6 weeks | Native valve: Symptoms < 3 months - 4 weeks therapy Symptoms > 3 months - 6 weeks therapy Prosthetic valve: minimum 6 weeks *In order to maximise synergistic effect, administer Gentamicin at the same time or temporally close to Ampicillin/Penicillin For Enterococcal Endocarditis with high level resistance to Gentamicin, consult Infectious Disease Specialist |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| Staphylococcus Aureus | | | |
| Native Valves <i>Methicillin-Susceptible Staphylococci</i> | Left sided endocarditis and complicated right sided (see comments): Cloxacillin 12g/24h IV in 4-6 divided doses for 6 weeks PLUS/MINUS Gentamicin ¹ 1mg/kg IV/IM q8h for 3-5 days Right sided endocarditis (tricuspid valve) in uncomplicated endocarditis (see comments): Cloxacillin 12g/24h IV in 4-6 divided doses for 2 weeks PLUS Gentamicin ¹ 1mg/kg IM/IV q8h for 2 weeks | Regimen for β-lactam allergic patients: <u>Immediate type hypersensitivity to penicillin (anaphylaxis):</u> Vancomycin ¹ 15mg/kg IV q12h for 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) <u>For non-immediate type hypersensitivity:</u> * Cefazolin 2g IV q8h for 6 weeks PLUS/MINUS Gentamicin ¹ 1mg/kg IM/IV q8h for 3-5 days | Uncomplicated right sided endocarditis: Absence of renal failure, extra pulmonary metastatic infections such as osteomyelitis, aortic or mitral valve involvement, meningitis, or infection by MRSA * If Cefazolin is not available, use of Cefuroxime may be considered |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Prosthetic Valves <i>Methicillin-Susceptible Staphylococci</i> | Cloxacillin 12g/24h IV in 4-6 divided doses for \geq 6 weeks PLUS Rifampicin ² 300mg PO q8h for \geq 6 weeks PLUS Gentamicin ¹ 1mg/kg IM/IV q8h for 2 weeks | Regimen for β -lactam allergic patients: <u>Immediate type hypersensitivity to Penicillin (anaphylaxis):</u> Vancomycin ¹ 15mg/kg IV q12h for \geq 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) PLUS Rifampicin ² 300mg PO q8h for \geq 6 weeks PLUS Gentamicin ¹ 1mg/kg IM/IV q8h for 2 weeks <u>For non-immediate type hypersensitivity:</u> *Cefazolin 2g IV q8h for 6 weeks PLUS Rifampicin ² 300mg PO q8h for \geq 6 weeks PLUS Gentamicin ¹ 1mg/kg IM/IV q8h for 2 weeks | *If Cefazolin is not available, use of Cefuroxime may be considered |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|----------|
| | Preferred | Alternative | |
| Native Valves Methicillin-Resistant Staphylococci | Vancomycin ¹ 15mg/kg IV q12h for 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) | | |
| Prosthetic Valves MRSA | Vancomycin ¹ 15mg/kg IV q12h for ≥ 6 weeks, not to exceed 2g/24h (unless serum levels are monitored) PLUS Rifampicin ² 300mg PO q8h for ≥ 6 weeks PLUS Gentamicin ¹ 1mg/kg IM/IV q8h for 2 weeks | | |
| HACEK Microorganisms (<i>Haemophilus parainfluenzae</i> , <i>Haemophilus aphrophilus</i> , <i>Actinobacillus actinomycetemcomitans</i> , <i>Cardiobacterium hominis</i> , <i>Eikenella corrodens</i> , and <i>Kingella kingae</i>) | | | |
| Native and Prosthetic valves | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV/IM q24h for 4 weeks | β -lactam/ β -lactamase inhibitors, e.g. Ampicillin/Sulbactam 3g IV q6h for 4 weeks | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| Therapy for Culture-Negative Endocarditis - Consultation with an infectious disease specialist needed | | | |
| Native Valves | Ampicillin/Sulbactam 3g IV q6h for 4-6 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 4-6 weeks | Vancomycin ¹ 15mg/kg IV q12h for 4-6 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 4-6 weeks PLUS Ciprofloxacin 500mg PO q12h OR 400mg IV q12h for 4-6 weeks | Vancomycin recommended only for patients unable to tolerate penicillins |
| Prosthetic valve (early, <1 y) | Vancomycin ¹ 15mg/kg IV q12h for 6 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks PLUS Cefepime 2g IV q8h for 6 weeks PLUS Rifampicin 300mg PO/IV q8h for 6 weeks | | |
| Prosthetic valve (late, >1 y) | Ampicillin/Sulbactam 3g IV q6h for 4-6 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 4-6 weeks PLUS Rifampicin 300mg PO/IV q8h for 6 weeks | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|---|
| | Preferred | Alternative | |
| Suspected Bartonella, culture negative | Ceftriaxone 2g IV/IM q24h for 6 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks OR Doxycycline 100mg IV/PO q12h for 6 weeks | | Patients with Bartonella endocarditis should be treated in consultation with an infectious disease specialist |
| Documented Bartonella, culture positive | Doxycycline 100mg IV/PO q12h PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks | If Gentamicin cannot be given, then replace with Rifampicin 600mg PO/IV q24h in 2 equally divided doses | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

²Rifampicin plays a unique role in the eradication of staphylococcal infection involving prosthetic material, combination therapy is essential to prevent emergence of rifampicin resistance

B. TREATMENT OF PACEMAKER INFECTIONS

| Antibiotic | Duration | Comments |
|--|-------------------------------------|--|
| <p>While awaiting microbiological diagnosis provide empirical cover for MRSA with:</p> <p>Vancomycin 15mg/kg IV q12h not to exceed 2g/24h (unless serum levels are monitored)</p> <ul style="list-style-type: none"> • Infection of pulse generator pocket with blood stream infection • Lead associated endocarditis <p>Change antibiotics according to culture results</p> | <p>10 to 14 days</p> <p>6 weeks</p> | <p><i>Complete removal of the entire implanted system including the cardiac leads is recommended even in patients with clinical infection of the pocket only</i></p> <p><i>The new implant can be placed on the contra lateral side 10 to 14 days after the removal of the implanted system in patients with infection of the pulse generator pocket and as late as 6 weeks in those with endocarditis</i></p> |

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Reference: American Heart Association Guideline 2005

CENTRAL NERVOUS INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Meningitis (acute) | | | |
| <p>Common organisms: <i>Streptococcus pneumoniae</i> <i>Neisseria meningitidis</i> <i>Haemophilus influenzae</i></p> <p>Other organisms: Gram negative rods Leptospirosis Scrub typhus Melioidosis <i>Mycoplasma pneumoniae</i></p> | <p>Empirical treatment on admission: Benzylpenicillin 4 mega units IV q4-6h</p> <p>PLUS <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 50-100mg/kg/24h IV in 2 divided doses (max: 4g/day). Usual dose is 2g q12h</p> <p>OR Cefotaxime 200mg/kg/24h IV in 3 divided doses (max: 12g/day). Usual dose is 2g q8h</p> | <p>Meropenem 120mg/kg/24h IV in 3 divided doses (max: 6g/day) Usual dose is 0.5-1.0g q8h</p> <p>Change to Meropenem if patient showed no clinical response after 3 days of antibiotics</p> <p>IV Dexamethasone in a dose of 0.15mg/kg (10mg) q6h is recommended to be administered 15 to 20 minutes before or at the time of first dose of antibiotics, for up to 4 days or until there is no evidence of pneumococcal meningitis</p> | <p>Antibiotic treatment must be started immediately, regardless of any investigations undertaken. If no organism isolated and patient is responding, continue antibiotics for 7-10 days</p> <p>Meropenem has slightly increased activity against gram negative organisms and slightly decreased activity against staphylococci and streptococci compared to imipenem</p> <p><u>Reference:</u> - Harrison's principles of Internal Medicine, 18th. Edition - de Gans J, van de Beek D. Dexamethasone in adults with bacterial meningitis. N Engl J Med 2002; 347:1549-1556</p> |

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Causative organism isolated: | | | |
| <i>Haemophilus influenzae</i> (Gram -ve bacilli) | <p>3rd gen. Cephalosporins, e.g. Ceftriaxone 50-100mg/kg/24h IV in 2 divided doses (max: 4g/day). Usual dose is 2g q12h</p> <p>OR</p> <p>Cefotaxime 200mg/kg/24h IV in 3 divided doses (max: 12g/day). Usual dose in 2g q8h</p> <p>Duration of treatment: 7-10 days</p> | <p>Meropenem 120mg/kg/24h IV in 3 divided doses (max: 6g/day). Usual dose is 0.5-1g q8h</p> <p>If organism is susceptible:</p> <p>Chloramphenicol 1g IV q6h for 14 days (max: 4g/day)</p> | Increasing primary resistance of <i>Haemophilus influenzae</i> to Chloramphenicol and Ampicillin - in HKL 7.7% and 23.1% respectively |
| <i>Streptococcus pneumoniae</i> (Gram +ve cocci) | <p>Penicillin-sensitive strains</p> <p>Benzylpenicillin 4 mega units IV q4-6h for 10-14 days</p> <p>Relatively-resistant strains</p> <p>3rd gen. Cephalosporins, e.g. Ceftriaxone IV OR Cefotaxime IV for 10-14 days, at doses for <i>H. influenzae</i></p> <p>Duration of treatment: 10-14 days</p> <p>Very ill patients may require treatment for 21 days</p> | <p>Vancomycin¹ 1g IV q12h</p> <p>PLUS</p> <p>3rd gen. Cephalosporins, e.g. Ceftriaxone IV or Cefotaxime IV</p> <p>(For penicillin and cephalosporins resistant strains)</p> | Resistance to penicillin in community acquired <i>Streptococcus pneumoniae</i> in HKL is 16.9% |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|--|
| | Preferred | Alternative | |
| <i>Neisseria meningitidis</i> (Gram -ve cocci) | Benzylicillin 4 mega units IV q4-6h for 7-10 days | 3 rd gen. Cephalosporins, e.g. Ceftriaxone IV OR Cefotaxime IV at doses for <i>H. influenzae</i> | For patients who do not have adequate response to penicillin, the treatment should be changed to 3rd gen. Cephalosporins, e.g. Ceftriaxone OR Cefotaxime |
| Prophylaxis for household and close contacts | Rifampicin 600mg PO q12h for 2 days (4 doses) [not recommended in pregnant women] OR Ciprofloxacin 500mg PO as single dose | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 250mg IM as single dose (especially in pregnancy) OR Azithromycin 500mg PO as single dose | Close contacts are defined as those individuals who have had contact with oropharyngeal secretions either through kissing or by sharing toys, beverages, or cigarettes |
| Viral encephalitis <i>Herpes simplex</i> | Acyclovir 5mg/kg IV q8h for 10-14 days | | |
| <i>Herpes zoster</i> | Acyclovir 10mg/kg IV q8h for 10-14 days | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| Meningitis (Chronic) | | | |
| Tuberculous meningitis <i>Mycobacterium tuberculosis</i> | Intensive 2 months treatment: Isoniazid 5-10mg/kg/24h PO [300mg] PLUS Pyridoxine 20-60mg PO q24h PLUS Rifampicin 10mg/kg/24h PO [600mg] PLUS Pyrazinamide 15-30mg/kg/24h PO [1.5-2g] PLUS Streptomycin 15-20mg/kg/24h IM [0.75-1g] OR Ethambutol 15-20mg/kg/24h PO [800mg] Refer to Page 143 (Tuberculosis Infections) Infection in HIV patients - refer to Page 53 (Human Immunodeficiency Virus) | Refer to Page 143 (Tuberculosis Infections) for management of tuberculosis for drug resistant tuberculosis | Treatment is continued for 12 months Medium dose steroid cover for MRC stage 2 and 3 patients: <i>Dexamethasone 4mg q8h for 2 weeks and then taper down within 4 weeks, or oral prednisolone 30-40mg/24h in tapering doses for 4-6 weeks</i> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|---|
| | Preferred | Alternative | |
| Cryptococcal Meningitis <i>Cryptococcus neoformans</i> | Amphotericin B 0.3-0.6mg/kg/24h IV until total dose of at least 1-1.5g PLUS Fluconazole 400mg PO q24h for 10-12 weeks <u>For fulminant cases:</u> 1st month - Amphotericin B at 0.3-0.6mg/kg/24h IV PLUS 5-Flucytosine 100-150mg/kg/24h IV/PO in 4 divided doses Followed by 2 months of Amphotericin B IV [same dose] + Fluconazole 400mg PO q24h <i>Infection in HIV patients - Refer to Page 53 (Human Immunodeficiency Virus)</i> | Fluconazole 400mg IV q24h initially and then 200-400mg IV q24h for 6-8 weeks Fluconazole "consolidation" therapy may be continued for as long as 6-12 months, depending on the clinical status of the patient If fluconazole is not tolerated: Itraconazole 200mg PO q12h | End point of treatment: till at least 1.5-2.0g of Amphotericin B given and CSF shows clearance of fungus by 2 negative C&S one month apart, and CSF Cryptococcal antigen titre becomes negative or at least 1:2 or shows a fourfold decrease Liposomal Amphotericin may be used in cases of severe toxicity to Amphotericin B e.g. *Abelcet 3-5mg/kg/day *Requires DG approval <u>Reference:</u> Infect Med 1998; 15(6): 396-409 |
| <i>Neurosyphilis</i> | Refer to Page 100 (Sexually Transmitted Infections) | | |
| HIV related CNS infection | Refer to Page 53 (Human Immunodeficiency Virus) | | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

Reference: Use of Antibiotics in Adults: CPG Guidelines. Ministry of Health, Singapore, 2006
 IDSA Practice Guidelines for Management of Cryptococcal Disease, CID 2000; 30:710-718

CHEMOPROPHYLAXIS

A. Surgical Chemoprophylaxis

It is the use of antibiotics to prevent infections at the surgical site. It should be considered when there is significant risk of post-operative infection or where post-operative infection would have severe consequences. Ideally the prophylaxis when given intravenously should be given as soon as the patient is stabilised after induction. Usually a single dose is sufficient. A second dose may be required in the following situations:

- a. delay in start of surgery
- b. in prolonged operations when the time is more than half of the usual dosing interval of the antibiotic

Giving more than 1 or 2 doses postoperatively is generally not advised. The practice of continuing prophylactic antibiotics until surgical drains have been removed is not RECOMMENDED

NATIONAL ANTI-BIOTIC GUIDELINE 2008

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| 1. OBSTETRICS | | | |
| C-Section a. Elective b. Emergency | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV to be given 10 minutes before the first incision | <i>2nd or 3rd gen. Cephalosporins, e.g.</i> Cefuroxime 1.5g IV OR Cefoperazone 1g IV In complicated LSCS (with bowel &/or bladder involvement or possibility of chorioamnionitis): ADD Metronidazole 500mg IV | RCOG Guidelines Antibiotics should be given for at least 5-7 days duration |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Peri/Postpartum Hysterectomy | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV</i> | <i>2nd or 3rd gen. Cephalosporins, e.g. Cefuroxime 1.5g IV</i> OR Cefoperazone 1g IV PLUS Metronidazole 500mg IV | Antibiotics should be given for 5-7 days |
| Repair of Vaginal/Birth tract trauma e.g. third and fourth degree tears | <i>2nd or 3rd gen. Cephalosporins, e.g. Cefuroxime 1.5g IV</i> OR Cefoperazone 1g IV PLUS Metronidazole 500mg IV Antibiotics should be given for at least 5-7 days duration | | RCOG Guideline |
| 2. GYNAECOLOGY | | | |
| Elective Surgery - TAH/TAHBSO - Vaginal hysterectomy <i>Coliforms, Enterococcus, Streptococcus, Clostridia and Bacteroides sp</i> | Cefuroxime 1.5g IV | <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5-3g IV 30-45 minutes before induction</i> | Second dose if procedure > 3 hours |
| Emergency Laparotomy | Cefuroxime 1.5g IV | <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g</i> OR Amoxicillin/Clavulanate 1.2g | ACOG Recommendations: If bowel or bladder perforation occurs add Metronidazole |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| 3. ORAL SURGERY | | | |
| Indication: | | | |
| Elective Minor Oral Surgery | Not Indicated | | Prophylaxis is recommended for all patients with an increased risk of surgical wound infection - i.e. in immunocompromised patients |
| Elective Major Oral Surgery | Indicated | | |
| Which Antibiotic / Route of Administration / Dose / Timing / Duration | | | |
| | <p>* Benzylpenicillin IV 1st Dose: 2 mega units IV (just before procedure) Subsequent Doses: 1 mega unit IV q3h (do not extend beyond surgery)</p> <p>PLUS ** Cloxacillin IV (if surgery involves skin) 1st Dose: 1g PO/IV Subsequent Doses: 500mg PO/IV (do not extend beyond surgery)</p> <p>If Penicillin Contraindicated *** Clindamycin IV 1st Dose*: 300mg IV (just before procedure) Subsequent Doses: 150mg IV q3h (do not extend beyond surgery)</p> | <p><i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate IV 1st Dose: 1.2g IV (just before procedure) Subsequent Doses: 0.6g IV q4h (do not extend beyond surgery)</p> <p>OR Cefuroxime IV 1st Dose: 1.5g (just before procedure) Subsequent Doses: 750mg IV q4h (do not extend beyond surgery)</p> <p>OR <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone IV (if all other above antibiotics contraindicated) 1g just before procedure (do not extend beyond surgery)</p> | <p>*Benzylpenicillin IV should be given by slow intravenous injection or by infusion</p> <p>**Cloxacillin IV should be given by slow intravenous injection or by infusion</p> <p>***Clindamycin IV should be given in 50ml of diluent over 10 min</p> |
| <ul style="list-style-type: none"> • Doses listed are adult doses - for paediatric patients adjust according to age/body weight • References from KKM CPG: Antibiotic Prophylaxis against Wound Infections for Oral Surgical Procedures 2003 (Reviewed 2007) | | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| 4. PLASTIC SURGERY | | | |
| Lip repair, Palatoplasty/ Pharyngoplasty | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV | Erythromycin 500mg IV | Skin, oral and nasal pathogen |
| Craniofacial surgery Maxillofacial surgery | Metronidazole 500mg IV PLUS <i>2nd or 3rd gen. Cephalosporins, e.g.</i> Cefuroxime 1.5g IV OR Ceftriaxone 2g IV (if craniotomy required) | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV | Skin, oral and nasal pathogen Prophylaxis against meningitis/encephalitis |
| Head and neck tumour | Metronidazole 500mg IV PLUS Cefuroxime 1.5g IV | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV | Skin, oral and nasal pathogen |
| Facial injuries | Cloxacillin 500mg-1g IV | Cefuroxime 1.5g IV <i>β-lactam/β-lactamase inhibitors, e.g.</i> | Gross contamination Skin pathogen |
| Breast surgery reconstructive | Cefuroxime 1.5g IV | Ampicillin/Sulbactam 1.5g IV | Skin pathogen |
| Hand replantation | Cefuroxime 1.5g IV | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV | Gross contamination Skin pathogen Prophylaxis against tenosynovitis |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--------------------|--|
| | Preferred | Alternative | |
| 5. VASCULAR SURGERY | | | |
| All Vascular Operations | <p><i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV OR Ampicillin/Sulbactam 1.5g IV</p> <p>OR Cefazolin 1g IV</p> <p>OR Cloxacillin 1g IV</p> | Cefuroxime 1.5g IV | In clean cases e.g aneurysectomy the antibiotic is given for 24 hours only. In cases where there is an infective foci, continue antibiotic as treatment |
| Implantation of prosthetic grafts in patients at risk to MRSA infection | Vancomycin ¹ 500mg IV | | In patients at risk, including patients on hemodialysis and long staying in-patients as well as units that have an MRSA outbreak; this is usually given for 24 hours |
| Burns | Cloxacillin 1g IV | Cefuroxime 1.5g IV | Debridement Monitor C&S |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| 6. HEPATOBIILIARY SURGERY | | | |
| Open Cholecystectomy ERCP ± stent | Cefuroxime 1.5g IV OR 3 rd gen. Cephalosporins, e.g. Cefoperazone 1g IV | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV OR Amoxycillin/Clavulanate 1.2g IV | Antibiotic prophylaxis NOT recommended for laparoscopic cholecystectomy |
| 7. GENERAL SURGERY | | | |
| Upper GIT oesophagus, stomach & upper small bowel | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxycillin/Clavulanate 1.2g IV OR 3 rd gen. Cephalosporins, e.g. Cefotaxime, Cefoperazone 1g IV | | |
| Distal small bowel Colo-rectal | Cefuroxime 1.5g IV PLUS Metronidazole 500mg IV | 3 rd gen. Cephalosporins, e.g. Cefoperazone 1g IV PLUS Metronidazole 500mg IV; OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxycillin/Clavulanate 1.2g IV OR Ampicillin/Sulbactam 1.5g IV | |
| Hernia repair with mesh | Cloxacillin 1g IV | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxycillin/Clavulanate 1.2g IV OR Ampicillin/Sulbactam 1.5g IV | Includes laparoscopic repair |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| Breast | Cloxacillin 1g IV | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV OR Ampicillin/Sulbactam 1.5g IV | Not recommended for minor excisions |
| 8. ORTHOPAEDIC SURGERY | | | |
| Internal fixation of all closed fracture Total Joint Replacement | Cloxacillin 1g IV | Cefuroxime 1.5g IV pre-operation, continue 750mg IV q8h (3 doses) post-operation; OR Cefazolin 1-2g IV | 30-45 minutes before skin incision and before tourniquet inflation |
| Spine surgery | | | |
| Arthroscopy | | | |
| Gunshot and other penetrating wounds <i>Staphylococcus</i> <i>Clostridium species</i> | Cloxacillin 1g IV OR 2 nd gen. Cephalosporins PLUS Metronidazole 500mg IV | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV OR Ampicillin/Sulbactam 1.5g IV | Thorough surgical debridement |
| Muscular, skeletal and soft tissue trauma, crush injuries and stab wounds | Cloxacillin 1-2g q6h PLUS Gentamicin ¹ 1.5mg/kg IV q8h PLUS Metronidazole 500mg slow IV q8h Duration: Should not be less than 5 days | If possible renal impairment: Cefuroxime 1.5g IV as a loading dose followed by 750mg IV q8h PLUS Metronidazole 500mg slow IV q8h Duration: Should not be less than 5 days | In all cases, a patient's tetanus immunisation status should be assessed |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|--|
| | Preferred | Alternative | |
| Compound fractures | Cloxacillin 1g IV q6h If wound soiling or tissue damage is severe and/or devitalised tissue is present: PLUS Gentamicin ¹ 5mg/kg IV q24h PLUS Metronidazole 500mg slow IV q8h | Cefuroxime 1.5g IV as a loading dose, followed by 750mg IV q8h | In all cases, a patient's tetanus immunisation status should be assessed Duration (<i>based on the grade of fracture</i>): Grade 1: 2 weeks Grade 2: 2-4 weeks Grade 3: 2-6 weeks |
| 9. UROLOGICAL SURGERY | | | |
| A. Diagnostic Procedures | | | |
| Transrectal ultrasound and prostate biopsy <i>E coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | Ciprofloxacin 500mg PO q12h | Trimethoprim/Sulfamethoxazole 160/800mg PO q12h | 5 days (pre-emptive therapy) Oral antibiotics to start 1 day before procedure |
| Cystoscopy/Urodynamics study/ Retrograde pyelogram/Ureteric stenting | None | None | Prophylaxis only for - High risk cases (immunocompromised patients e.g. debilitated patients on long term catheters, patient with prosthesis/heart valves, diabetics, transplant recipients) - If heart valve: ❖ follow recommendation for SBE prophylaxis - Other patients: ❖ Cefuroxime 250mg PO stat |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|----------|
| | Preferred | Alternative | |
| B. Endourology | | | |
| Endourological surgery e.g. PCNL, URS, RIRS, TURP <i>E coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV</i> OR Ampicillin/Sulbactam 1.5g IV | <i>3rd gen. Cephalosporins, e.g. Cefoperazone 1g IV</i> | |
| C. Open Surgery | | | |
| Clean operations e.g. orchidectomy, orchidopexy, varicocelectomy, deroofting renal cysts <i>Staph aureus</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV stat</i> OR Ampicillin/Sulbactam 1.5g IV stat | Cefuroxime 750mg IV stat | |
| Clean-contaminated (with opening of urinary tract) e.g. <i>nephrectomy, prostatectomy, open stone surgery.</i> <i>E coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV q8h</i> OR Ampicillin/Sulbactam 1.5g IV q8h for 1 day | <i>3rd gen. Cephalosporins, e.g. Cefoperazone 1g IV q12h for 1 day</i> | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| Clean-contaminated (with use of bowel segments) e.g. Cystectomy with urinary diversion, cystoplasty. <i>E. coli, Klebsiella, Proteus, Enterococcus, Pseudomonas, Anaerobes</i> | <i>3rd gen. Cephalosporins, e.g. Cefoperazone 1g IV q12h</i> PLUS Metronidazole 500mg IV q8h | Gentamicin ¹ 1.5mg/kg IV q8h PLUS Metronidazole 500mg IV q8h | For duration of catheter presence |
| Implant of prosthetic devices e.g. Insertion of penile prosthesis or artificial urinary sphincter, artificial slings <i>Staph aureus</i> | Cefuroxime 1.5g IV q8h for 1 week | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV q8h</i> OR Ampicillin/Sulbactam 1.5g IV q8h for 1 week | Pre-emptive therapy |
| Laparoscopic surgery | As for open surgery | As for open surgery | Depending on type of procedure performed whether clean or clean - contaminated |
| Reference: European Association of Urology Guidelines 2006 | | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|----------|
| | Preferred | Alternative | |
| 10. NEUROLOGICAL SURGERY | | | |
| Clean, non-implant surgery (procedure does not cross the cranial sinuses) e.g. Tumour excision, evacuation of intracerebral clots <i>Staphylococcus aureus</i> Gram-positive cocci Gram-negative bacilli | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV stat at induction of anaesthesia and q6h during surgery | Cefuroxime 1.5g IV at induction of anaesthesia and q3h during surgery | |
| Clean-contaminated surgery (procedure crosses the cranial sinuses) e.g. Transphenoidal surgery | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV PLUS Metronidazole 500mg IV at induction of anaesthesia and q3h during surgery | Cefuroxime 1.5g IV PLUS Metronidazole 500mg IV at induction of anaesthesia and q3h during surgery | |
| CSF shunt surgery <i>Coagulase - Negative</i> <i>Staphylococcus spp Staphylococcus aureus</i> <i>Aerobic gram-ve bacilli</i> (Aerobic gram-ve bacilli are late infections) | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV OR Cefuroxime 1.5g IV | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| 11. GASTROENTEROLOGY | | | |
| ERCP ANTIBIOTIC PROPHYLAXIS | | | |
| - Bile stasis - Pancreatic Pseudocyst - Previous Cholangitis | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 2g IV 30 minutes before procedure | Gentamicin ¹ 120mg IV just before procedure OR Ciprofloxacin 750mg PO 60-90 minutes before procedure | Prompt and adequate biliary drainage is essential in biliary obstruction |
| PERCUTANEOUS ENDOSCOPIC GASTROSTOMY (PEG) | | | |
| PEG PEJ* | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV OR Cefuroxime 1.5g IV given 30 minutes before procedure | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 2g IV 30 minutes before procedure | * Percutaneous endoscopic Jejunostomy <u>Reference:</u> <i>Am J Gastro 95:3133, 2000</i> |
| UPPER GI BLEEDING IN CIRRHOSIS (Antibiotic Prophylaxis) | | | |
| Upper GI bleeding in cirrhosis | Ciprofloxacin 500mg PO q12h OR 200mg IV q12h for 7 days | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV q24h for 7 days OR Cefotaxime 2g IV q8h for 7 days | Should be offered to all cirrhotics with upper GI bleeding <u>Reference:</u> <i>Cochrane database 2002(2): CD002907</i> |
| <u>Reference:</u> British Society of Gastroenterology | | | |

12. OPHTHALMOLOGY

Use of povidone iodine 5% as an antiseptic agent for preparation of skin and conjunctival sac preoperatively is recommended

Proper draping of the eyelid margin using an adhesive non porous drape and the use of speculum to cover all the eyelashes is recommended

Intracameral injection of 1mg Cefuroxime in 0.1ml at the end of cataract surgery is recommended. Careful dilution should be undertaken to prevent potential toxicity

Reference:

Prophylaxis for intraocular surgery-CPG for Management of Post-Operative Endophthalmitis, Ministry of Health Malaysia, August 2006

*Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

B. Non-Surgical Chemoprophylaxis

1. PREVENTION OF BACTERIAL ENDOCARDITIS

(a) Cardiac conditions for which prophylaxis is recommended

High risk category

- ♦ Prosthetic cardiac valves, including bioprosthetic and homograft valves
- ♦ Previous bacterial endocarditis
- ♦ Complex cyanotic congenital heart disease (e.g. single ventricle states, transposition of the great arteries, Tetralogy of Fallot)
- ♦ Surgically constructed systemic pulmonary shunts or conduits

Moderate risk category

- ♦ Most other congenital cardiac malformations (other than above & below)
- ♦ Acquired valvular dysfunction (e.g. rheumatic heart disease)
- ♦ Hypertrophic cardiomyopathy
- ♦ Mitral valve prolapse with valvular regurgitation and/or thickened leaflets

(b) Dental Procedures for which prophylaxis is recommended

- ♦ Dental Extractions
- ♦ Periodontal procedures including surgery, scaling and root planing, probing and recall maintenance
- ♦ Dental implant placement and reimplantation of avulsed teeth
- ♦ Endodontic (root canal) instrumentation or surgery only beyond the apex
- ♦ Subgingival placement of antibiotic fibers or strips
- ♦ Initial placement of orthodontic bands but not brackets
- ♦ Intraalveolar local anaesthetic injections
- ♦ Prophylactic cleaning of teeth or implants where bleeding is anticipated

(c) Other Procedures for which prophylaxis is recommended

Respiratory Tract

- ♦ Tonsillectomy and/or adenoidectomy
- ♦ Surgical operations that involve respiratory mucosa
- ♦ Bronchoscopy with a rigid bronchoscope

Gastrointestinal Tract

- ♦ Sclerotherapy for esophageal varices
- ♦ Esophageal stricture dilation
- ♦ Endoscopic retrograde cholangiography with biliary obstruction
- ♦ Biliary tract surgery
- ♦ Surgical operations that involve intestinal mucosa

Genitourinary Tract

- ♦ Prosthetic surgery
- ♦ Cytoscopy
- ♦ Urethral dilation

NATIONAL ANTIBIOTIC GUIDELINE 2008

**PROPHYLACTIC REGIMENS FOR DENTAL, ORAL RESPIRATORY TRACT OR
OESOPHAGEAL PROCEDURES**

| Situation | Agents | Regimens |
|---|--------------------------------------|--|
| Standard General Prophylaxis | Amoxicillin | 2g PO 1h prior to procedure |
| Unable to take oral medications | Ampicillin | 2g IM/IV within 30min prior to procedure |
| Allergic to penicillin | Clindamycin | 600mg PO 1h prior to procedure |
| | Cephalexin | 2g PO 1h prior to procedure |
| | Azithromycin OR Clarithromycin | 500mg PO 1h prior to procedure |
| Allergic to penicillin and unable to take oral medication | Cefazolin/ Ceftriaxone | 1g IM/IV within 30min prior to procedure |
| | OR Clindamycin | 600mg IV within 30min prior to procedure |

- Note:
1. Cephalosporins should not be used in individuals with immediate type hypersensitivity reaction (urticaria, angioedema, or anaphylaxis) to penicillins
 2. For established respiratory infection, if Staphylococcus is suspected, give prophylactic regimens containing anti-staphylococcal penicillins or cephalosporins or Vancomycin¹ if unable to tolerate beta lactams

NATIONAL ANTIBIOTIC GUIDELINE 2008

PROPHYLACTIC REGIMENS GENITOURINARY/GASTROINTESTINAL (EXCLUDING OESOPHAGEAL) PROCEDURES

| Situation | Agents | Regimens |
|--|---|--|
| High risk patients | Ampicillin PLUS Gentamicin ¹ | Ampicillin 2g IM/IV PLUS Gentamicin ¹ 1.5mg/kg (not to exceed 120mg) within 30min prior to procedure FOLLOWED BY Ampicillin 1g IM/IV OR Amoxicillin 1g PO 6h later |
| High risk patients allergic to Ampicillin/ Amoxicillin | Vancomycin ¹ PLUS Gentamicin ¹ | Vancomycin ¹ 1g IV over 1-2h PLUS Gentamicin ¹ 1.5mg/kg IV/IM (not to exceed 120mg). Complete infusion within 30min of starting procedure |
| Moderate risk patients | Amoxicillin OR Ampicillin | Amoxicillin 2g PO 1h prior to procedure OR Ampicillin 2g IM/IV within 30min prior to procedure |
| Moderate risk patients allergic to Ampicillin/ Amoxicillin | Vancomycin ¹ | Vancomycin ¹ 1g IV over 1-2h complete infusion within 30min of starting procedure |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

Note: No second dose of Vancomycin or Gentamicin is recommended

NATIONAL ANTIBIOTIC GUIDELINE 2008

2. RHEUMATIC FEVER**a) SECONDARY PREVENTION OF RHEUMATIC FEVER (Prevention of recurrent attacks)**

Benzathine Penicillin 1.2 mega units IM every 4 weeks (in high risk situations give every 3 weeks) **OR**

Phenoxyethylpenicillin 250mg PO q12h

If allergic to Penicillin:

EES 400mg PO q12h

b) DURATION OF SECONDARY PREVENTION OF RHEUMATIC FEVER PROPHYLAXIS

| | |
|---|---|
| Rheumatic fever with carditis and residual heart disease (persistent valvular disease - clinical or echocardiograph evidence) | At least 10 years since last episode and at least until age of 40 years, sometimes lifelong prophylaxis |
| Rheumatic fever with carditis but no residual heart disease (no valvular disease) | 10 years or well into adulthood, whichever is longer |
| Rheumatic fever without carditis | 5 years or until age 21 years, whichever is longer |

3. RECOMMENDATIONS FOR PREVENTION OF INFECTION IN ASPLENIA (OR HYPOSPLENIA) ADULT PATIENTS**A. Antibiotics Prophylaxis**

| | |
|-------------------------|---|
| Antibiotics Prophylaxis | <ol style="list-style-type: none"> 1. Phenoxyethylpenicillin 250-500mg PO q12h OR Amoxicillin 500mg PO q12h 2. Penicillin allergy - EES 400mg PO q12h OR Azithromycin 250mg PO q24h 3. Duration: Minimum 2 years post splenectomy is encouraged in adults. Up to 16 years of age in children. Life long is not recommended (<i>McMullin 1993</i>). <i>Long term management of patients after splenectomy. BMJ 307, 1372-1373</i> 4. Emergency supply of antibiotic: Alternative to OR in addition to long term prophylaxis <ol style="list-style-type: none"> a) Amoxicillin 3g PO should be kept at home if fever occurs OR b) Cefuroxime 1g PO OR c) Amoxicillin/Clavulanate 625mg PO OR d) If taking EES, increase dose to 800mg PO q12h OR e) If taking Azithromycin, increase dose to 500mg PO q24h OR f) Clindamycin 600mg PO OR g) Trimethoprim/Sulphamethoxazole 960mg PO <p>Take higher regime as stat dose and seek medical advice as soon as possible</p> |
|-------------------------|---|

NATIONAL ANTIBIOTIC GUIDELINE 2008

| | |
|-------------------------------|--|
| Patient Education | Inform patient (and relative/friend) of increased risk of infection and strategies to prevent bacterial infections. Discuss OPSI (overwhelming post splenectomy infection), tick and animal bites/scratches. Provide immunisation card |
| Blood test | FBC and PBF-assessing presence of Howell Jolly bodies |
| Travel Recommendations | 1. Seek medical advice before travel 2. Ensure meningococcal vaccination is current for travel to high incidence countries 3. Always carry the immunisation card |
| Alerts | Patient is encouraged to wear/carry medic alert medallion or wallet card |
| SEEK MEDICAL ATTENTION | Fever, shivers, vomiting, prolonged sore throat (signs of bacterial infection) |

B. Vaccine

| Vaccine Recommendation | Which vaccine | Route | Timing | Re-vaccination |
|---|--|--------------------------|---|--|
| Pneumococcal vaccine | Pneumococcal 23-valent polysaccharide vaccine (Pneumo 23) | 0.5ml S/C or IM | > 2 weeks before elective surgery. 7-14 days after emergency splenectomy or prior to discharge | Booster every 5 years |
| Meningococcal vaccines <i>polysaccharide</i> | Meningococcal quadrivalent polysaccharide ACWY vaccine (Mencevax ACWY or Menomune) | 0.5ml S/C | As above | Polysaccharide ACWY Booster every 5 years |
| Hemophilus influenzae type B | HiB (Liquid Pedvax Hib) Annually | 0.5ml IM thigh/upper arm | As above | No booster required |
| Influenza | | 0.5ml deep S/C | | Annual |

For patient with bleeding disorder and there is concern about giving vaccinations, vaccinations are given subcutaneously including HiB vaccine. Any doubt please contact Haematology Registrar

GASTROINTESTINAL INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| 1. OESOPHAGITIS | | | |
| a. Fungal Infections | Refer to Page 53 (Human Immunodeficiency Virus) | Acyclovir 400mg PO q8h for 7-10 days | Duration of therapy represents total time IV, PO, or IV + PO. Most patients on IV therapy able to take PO medications should be switched to PO therapy soon after clinical improvement (usually < 72 hours) |
| b. Viral <i>HSV-1</i> | Acyclovir 5mg/kg IV q8h for 7-10 days | | |
| <i>CMV</i> | Ganciclovir 5mg/kg IV q12h for 3-6 weeks | | |
| 2. Helicobacter Pylori INFECTION (Ref. P. Malfertheiner et al. GUT 2007; 56:772-781) | | | |
| <ul style="list-style-type: none"> - Peptic ulcer disease (Including complicated PUD) - MALToma - Atrophic gastritis - After gastric cancer resection - Patient who are first-degree relatives of patients with gastric cancer - Non-ulcer dyspepsia - Naive NSAID users - Chronic NSAID users - Long term aspirin use - Long term PPI therapy - Immune Thrombocytopenic Purpura and iron deficiency anaemia | <p>*Proton Pump Inhibitors (PPI) e.g. Omeprazole, Pantoprazole, Lansoprazole, Rabeprazole, Esomeprazole PO q12h for 7 days</p> <p>PLUS Clarithromycin 500mg PO q12h for 7 days</p> <p>PLUS Metronidazole 400mg PO q12h for 7 days</p> <p>OR Amoxicillin 1g PO q12h for 7 days</p> | <p>PPI, e.g. Omeprazole 20mg PO q12h</p> <p>PLUS Amoxicillin 1g PO q12h</p> <p>OR Tetracycline 500mg PO q8h</p> <p>PLUS Metronidazole 400mg PO q8h for 10 days</p> | <ul style="list-style-type: none"> - First choice therapy recommended in areas with <15-20% Clarithromycin resistance. - Bismuth-based quadruple therapy for 7-10 days may be used as second choice therapy if available. - Third choice or rescue treatment should be based on antibiotic susceptibility testing <p>* Dosages:- Omeprazole 20mg q12h Pantoprazole 40mg q12h Lansoprazole 30mg q12h Rabeprazole 20mg q12h Esomeprazole 20mg q12h</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| 3. INFECTIOUS DIARRHOEA (Reference: NEJM 342: 1716, 2000; JID 185: 133, 2002; CID 39: 504, 2004) | | | |
| a. Acute Watery Diarrhoea Campylobacter Yersinia Salmonella Aeromonas Plesiomonas sp | Ciprofloxacin 500mg PO q12h for 3-5 days | Trimethoprim/Sulfamethoxazole 160/800mg PO q12h for 3-5 days | - Antibiotics are not indicated in acute or uncomplicated diarrhoea (<i>Oral Rehydration Solution will be sufficient</i>) - Antibiotics may be considered when patients have fever (>38.5°C) and severe diarrhoea in the elderly |
| b. Acute Dysentery <i>E. histolytica</i> | Metronidazole 800mg PO q8h for 10 days | Tinidazole 1g PO q12h for 3 days | Fever and bloody stool are features of dysentery |
| <i>Shigella</i> | Ciprofloxacin 200-400mg IV or 500mg PO q12h for 3 days | Trimethoprim/Sulfamethoxazole 160/800mg PO q12h for 3 days OR Azithromycin 500mg IV or PO q24h for 3 days | |
| c. Chronic Watery Diarrhoea <i>Giardia lamblia</i> | Metronidazole 400-800mg PO q8h for 5 days | Albendazole 400mg PO q24h for 5 days OR Tinidazole 2g stat | |
| <i>Cryptosporidia</i> | Treatment is unsatisfactory | | |
| <i>Cyclospora</i> | Trimethoprim/Sulfamethoxazole 160/800mg PO q12h for 7-10 days | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|--|
| | Preferred | Alternative | |
| d. Antibiotic-associated Diarrhoea <i>Clostridium difficile</i> | | | |
| Uncomplicated | Metronidazole 400mg PO q8h for 14 days | Vancomycin 125mg PO q6h for 14 days | - Discontinue offending antibiotic if possible. Avoid antimotility agents |
| Severe with ileus or toxic mega colon | Metronidazole 500mg IV q8h | Vancomycin 500mg PO q6h (via nasogastric tube) | - Rifampicin may be added to Vancomycin for relapsing disease |
| Relapsing disease | Metronidazole 400mg PO q8h for 10 days | Vancomycin PO tapering dose over 4 weeks or 125mg EOD for 6 weeks | - The IV preparation of Vancomycin may be taken orally if oral Vancomycin is not available |
| 4. LIVER ABSCESS | | | |
| a. Pyogenic Liver Abscess | | | |
| <i>Enterobacteriaceae</i> <i>Enterococci</i> <i>Bacteroides</i> | Ampicillin 1-2g IV q6h PLUS Gentamicin ¹ 1.5mg/kg IV q8h PLUS Metronidazole 500mg IV q8h for 14 days; OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5-3g IV q6h for 14 days | Metronidazole 500mg IV q8h PLUS <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1-2g IV q24h OR Ciprofloxacin 400mg IV q12h for 14 days | Treat until clinical improvement achieved Surgical or percutaneous drainage may be required Follow-up ultrasound scans recommended Metronidazole may be added to the regimen if an amoebic liver abscess cannot be excluded |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|------------------------------------|---|
| | Preferred | Alternative | |
| b. Amoebic Liver Abscess <i>Entamoeba histolytica</i> | Metronidazole 500mg IV q8h for 10 days (May switch to PO when clinical improvement occurs) | Tinidazole 2g PO q24h for 3-5 days | |
| 5. CHOLECYSTITIS (Ref: M. Yoshida et al. J. Hepatobiliary Pancreat. Surg (2007) 14:83-90) | | | |
| a. Mild <i>E. coli</i> <i>Klebsiella</i> <i>Enterococci</i> | β -lactam/ β -lactamase inhibitors, e.g. Ampicillin/Sulbactam 3g IV q6h for 7 days OR Ciprofloxacin 500mg PO q12h for 7 days | | Grade I (mild) acute cholecystitis is defined as acute cholecystitis in a patient with limited gallbladder disease, making cholecystectomy a low risk procedure |
| b. Moderate <i>E. coli</i> <i>Klebsiella</i> <i>Enterococci</i> | β -lactam/ β -lactamase inhibitors, e.g. Ampicillin/Sulbactam 3g IV q6h for 7 days | | Grade II (moderate) acute cholecystitis is associated with extensive gallbladder disease resulting in difficulty in safely performing a cholecystectomy |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|--|
| | Preferred | Alternative | |
| c. Severe <i>E. coli</i> <i>Klebsiella</i> <i>Enterococci</i> | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q12h for 7 days PLUS Metronidazole 500mg IV q8h for 7 days | Ciprofloxacin 400mg IV q12h for 7 days PLUS Metronidazole 500mg IV q8h for 7 days OR *Cefoperazone/Sulbactam 2g IV q12h for 7 days PLUS Metronidazole 500mg IV q8h for 7 days OR Imipenem 500mg IV q6h for 7 days OR Meropenem 1g IV q8h for 7 days | Grade III (severe) acute cholecystitis is defined as acute cholecystitis with organ dysfunction *Reserved for <i>Acinetobacter</i> |
| 6. CHOLANGITIS (Reference: A. Tanaka et al. <i>J. Hepatobiliary Pancreat Surg</i> (2007) 14:59-67) | | | |
| Normal host <i>E. coli</i> <i>Klebsiella</i> <i>Enterococci</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 3g IV q6h for 7 days OR <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q24h for 7 days OR Cefoperazone 2g IV q12h for 7 days PLUS Metronidazole 500mg IV q8h for 7 days | Ciprofloxacin 400mg IV q12h PLUS Metronidazole 500mg IV q8h for 7 days OR Imipenem 500mg IV q6h for 7 days OR Piperacillin/Tazobactam 4.5g IV q8h for 7 days (If <i>Pseudomonas</i>) | Duration of treatment is a minimum of 7 days Antimicrobial therapy should be selected according to the severity assessment Empirical agents should be changed according to bile C&S reports Biliary drainage should be performed for moderate to severe cholangitis |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| 7. ACUTE PANCREATITIS (ANTIBIOTIC PROPHYLAXIS) | | | |
| <i>(Ref: UK guidelines for the management of Acute Pancreatitis GUT 2005; 54:1-9)</i> | | | |
| Severe acute pancreatitis (CT evidence of >30% necrosis) | Imipenem 500mg IV q6h for 7-14 days | | The evidence for antibiotic prophylaxis in severe acute pancreatitis is conflicting. There is currently no clear consensus |
| 8. PANCREATIC INFECTIONS | | | |
| <i>(Am J Gastroenterol 2006; 101:2379-2400)</i> | | | |
| Infected pancreatic necrosis <i>Enterobacteriaceae</i> <i>B. fragilis</i> Pancreatic abscess Infected Pseudocyst | Ciprofloxacin 400mg IV q12h PLUS Metronidazole 500mg IV q8h for 14 days | Imipenem 500mg IV q6h for 14 days OR Meropenem 1g IV q8h for 14 days OR Piperacillin/Tazobactam 4.5g IV q8h for 14 days | CT-guided percutaneous aspiration with Gram's stain and culture is recommended when infected necrosis is suspected Culture of Abscess, infected pseudocyst or infected necrosis should guide treatment Drainage of the abscess and/or surgery may be required |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| 9. DIVERTICULAR DISEASE | | | |
| <i>(Ref: World Gastroenterology Organization (WGO) Practice Guidelines)</i> | | | |
| Diverticulitis <i>E. coli</i> <i>B. fragilis</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h for 7days OR Ciprofloxacin 200-400mg IV q12h PLUS Metronidazole 500mg IV q8h for 7days | | If there is no improvement in 48-72 hours, look for complications e.g. abscess and perforation |
| 10. INTRA-ABDOMINAL INFECTIONS/PERITONITIS | | | |
| <i>(Reference: Clin Infection. Dec. 2003; 37:997-1005)</i> | | | |
| Spontaneous bacterial peritonitis (SBP)* <i>Enterobacteriaceae</i> <i>For other on Intra-abdominal Infections/peritonitis - Refer to Page 120 (Surgical Infections)</i> | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 2g IV q8h for 5 days | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q24h for 5 days | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|----------|
| | Preferred | Alternative | |
| 11. HEPATOSPLENIC CANDIDIASIS | | | |
| Hepato-splenic candidiasis <i>Candida albicans</i> | Fluconazole 400mg IV/PO q24h for 21 days (or at least 2 weeks after being culture negative) | Amphotericin B 0.5mg/kg IV q24h for 21 days | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

INFECTIONS IN IMMUNOCOMPROMISED PATIENTS

A. HAEMATOLOGY

- Any infection in the immunocompromised host is life-threatening and needs immediate attention. Neutropaenic sepsis is defined as a temperature of $> 38.3^{\circ}\text{C}$ or $> 38^{\circ}\text{C}$ over one hour and ANC < 500 cells/uL or < 1000 cells/uL in those with anticipated declining counts.
- Cultures may be positive in less than 40% of cases. Patients have impaired inflammatory responses and hence may have no localizing signs. The usual sign is fever $> 38^{\circ}\text{C}$ or hypothermia. Empirical antibiotics must be started immediately after appropriate blood cultures are taken. The common portals of infection include the oral cavity, gastrointestinal tract, perianal region, lungs and IV lines.
- Potential pathogens are dependent on the underlying defect, e.g.

| | |
|-----------------------------|--|
| Neutropaenia | Gm -ve organisms Gm +ve organisms Fungi |
| Hypogammaglobulinaemia | Encapsulated organisms |
| Defective cellular immunity | Pneumocystis, Toxoplasma Fungi Viruses Mycobacteria |

- The choice of antibiotics is based on local organisms and sensitivity patterns. This should depend on sound clinical judgement, the clinical state of the patient, prior infections, recent outbreaks e.g. MRSA or multiresistant Klebsiella, E coli as well as the availability and cost of the antibiotics. The incidence of ESBL-producing organisms in the local setting must be borne in mind when selecting agents for use in the first line setting. Many less virulent or uncommon organisms are also increasingly seen e.g. *Stenotrophomonas maltophilia*, *Acinetobacter spp.*
- For neutropaenic adult patient, the following regimens are suggested:
 - 1st line** Piperacillin/Tazobactam 4.5g IV q6h **OR** Cefepime 2g IV q8h. Aminoglycosides e.g. Gentamicin or Amikacin may be added in combination therapy.
 - 2nd line** Carbapenem: Imipenem 500mg IV q8h/q6h **OR** Meropenem 1g q8h. Imipenem 1g q8h is used in severe sepsis.
 - Monotherapy** is likely just as efficacious and less toxic. Drugs that can be used as monotherapy are Piperacillin/Tazobactam, Cefepime, Imipenem or Meropenem.
 - Anaerobic infections** account for $< 5\%$ of all cases of bacteraemia. Piperacillin/Tazobactam and Carbapenems generally have good anaerobic coverage. Metronidazole 500mg IV q8h may be added in the presence of severe mucositis, intraabdominal infections, perirectal abscesses or colitis.

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- e. **Glycopeptide therapy** e.g. Vancomycin **OR** Teicoplanin can be delayed 48-72h without risk. Vancomycin 15mg/kg IV q12h or q8h may be added in suspected central device infections, known colonizers by MRSA, severe mucositis, suspected MRSA/MRSE infections and severe sepsis, septic shock or respiratory distress. Linezolid is an alternative in those patients with no clinical response to Vancomycin and in those with VRE, VISA or VRSA.
- f. **Antifungal therapy** is added from day 5 to 7 or earlier especially for severe mucositis, thrush, painful swallowing, suspicious skin infiltrates or pulmonary infiltrates, fundal exudates or after prolonged steroid/antibiotic use > 2 weeks. Amphotericin B remains the empirical therapy of choice for invasive fungal treatments. For patients who are intolerant, refractory or those with toxicity, the lipid formulations and Caspofungin are alternative as empirical therapy. Voriconazole is an alternative to Amphotericin B for the treatment of invasive aspergillosis.
- g. **The use of growth factors** e.g. G-CSF or GM-CSF may be considered but the benefits in this setting have not been proven. It should be considered in high-risk patients with ANC < 100/uL, MODS, pneumonia, invasive fungal infections or septic shock.
- h. **The use of immunoglobulins** and IgM enriched preparations has not shown survival benefits in adult patients with sepsis.
- i. **The role of granulocytes** remains controversial. Granulocyte transfusions may be used in patients with serious bacterial or fungal infections not responding to appropriate treatment and who will likely recover in the neutrophil count in the short term. The risk of disease transmission e.g. CMV must be borne in mind.
- j. **The use of oral antibiotics** in an outpatient setting for low risk patients is currently not advised as the risks stratification have not been validated in a local setting, the local resistance patterns of organisms to the oral therapy e.g. Ciprofloxacin and Amoxicillin/Clavulanate as well as the lack of local facilities for immediate access to prompt medical attention in the outpatient.
- k. **Prophylaxis against bacterial or fungal infections** is advised after bone marrow transplantation or in the high-risk patient after chemotherapy. In the routine setting, it results in increasing resistance and is expensive.
- l. Infections following stem cell transplant are generally similar to that in the solid organ transplant setting. In addition to the usual bacterial and fungal infections, viral infections especially CMV reactivation and parasitic infections e.g. Pneumocystis carinii and Toxoplasma infection can occur. It is recommended that prophylactic use of Ganciclovir or preemptive monitoring for CMV reactivation should be carried out during the first 100 days. Trimethoprim/Sulphamethoxazole 6-8 tablets per week is also extremely effective in the prevention of PCP or toxoplasmosis. It is recommended that these measures be continued in patients with active graft-vs-host disease and in those remaining on high dose immunosuppressives.

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| | | |
|----------------------|--|---|
| 1 st line | Piperacillin/Tazobactam 4.5g IV q6h OR Cefepime 2g IV q8h | Aminoglycosides e.g. Gentamicin or Amikacin may be added in combination |
| 2 nd line | Imipenem 500mg IV q8h or q6h or 1g q8h (severe sepsis) OR Meropenem 1g q8h | |
| Glycopeptides | Vancomycin 15mg/kg IV q12h or q8h | May be delayed 48-72h until cultures, unless indicated |
| Antifungal agents | Conventional Amphotericin B Liposomal Amphotericin B Caspofungin | May be added as empirical therapy from D5-7 Voriconazole preferred in invasive aspergillosis |

6. Attention must be paid to:
- Strict isolation measures
 - Patient's personal hygiene and diet
 - Modification of antibiotic regimen if deterioration of clinical status or if there is no clinical improvement in 72-96h in a stable patient
 - The antibiotics are generally kept for a minimal duration of 5 to 7 days or stopped if afebrile for 3 days in patients with improving neutrophil counts
 - Regular culture and surveillance
 - HAND WASHING and strict aseptic technique
 - Venous canula must be inspected daily for signs of phlebitis and changed every 72h or when necessary. Central devices are removed if there is clinical deterioration in spite of appropriate antibiotics for 48-72h

References:

- NCCN Clinical Practice Guidelines in Oncology V.1 2006. Fever and Neutropaenia
- Hughes WT, Armstrong D, Bodey GP et al. 2002 Guidelines for the use of antimicrobial agents in neutropenic patients with cancer. Clin Infect Dis 2002; 34:730-751
- Herbrecht R, Denning DW, Patterson TF et al. Voriconazole versus amphotericin B for primary therapy of invasive aspergillosis. NEJM 2002; 347:408-415
- Walsh TJ, Tepler H, Donowitz GR et al. Caspofungin versus liposomal amphotericin B for empirical antifungal therapy in patients with persistent fever and neutropaenia. NEJM 2004; 351(14):1391-1402

B. Human Immunodeficiency Virus (HIV)

*Important cut-offs for CD4 T cells, above which particular AIDS illnesses are improbable.
These CD4 counts are only reference values; exceptions are always possible.*

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| | |
|----------------|--|
| No cut-off | Kaposi's sarcoma, pulmonary tuberculosis, HZV, bacterial pneumonia, lymphoma |
| < 250/ μ l | PCP, esophageal candidiasis, PML, HSV |
| < 100/ μ l | Cerebral toxoplasmosis, HIV encephalopathy, cryptococcosis, miliary tuberculosis |
| < 50/ μ l | CMV retinitis, cryptosporidiosis, atypical mycobacteriosis |

The treatment regimes are based on drugs available in the Ministry of Health National Formulary and hence in some instances may vary from internationally accepted treatments. Some regimes are chosen as preferred regimes due to cost considerations

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Pneumocystic Jiroveci (Carinii) | | | |
| Interstitial Pneumonia | Trimethoprim 15-20mg/kg/24h PLUS Sulfamethoxazole 75-100mg/kg/24h PO (excellent bioavailability) or IV q6h or q8h for 21 days | For severe cases: (PO₂ < 70mmHg) Pentamidine 4mg/kg/24h IV (in 1 pint D5% or N/S run over 1-2 hours) For mild to moderate cases: (PO₂ 70-80mmHg) Clindamycin 600mg IV q8h OR 300-450mg PO q6h PLUS Primaquine 30mg base PO/24h for 21 days OR Dapsone 100mg PO q24h PLUS Trimethoprim 15mg/kg/day PO (3 divided doses) | Patients with severe disease should receive steroids as soon as possible (within 72 hours of starting PCP treatment): Prednisolone 40mg PO q12h for 5 days then 40mg PO q24h for 5 days then 20mg PO q24h for 11 days |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Prophylaxis Indications: H/o PCP, CD4 < 200 or <14% HIV associated thrush, or unexplained fever > 2 weeks | Trimethoprim/Sulfamethoxazole 160/800mg q24h OR 80/400mg q24h | Dapsone 100mg PO q24h Aerosolized Pentamidine 300mg monthly via Respiguard II nebulizer or ultrasonic nebulizer ±O ₂ agonist | Patients given Dapsone should be tested for G6-PD deficiency if at risk Discontinuation: Consider in patients on HAART with CD4 > 200 for > 3-6 months Secondary prophylaxis: Should be re-introduced if the CD4+ T lymphocyte count decreases to < 200 cells/μL OR if PCP recurs at a CD4+T lymphocyte count of > 200 |
| Candidal | | | |
| Oropharyngeal (thrush) | Itraconazole 200mg PO q24h OR Nystatin suspension 400,000-600,000 units (4-6ml) q6h for 7-14 days | Fluconazole 100mg PO q24h | Suppressive therapy - generally not recommended unless patients have frequent or severe recurrences |
| Vaginitis | Azoles pessary (Clotrimazole, Miconazole) for 3-7 days | Fluconazole 150mg PO x 1 dose OR Itraconazole 200mg PO q12h for 1 day or 200mg PO q24h for 3 days | Prolonged or refractory episodes is observed in approximately 10% of patients and requires antimycotic therapy for >7 days |

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|--|
| | Preferred | Alternative | |
| Esophagitis | Fluconazole 200mg PO q24h up to 400mg q24h for 2 weeks | Itraconazole 200mg PO q12h OR Amphotericin B 0.3-0.7mg/kg IV q24h | Candidiasis is the most common cause of esophagitis with HIV infection, but CMV, HSV and aphthous ulcerations can present with similar complaints Endoscopy required with unusual presentations or lack of response to azole within several days |
| Cryptococcal meningitis or meningoenkephalitis (by <i>Cryptococcus neoformans var neoformans</i>) | | | |
| Initial Treatment | <u>Induction therapy:</u> Amphotericin B 0.7mg/kg/24h PLUS/MINUS Flucytosine 25mg/kg PO q6h for 2 weeks <u>Consolidation therapy:</u> Fluconazole 400mg PO q24h for 8 weeks or until CSF cultures are sterile | <u>Induction therapy:</u> Fluconazole 400-800mg q24h PO PLUS Flucytosine 25mg/kg PO q6h for 4-6 weeks <u>Consolidation therapy:</u> Itraconazole 200mg PO q12h | If ICP >250mm and signs of cerebral oedema present, do daily LP to reduce pressure until patient is improved If clinical signs of cerebral oedema do not improve after about 2 weeks of daily LPs, consider placement of a lumbar drain or ventriculoperitoneal shunt |
| Maintenance Therapy | Fluconazole 200mg PO q24h | Itraconazole 200mg PO q24h for patients intolerant or failed Fluconazole | Discontinuation: Consider if patient on HAART with good viral suppression and CD4>200 >6 months |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|--|
| | Preferred | Alternative | |
| Toxoplasma Gondii Encephalitis | | | |
| Acute Infection (up to 97% patients are Toxo IgG +ve) | *Pyrimethamine 100-200mg PO loading dose followed by Pyrimethamine 50-100mg PO q24h (Fansidar 1 tab q12h) PLUS Folinic acid 10-25mg PO q24h PLUS Clindamycin 600mg IV/PO q6h for at least 6 weeks | *Pyrimethamine PLUS Folinic acid (see preferred regime) PLUS Sulfadiazine 1g PO q6h OR Trimethoprim/Sulfamethoxazole (5mg/kg TMP and 25mg/kg SMX) IV or PO q12h | *1 tab Fansidar (Sulfadoxine/Pyrimethamine) contains 25mg of pyrimethamine Adjunctive corticosteroids (e.g. dexamethasone) should be administered when clinically indicated only for treatment of a mass effect associated with focal lesions or associated edema. Because of the potential immunosuppressive effects of corticosteroids, they should be discontinued as soon as clinically feasible |
| Suppressive/ Maintenance Therapy | Pyrimethamine 25-75mg PO q24h PLUS Clindamycin 300-450mg PO q6-8h PLUS Folinic acid 10-25mg q24h | Pyrimethamine 25-75mg PO q24h PLUS Folinic acid 10-25mg q24h PLUS Sulphadiazine 0.5-1g PO q24h | Discontinuation: Consider when on HAART, CD4 > 200 > 3 months and viral load well suppressed |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|----------|
| | Preferred | Alternative | |
| 1^o Prophylaxis Indications: ToxolG +ve and CD4<100 | Trimethoprim/Sulfamethoxazole 160/800mg PO q24h | Trimethoprim/Sulfamethoxazole 80/400mg PO q24h OR Dapsone 50mg/day PO PLUS Pyrimethamine 50mg/week PO PLUS Folinic acid 25mg/week PO OR Dapsone 200mg/week PO PLUS Pyrimethamine 75mg/week PO PLUS Folinic Acid 25mg/week PO | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|---|
| | Preferred | Alternative | |
| Mycobacterium Avium Complex Disease | | | |
| Treatment 1° Prophylaxis Indications: CD4 < 50 cells Ruled out MAC bacteremia and active TB | Clarithromycin 500mg PO q12h PLUS Ethambutol 15mg/kg/24h PO Clarithromycin 500mg PO q12h OR Azithromycin 1.2g weekly | Azithromycin 500-1000mg/24h PO PLUS Ethambutol (same dose) Alternate 3rd or 4th drug PLUS Amikacin 10-15mg/kg/24h IV OR Ciprofloxacin 500-750mg PO q12h OR Levofloxacin 500mg PO q24h | Discontinuation: Consider if patient is on HAART and viral load well suppressed, CD4 > 100 > 6 months, asymptomatic of MAC, and has completed > 12 months of MAC treatment Caution with Clarithromycin PLUS Efavirenz: high rates of rash |
| Cytomegalovirus Retinitis | | | |
| Initial Therapy (until scar formation on the lesion) | Ganciclovir 5mg/kg IV q12h for 2-3 weeks Maintenance Regime: Intravitreal Ganciclovir 400µg/week | Alternative maintenance: Ganciclovir 5mg/kg IV q24h | Initial therapy should also include optimisation of HAART |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| Extraocular CMV diseases: Esophageal ulcer, colitis Interstitial pneumonitis | Ganciclovir 5mg/kg IV q12h for 21-28 days or until signs and symptoms have been resolved | | Maintenance therapy is generally not necessary; HAART offers best hope for prevention of relapses |
| Salmonella (non-typhi) | | | |
| Initial Therapy | Salmonella gastroenteritis: Ciprofloxacin 500-750mg PO q12h OR 400mg IV q12h Duration: - Mild gastroenteritis without bacteremia = 7-14 days - Advanced HIV (CD4+ <200) and/or bacteremia = at least 4-6 weeks | Trimethoprim/Sulfamethoxazole PO OR <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone IV OR Cefotaxime IV | |
| Maintenance Therapy | Trimethoprim/Sulfamethoxazole 160/800 PO q12h | | Discontinuation: Consider once patient on HAART, viral load well suppressed and CD4 > 200 > 6 months |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|---|-------------|--|
| | Preferred | Alternative | |
| Herpes Simplex | | | |
| | Genital or orolabial herpes: Acyclovir 400mg PO q8h OR 800mg PO q12h for 5-10 days Moderate-to-severe mucocutaneous HSV infections: Initial therapy - Acyclovir 5mg/kg IV q8h After lesion begins to regress, Acyclovir 400mg PO q8h until lesions have completely healed Suppressive therapy: Acyclovir 400mg PO q12h | | Suppressive therapy indicated if herpes outbreaks frequent or severe |
| Herpes Zoster | | | |
| Initial Therapy | Acyclovir 800mg PO 5x/day for 7-10 days Severe infection (CNS, ocular, disseminated): Acyclovir 10mg/kg IV q8h for 14-21 days | | Effective in immune competent patients only if initiated within 72h, but for immune suppressed, treat unless lesions crusted Consider treatment for severe infection whenever clinical diagnosis of zoster likely + altered mental status or visual symptoms while definitive diagnosis pursued |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|--|---|
| | Preferred | Alternative | |
| Histoplasmosis | | | |
| Initial Therapy | Induction regime: Amphotericin B 0.6-0.7mg/kg IV q24h for 2 weeks Continuation phase: (12 weeks) Itraconazole 200mg PO q12h Chronic maintenance therapy: Itraconazole 200mg PO q24h | In less severe disease: Itraconazole 200mg PO q8h for 3 days, then 200mg PO q12h for 12 weeks | Consider discontinuation among patients who remain asymptomatic, with CD4+ count > 100-200 cells/ μ L for > 6months Syrup Itraconazole has better bioavailability and hence preferred by some for the induction phase in less severe disease |
| Isospora Belli Infection | | | |
| Initial Therapy | Trimethoprim/Sulfamethoxazole 160/800mg PO/IV q6h for 10 days OR Trimethoprim/Sulfamethoxazole 320/1600mg PO/IV q12h for 10-14 days | Pyrimethamine 50-75mg PO q24h PLUS Folinic acid 5-10mg PO q24h; OR Ciprofloxacin 500mg PO q12h | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|---|--|
| | Preferred | Alternative | |
| Nocardia | | | |
| Initial Therapy | Trimethoprim PLUS Sulfamethoxazole (TMP 15mg/kg/24h + SMX 75mg/kg/24h) IV or PO in four divided doses. May consider decreasing to SMX/TMP (TMP 10mg/kg/24h) after clinical improvement | Imipenem/Cilastatin 500mg IV q6h PLUS Amikacin ¹ 7.5mg/kg IV q12h for 2-4 weeks or clinical improvement followed by oral regimen OR <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q12-24h PLUS Amikacin ¹ 7.5mg/kg IV q12h for 2-4 weeks or clinical improvement followed by oral regimen | Use indefinite low dose oral suppression in patients with advanced HIV or significant immunosuppression to prevent relapse with TMP-SMX 160/800 q12h |
| Penicilliosis | | | |
| Initial Therapy | Induction regime: Amphotericin B 0.6-0.7mg/kg IV q24h for 2 weeks Continuation phase: (12 weeks) Itraconazole 200mg PO q12h Chronic maintenance therapy: Itraconazole 200mg PO q24h | In less severe disease: Itraconazole 200mg PO q8h for 3 days, then 200mg PO q12h for 12 weeks | Consider discontinuation among patients who remain asymptomatic, with CD4+ count >100-200 cells/ μ L for >6 months Syrup Itraconazole has better bioavailability and hence preferred by some for the induction phase in less severe disease (same dose) |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|------------------------------------|-------------|--|
| | Preferred | Alternative | |
| Progressive Multifocal Leukoencephalopathy (PML) | | | |
| Initial Therapy | No effective therapy exists | | With HAART, some patients improve and others stabilise. Few may deteriorate due to immune reconstitution |
| Cryptosporidiosis | | | |
| Initial Therapy | Symptomatic treatment of diarrhoea | | Effective ART (to increase CD4+ count to >100) can result in complete, sustained clinical, microbiological and histologic resolution |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

C. SOLID TRANSPLANT

Approach to Post-Solid Organ Transplant - related Infections

(Renal and Liver Transplantation)

As most organ transplant recipients require immunosuppression, which though remarkably effective at controlling rejection, can produce a wide range of undesirable side-effects, especially a predisposition to serious infections. This chronic risk of infection, with its diagnostic problems and potentially fatal outcome, mandates an understanding of the principles of transplant-associated infections.

The following brief discussion of the approach to transplant-associated infections is meant to assist, alert and orient the physician who does not deal routinely with infections in the compromised host.

Consultation with infectious disease physician is recommended.

Important considerations in transplant-related infection;

- ◆ Tissue rejection notoriously mimics infections in solid organ transplantation. In all febrile episodes, the clinician must first consider rejections as a cause of fever.
- ◆ Medication side effects can cause fevers; thus the drug list should be reviewed for possible causative agents.
- ◆ The presenting features of infection in patients on immunosuppressive therapy may be vague as the impaired inflammatory response results in a paucity of physical signs and atypical presentation of infective processes. The insidious onset and rapid progression of infections warrant a prompt, thorough evaluation early in the course of any febrile event. The initiation of empiric broad-spectrum antibiotics is reasonable in patients with rigors or leucopenia. Opportunistic organisms are important considerations in the evaluation of febrile episodes in transplant patients and these include the following: cytomegalovirus (CMV), herpes simplex virus (HSV), fungal infections eg. candida and aspergillus, pneumocystis, mycobacteria, etc. There exist an 'infection timetable' especially in renal and heart transplant, whereby some specific pathogens often cause infections at certain time intervals from onset of immunosuppressions. (Figure 1)

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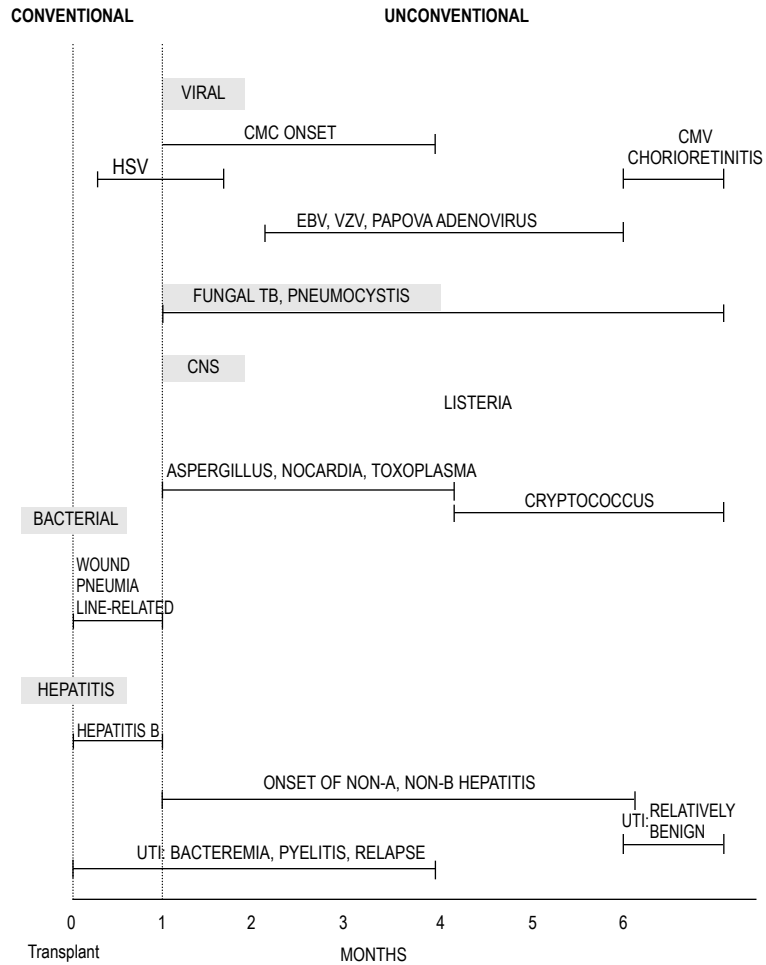


Figure 1
Timetable of occurrence of infection in renal transplant recipient

Post Liver Transplant-related Infections:

Febrile episodes in orthotopic liver transplant (OLT) are caused by infections in 80% of cases. Predominant causes of fever are bacterial infections (62%), viral (6%); whereas rejection accounts for only 4% of febrile episodes.

Bacteraemic infections are a major cause of death among organ transplant patients; for liver transplant patients the portal of entry is mainly the gastrointestinal and biliary tract with *Pseudomonas aeruginosa* and *Enterobacter* species having particularly high fatality rates. These infections are often seen in the early post transplant period (< 100 days). Stool cultures obtained before OLT are useful for choice of perioperative prophylactic/empirical antibiotics.

The most common sites of infection are generally in the abdomen followed by the blood stream. Commonest infections are bacterials followed by fungal infections. Gram positive aerobic bacterial infections are more common than Gram negative infections with portal vein thrombosis being an important risk factor for early bacterial infection.

The need of empirical antibiotic therapy in transplant patients with pulmonary infiltrates in intensive care units (ICU) can be assessed using several factors including; clinical pulmonary infection score (Pugin score) > 6, abnormal temperature and serum creatinine > 1.5mg/dl. Pugin score > 6 warrants antimicrobial therapy. Common causative bacterial organisms include; *Methicillin Resistant Staphylococcus aureus* (MRSA), *Pseudomonas aeruginosa*, *Enterobacter spp.* and *Serratia marcescens*. *Aspergillus* pulmonary infections should also be suspected in early onset pneumonia within 30 days of transplantation.

CMV infection is a common post-transplant occurrence; it maybe primary or secondary (ie. reactivation); being the most common cause of hepatitis in liver allograft patients. Infection usually presents within 90 days of transplant and continue for months (even years) in those with poor graft function requiring heavy immunosuppression. Long term Ganciclovir for the first 100 days post-transplant largely eliminates CMV infection.

INFECTIONS IN INTENSIVE CARE UNIT

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|---|
| | Preferred | Alternative | |
| A. Severe Sepsis Or Septic Shock Where Site Of Infection Is Not Identified | | | |
| Severe sepsis or septic shock (site of infection is unknown) <i>Gram-negative bacilli</i> <i>Gram-positive cocci</i> | Cefepime 2g IV q12h OR Piperacillin/Tazobactam 4.5g IV q8h | Meropenem 1g IV q8h OR Imipenem 500mg IV q6h | Current evidence suggests that carbapenems, 4 th generation cephalosporins or Piperacillin/Tazobactam are equally effective in treatment of septic shock If melioidosis cannot be ruled out, carbapenem should be used as the empirical agent |
| <i>Methicillin-resistant S. Aureus</i> <i>Penicillin-resistant S. Pneumoniae</i> <i>Ampicillin-resistant Enterococci</i> | PLUS OPTIONAL Vancomycin ¹ 1g IV q12h | | Empirical use of Vancomycin ¹ is only justified in areas with high endemic levels of MRSA or high levels of penicillin-resistant <i>S. pneumoniae</i> |
| <i>Candida</i> | PLUS OPTIONAL Fluconazole 400-800mg IV q24h | PLUS OPTIONAL Amphotericin B 0.6-1.0mg/kg IV q24h | Empirical antifungal agents should not be used on a routine basis <i>Reference 1, 2</i> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|--|
| | Preferred | Alternative | |
| B. Severe Community-Acquired Pneumonia Requiring Mechanical Ventilation | | | |
| Severe community-acquired pneumonia requiring mechanical ventilation <i>S. Pneumoniae</i> <i>H. Influenzae</i> <i>S. Aureus</i> <i>K. Pneumoniae</i> <i>M. Pneumoniae</i> <i>L. Pneumophila</i> <i>C. Pneumoniae</i> <i>*B. Pseudomallei</i> | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q24h PLUS Erythromycin 500mg IV q6h OR Azithromycin 500mg IV q24h *If risk factors present, consider Ceftazidime (Please refer to Page 95 (LRTI)) | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h PLUS Erythromycin 500mg IV q6h OR Azithromycin 500mg IV q24h | Reference 3, 4, 5 |
| C. Severe Nosocomial Pneumonia Requiring Mechanical Ventilation (Including Ventilator-Associated Pneumonia) | | | |
| Nosocomial pneumonia requiring mechanical ventilation (including VAP) Low risk for infection with multi-drug resistant (MDR) organisms - < 5 days <i>S. Pneumoniae</i> <i>H. Influenzae</i> <i>S. Aureus</i> <i>E. Coli</i> <i>K. Pneumoniae</i> <i>Enterobacter spp.</i> <i>Proteus spp.</i> <i>Serratia Marcescens</i> | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q24h OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV q6h | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h | <i>S. aureus</i> is more common in diabetes mellitus, head trauma Monotherapy is recommended for early onset HAP/VAP/HCAP Reference 6, 7 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|---|
| | Preferred | Alternative | |
| High risk for infection with multi-drug resistant (MDR) organisms <i>P. Aeruginosa</i> <i>Acinetobacter spp.</i> <i>K. Pneumoniae (ESBL)</i> Methicillin-resistant <i>S. Aureus</i> | Piperacillin/Tazobactam 4.5g IV q6h OR Cefepime 2g IV q12h PLUS Amikacin ¹ 15mg/kg/24h IV OR Ciprofloxacin 400mg IV q8h Cefoperazone/Sulbactam 2g IV q12h Meropenem 1g IV q8h OR Imipenem 500mg IV q6h PLUS (if MRSA is suspected) Vancomycin ¹ 1g IV q12h | Imipenem 500mg IV q6h OR Meropenem 1g IV q8h PLUS Amikacin ¹ 15mg/kg/24h IV OR Ciprofloxacin 400mg IV q8h <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV q6h | Use combination therapy if MDR pathogen is suspected Aminoglycoside can be stopped after 5-7 days in patients on combination therapy who are responding to treatment |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

References:

- | | | |
|--|--|---|
| 1. Crit Care Med 2003; 31:1250-1256 | 3. Am J Respir Crit Care Med 2002, 166:717-723 | 6. Am J Respir Crit Care Med. 2005; 171:388-416 |
| 2. Crit Care Med 2004; 32(11)S495 S512 | 4. Clin Infect Dis 2003; 37:1405-33 | 7. Curr Anaes and Crit Care 2005;16:209-219 |
| | 5. Curr Opin Crit Care 2004; 10:59-64 | |

OBSTETRICS & GYNAECOLOGICAL INFECTIONS

A. OBSTETRICS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|-----------------|
| | Preferred | Alternative | |
| Intrapartum prophylaxis for GBS (<i>Group B. Streptococcus</i>), positive mothers | Intrapartum Benzylpenicillin 5 mega units IV followed by 2.5 mega units IV q4h | Intrapartum <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV followed by 750mg q8h OR Ampicillin 2g IV as loading dose followed by 1g IV q4h, to stop after delivery If allergic to penicillin (non-anaphylactic): Cefuroxime 1.5g IV followed by 750mg IV q6-8h If life threatening (anaphylactic): Erythromycin 500mg IV q6h, if susceptible | RCOG Guidelines |
| PPROM (Preterm Premature Rupture of Membranes) <i>Mixed</i> | EES 400mg PO q12h for 10 days | Amoxycillin 500mg PO q8h OR Cefuroxime 250mg PO q12h for 10 days | RCOG guidelines |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|-----------------|
| | Preferred | Alternative | |
| Chorioamnionitis <i>Gram (-) rods/ Gram (+) coccus/ Anaerobes</i> | <i>2nd or 3rd gen. Cephalosporins, e.g.</i> Cefuroxime 750mg IV q8h OR Cefoperazone 1g IV q12h PLUS Metronidazole 500mg IV q8h for 3 days followed by oral treatment for 7 days | Ampicillin 1g IV q6h PLUS Metronidazole 500mg IV q8h PLUS Gentamicin ¹ 5mg/kg IV q24h for 7 days | RCOG Guidelines |
| Puerperal Sepsis Mixed:- <i>Streptococcus Staphylococcus Gram Negative Bacilli Anaerobes</i> | OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV q8h for 3 days followed by oral treatment for 7 days | | |

B. GYNAECOLOGY

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| Pelvic Inflammatory Disease | | | |
| <p><i>C. Trachomatis</i> <i>Bacteroides sp.</i> <i>Gardnerella Vaginalis</i> <i>E. Coli</i> <i>Streptococcus</i> <i>Coagulase-negative</i> <i>Staphylococcus</i></p> | <p>IV THERAPY (for moderate to severe disease): 2nd or 3rd gen. Cephalosporins, e.g. Cefuroxime 750mg IV q8h OR Ceftriaxone 2g IV q24h PLUS Doxycycline 100mg PO q12h PLUS Metronidazole 400mg PO q8h Duration of treatment is 14 days OUTPATIENT THERAPY (for mild disease): Cefuroxime 250-500mg PO q12h PLUS Doxycycline 100mg PO q12h PLUS Metronidazole 400mg PO q8h If gonococcal infection suspected, Refer to Page 100 (Sexually Transmitted Infections)</p> | <p><i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5-3g IV q6h PLUS Doxycycline 100mg PO q12h</p> | <p>Antibiotic should be changed accordingly after C&S results available</p> |

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| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|---|
| | Preferred | Alternative | |
| Vaginitis | | | |
| Bacterial Vaginosis <i>Gardnerella Vaginalis</i> | Metronidazole 400mg PO q12h for 7 days | Clindamycin 300mg PO q12h for 7 days | <ul style="list-style-type: none"> - Metronidazole is best avoided in the first trimester of pregnancy - In pregnancy, treatment is indicated for symptomatic disease and asymptomatic women at high risk for preterm delivery - Avoid alcohol (antabuse effect) |
| Candidiasis <i>Candida Albicans</i> | Clotrimazole 500mg as a single vaginal pessary (stat dose) Clotrimazole 200mg as vaginal pessary for 3 nights | Tinidazole 500mg PO q12h for 5 days OR Tinidazole 2g PO stat | Metronidazole/Tinidazole are best avoided in the first trimester of pregnancy |
| Trichomoniasis <i>Trichomonas Vaginalis</i> | Metronidazole 200mg PO q8h for 7 days OR Metronidazole 400mg PO q12h for 7 days OR Metronidazole 2g PO stat | <u>In pregnancy:</u> Clotrimazole pessary 100mg daily for 7 days, but systemic treatment will ultimately be necessary to eradicate the infection | Avoid alcohol (antabuse effect) |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|----------|
| | Preferred | Alternative | |
| Septic Miscarriage | | | |
| <i>Streptococcus</i> <i>Staphylococcus</i> <i>Gram Negative Bacilli</i> <i>Anaerobes</i> | 2 nd or 3 rd gen. Cephalosporins, e.g. Cefuroxime 750mg IV q8h OR Cefoperazone 1g IV q12h PLUS Metronidazole 500mg IV q8h for 3 days followed by oral treatment for 7 days OR β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g IV q8h for 3 days followed by oral treatment for 7 days | Ampicillin 500mg IV q6h PLUS Metronidazole 500mg IV q8h PLUS Gentamicin ¹ 5mg/kg IV q24h for 7 days | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

OCULAR INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Blepharitis <i>Staph. Aureus</i> <i>Staph. Epidermidis</i> | Chloramphenicol 1% eye ointment applies q6h to lid margins Duration as required | Chlortetracycline 1% eye ointment apply q6h OR Fusidic Acid 1% eye ointment apply q6h | In resistant cases, Doxycycline 100mg PO q24h or Tetracycline 250mg PO q6h for 2 to 4 weeks or as necessary Incision and curettage may be required |
| Internal Hordeolum with Secondary Infection <i>Staph. Aureus</i> | Chloramphenicol 1% eye ointment apply q6h for 1 week | Chlortetracycline 1% eye ointment apply q6h for 1 week | Topical antibiotics NOT indicated unless keratitis is present. Topical saline drops for toilet |
| External Hordeolum (stye) <i>Staph. Aureus</i> | Chloramphenicol 1% eye ointment apply q6h for 1 week | Chlortetracycline 1% eye ointment apply q6h for 1 week | |
| Gonococcal Conjunctivitis (including neonates) <i>Neisseria Gonorrhoea</i> | Needs systemic therapy Refer to Page 100 (Sexually Transmitted Infections) & Page 177 (Neonatal Infections) | | Topical antibiotics NOT indicated unless keratitis is present. Topical saline drops for toilet |
| Chlamydial Conjunctivitis (including neonates) <i>Chlamydial Trachomatis</i> | Needs systemic therapy Refer to Page 99 (Sexually Transmitted Infections) & Page 177 (Neonatal Infections) | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| Adult Inclusion Conjunctivitis or Trachoma <i>Chlamydia Trachomatis</i> | Needs systemic therapy Refer to Page 100 (Sexually Transmitted Infections) and Page 177 (Neonatal Infections) | | Exclude other STD's. Treat sexual partners |
| Bacterial Conjunctivitis <i>Staph Aureus, Strep Pneumonia, H. Influenzae</i> | Chloramphenicol 0.5% eye drop apply q2-4h for 1 week | Gentamicin 0.3% eye drop apply q2-4h for 1 week | |
| Bacterial Keratitis Mixed Growth/ No Growth | *Cefuroxime 5% eye drop apply hrly PLUS *Gentamicin 0.9% or 1.4% eye drop apply hrly | Ciprofloxacin 0.3% eye drop apply hrly | In severe keratitis, commence a loading dose of one drop every 15 minutes for 3 hours followed by hourly drops around the clock. Taper based on clinical response *prepare ready to use extemporaneous by using injectable forms |
| Bacterial Keratitis Gram-Positive Cocci Gram-Negative Rods Gram-Negative Cocci | *Cefuroxime 5% eye drop apply hrly **Gentamicin 0.9% or 1.4% eye drop apply hrly *Ceftazidime 5% eye drop apply hrly | Ciprofloxacin 0.3% eye drop apply hrly *Ceftazidime 5% eye drop apply hrly Ciprofloxacin 0.3% eye drop apply hrly | *Vancomycin 5% eye drop may be indicated for MRSA *Cefuroxime 5% eye drop, Ceftazidime 5% eye drop, Vancomycin 5% eye drop - prepare ready to use extemporaneous by using injectable forms. **Gentamicin 0.9% & 1.4% eye drop - prepare Fortified Gentamicin Eye Drops |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|--|
| | Preferred | Alternative | |
| Contact Lens Related Bacterial Keratitis <i>Pseudomonas</i> | **Gentamicin 0.9% or 1.4% eye drop apply q1-2h PLUS *Ceftazidime 5% eye drop apply q1-2h | Ciprofloxacin 0.3% eye drop apply hrly | *Ceftazidime 5% eye drop - prepare ready to use extemporaneous by using injectable forms **Gentamicin 0.9% & 1.4% eye drop - prepare Fortified Gentamicin Eye Drops |
| Gonococcal Keratoconjunctivitis <i>Neisseria Gonorrhoea</i> | Ocular Treatment: Ciprofloxacin 0.3% eye drop apply hrly Refer to Page 100 (Sexually Transmitted Infections) & Page 177 (Neonatal Infections) | Ocular Treatment: *Ceftazidime 5% eye drop apply hrly | *Ceftazidime 5% eye drop - prepare ready to use extemporaneous using injectable forms |
| Herpes Simplex Keratitis <i>Herpes Simplex</i> Type 1 and 2 | Acyclovir 3% eye ointment apply 5 times/day until the epithelium heals then taper | | Acyclovir 3% eye ointment should not be used for more than 6 weeks due to toxicity |
| Herper Zoster Ophthalmicus <i>Herpes Zoster Virus</i> | Needs systemic therapy Refer to Page 108 (Skin & Soft Tissue Infections) | | |
| Acanthamoeba Keratitis <i>Acanthamoeba sp.</i> | *Chlorhexidine 0.02% eye drop PLUS Neomycin 0.5% eye ointment apply hrly | | *Chlorhexidine 0.02% eye drop prepare ready to use extemporaneous |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|--|
| | Preferred | Alternative | |
| Fungal Keratitis Filamentous Fungi/Yeast | ***Fluconazole 0.2% eye drop q1-2h PLUS/MINUS Amphotericin B 0.15%-0.2% eye drop q1-2h PLUS Fluconazole 200mg PO q24h | **Natamycin 5% q1-2h for 3-4 days, then q3-4h for 2-3 weeks PLUS Amphotericin B 0.15% to 0.2% eye drop q1-2h PLUS Ketoconazole 200mg PO q24h | Treatment depending on the severity of the infection **requires DG approval ***Fluconazole 0.2% eye drop - prepare ready to use extemporaneous |
| Dacryocystitis <i>Strep Pneumonia, Staph Aureus</i> <i>Gram -ve Anaerobes</i> | Amoxicillin 500mg PO q8h for at least 5 days | Cephalexin 500mg PO q6h for at least 5 days | Consider corresponding intravenous antibiotics in severe infections |
| Preseptal Cellulitis <i>Strep Pneumoniae, Staph Aureus, Streptococcus sp.</i> | Cloxacillin 500mg-1g PO q6h for 5 days | Amoxicillin 500mg PO q8h | Consider corresponding intravenous antibiotics: - in severe infections - if secondary to sinusitis |
| Ocular Toxoplasmosis <i>Toxoplasma Gondii</i> | Needs systemic therapy Refer to Page 53 (Human Immunodeficiency Virus) | | |
| Acute Retinal Necrosis Herpes Simplex | Needs systemic therapy Refer to Page 53 (Human Immunodeficiency Virus) | | Systemic steroid is indicated depending on location or severity of the infection |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|---|
| | Preferred | Alternative | |
| CMV Retinitis Cytomegalovirus | Needs systemic therapy Refer to Page 53 (Human Immunodeficiency Virus) Ocular Treatment: Intravitreal Ganciclovir 2mg/0.1ml (weekly) - (Prefer: Ganciclovir implant: 4.5g - if available) | Ocular Treatment: Intravitreal *Foscarnet 2.4mg/0.1ml (1-2 weekly) | Intravitreal to be repeated according to clinical response *Requires DG approval To continue until CD4 count is > 150 cell/mm ³ |
| Ocular Syphilis <i>Treponema Pallidum</i> | Needs systemic therapy Refer to Page 100 (Sexually Transmitted Infections) | | Referral to neurologist prior to starting treatment |
| Ocular Tuberculosis <i>Mycobacterium Tuberculosis</i> | Needs systemic therapy Refer to Page 143 (Tuberculosis Infections) | | Systemic steroid may be indicated but is only for <ul style="list-style-type: none"> - non-active systemic TB - severe ocular inflammation and vision threatening condition |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|--|
| | Preferred | Alternative | |
| Orbital Cellulitis/abscess <i>Strep Pneumoniae, Staph Aureus, Streptococcus sp. Gram -ve Anaerobes</i> | Cefuroxime 750mg-1.5g q8h OR Cloxacillin 1-2g IV q6h PLUS Ceftriaxone 1-2g IV q24h If sinusitis is suspected as the cause ADD: Initial Metronidazole 15mg/kg IV infused over 1 hr Anaerobic infection: maintenance, 7.5mg/kg/hr IV q6h, starting 6 hrs after initial dose; maximum 4g/day Treat for 5 days | | Treat underlying cause (e.g. sinusitis) In orbital abscess, surgical drainage is often necessary References: 1. Medical and Surgical Management of Orbital Cellulitis Michael T. Yen, M.D. Contemporary Ophthalmology, June 2005, Vol. 4, No. 11, Page 1-6 2. Role of Inflammation in Orbital Cellulitis Carolyn E. Kloek, MD Peter A.D. Rubin, MD Manuscript on Role of Inflammation in Orbital Cellulitis Page 57-68 |
| Post Operative Fungal Endophthalmitis | Intravitreal Amphotericin B 0.005mg in 0.1ml | *Intravitreal Miconazole: (0.01mg in 0.1ml) | *Requires DG approval CPG for Management of Post-Operative Endophthalmitis, Ministry of Health Malaysia, August 2006 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| <p>Post Operative Bacterial Endophthalmitis <i>Staphylococcus Epidermidis</i> <i>Staphylococcus Aureus</i> <i>Pseudomonas Aeruginosa</i>, <i>Bacteroids Species</i> <i>Streptococcus Pneumoniae</i>, <i>Alpha-Haemolytic Streptococci</i></p> | <p>Intravitreal antibiotic injections: Vancomycin 1-2mg in 0.1ml and Cefazidime 2mg in 0.1ml</p> <p>If suspicious of fungal endophthalmitis, ADD: Intravitreal Amphotericin B 0.005mg in 0.1ml</p> <p>ALSO consider in culture negative cases with poor clinical response: Ciprofloxacin 250mg PO q12h</p> | <p>Intravitreal antibiotic injections Vancomycin 1-2mg in 0.1ml and Amikacin 0.4mg in 0.1ml</p> <p>Clarithromycin 250-500mg PO q12h for 7-14 days</p> | <ol style="list-style-type: none"> 1. Begin intensive topical antibiotics and topical steroid soon after intravitreal antibiotic injection 2. Systemic antibiotics for severe, virulent endophthalmitis 3. Oral prednisolone to be considered and may be given 24 hours following intravitreal antibiotics injection 4. Review antibiotic regimen after microbiology results 5. Repeat intravitreal antibiotics after 48 to 72 hours if indicated <p>EARLY REFERRAL TO A VITREORETINAL CENTER IS RECOMMENDED</p> <p><i>CPG for Management of Post-Operative Endophthalmitis, Ministry of Health Malaysia, August 2006</i></p> |

ORAL/DENTAL INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|-------------|--|
| | Preferred | Alternative | |
| 1. ANTIMICROBIAL USE FOR BACTERIAL INFECTIONS | | | |
| A. Infections of the Teeth and Supporting Structures | | | |
| Reversible/Irreversible Pulpitis | Systemic antibiotic use not recommended | | Endodontic treatment and symptomatic relief of pain <i>Cochrane Database of Systematic Reviews</i> 2005, Issue 2. Art. No: CD004969. DOI: 10.1002/14651858.CD004969.pub2 |
| Localised Dentoalveolar Abscess | Systemic antibiotic use not recommended | | Incision and Drainage and management of cause of abscess and symptomatic relief of pain <i>J Can Dent Assoc</i> 2003 Nov 69(10):660 |
| Dry Socket | Systemic antibiotic use not recommended | | Local treatment with saline irrigation and antiseptic/analgesic dressings and symptomatic relief of pain <i>Med Oral Patol Oral Cir Bucal</i> 2005; 10:77-85 |
| Localised Pericoronitis | Systemic antibiotic use not recommended | | Local treatment with antiseptic irrigation and mouthwash and symptomatic relief of pain <i>J Clin Microbiol.</i> 2003; 41(12):5794-7 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|---|
| | Preferred | Alternative | |
| Chronic Gingivitis | Systemic antibiotic use not recommended | | 1st line treatment - Mechanical plaque control 2nd line treatment - Antimicrobial mouth rinse <i>Clinical Periodontology - 9th ed. 2002</i> |
| Chronic Periodontitis | Systemic antibiotic use not recommended | | 1st line treatment - Mechanical plaque control <i>Eur J Prosthodont Restor Dent. 2004 Jun; 12(2): 63-9</i> CPG Management of chronic periodontitis 2005 MOH, Malaysia |
| Aggressive Periodontitis <i>A. Actinomycetemcomitans,</i> <i>P. Gingivalis,</i> <i>Tannerella Forsythensis,</i> <i>P. Intermedia,</i> <i>Spirochaetes</i> | *Amoxicillin 500mg PO q8h PLUS *Metronidazole 400mg PO q8h | *Doxycycline 100mg PO q12-24h OR *Clindamycin 150-300mg PO q6h | Antibiotics are not used alone but are used as an adjunct to scaling and root debridement <i>J Periodontol 2004; 75: 1553-1565</i> <i>J Clin Periodontol. 2005 Oct; 32(10): 1096-107</i> <i>Evid Based Dent. 2006; 7(3): 67.</i> *Treatment depending on severity of infection |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|--|
| | Preferred | Alternative | |
| Localised Periodontal Abscess | Systemic antibiotic use not recommended | | <p>Incision and Drainage and management of cause of abscess and symptomatic relief of pain</p> <p><i>CPG = Management of periodontal abscess - MOH, Malaysia April 2004</i></p> |
| B. Infections of the Jaws | | | |
| <p>Osteomyelitis of the jaws of dental origin</p> <p><i>Different organisms may be involved</i></p> | <p>For acute cases, start with: Phenoxymethylpenicillin 250-500mg PO q6h</p> <p>OR *Benzylpenicillin 1-2 mega units IV q6h</p> | <p>*Clindamycin 150-300mg PO q6h</p> <p>OR *Clindamycin 150-450mg IV q6h</p> | <p>Culture and sensitivity is necessary</p> <p>For chronic cases, start with surgical treatment first. Antibiotics only when causative organisms are identified</p> <p>*Treatment depending on severity of infection</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| C. Spreading Infections and Infections of Fascial Spaces (with/without Systemic Signs) | | | |
| <p><u>Cellulitis ± Abscess of dental origin</u> <i>Viridans Streptococci, Staphylococci, Prevotella, Peptostreptococcus</i></p> <p><u>Surgical site infection & Traumatic wound infection (Infection is usually by endogenous organisms rather than exogenous)</u> <i>Viridans Streptococci Staphylococci Prevotella, Peptostreptococcus, Eubacterium, and Fusobacterium</i></p> | <p>Benzylpenicillin 2-4 mega units IV stat then 1-2 mega units IV q4-6h* PLUS/MINUS Metronidazole 500mg IV q8h (or 1g q12h)*</p> <p>PLUS Cloxacillin 500mg-1g IV q6h (in skin involvement - if Staph. expected) OR Clindamycin 150-450mg IV q6h*</p> <p><u>Oral administration:</u> Amoxycillin 250-750mg PO q8h* PLUS/MINUS Metronidazole 400mg PO q8-12h*</p> <p>OR Clindamycin 150-450mg PO q6h*</p> | <p><i>β-lactam/β-lactamase inhibitors, e.g. Amoxycillin/Clavulanate 1.2g IV q6-8h (not more than 1.2g in a single dose - max 7.2g daily)*</i></p> <p>OR Cefuroxime 750mg-1.5g IV q8h PLUS/MINUS Metronidazole 500mg IV q8h (or 1g q12h)* OR If not responding to above antibiotics, 3rd gen. Cephalosporins, e.g. Ceftriaxone 1-2g IV q24h* (may be given up to 4g per day)</p> <p><u>Oral administration:</u> <i>β-lactam/β-lactamase inhibitors, e.g. Amoxycillin/Clavulanate 625mg PO q12h. If severe, 625mg PO q8h*</i></p> <p>OR Cefuroxime 250-500mg PO q12h* D. Post Implant Infections ("Periimplantitis")</p> | <p><i>J Oral Maxillofac Surg 2006; 64:1377-1380</i></p> <p><i>Asian J Oral Maxillofac Surg 2005; 17:168-172</i></p> <p><i>Antimicrobial Agents and Chemotherapy, 1995; 39(10):2243-47</i></p> <p><i>Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2000; 90:600-8</i></p> <p><i>Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2005; 100:550-8</i></p> <p><i>J Craniomaxillofac Surg 1995; 23:38-41</i></p> <p><i>Int J Antimicrobial Agents 2000; 15:1-9</i></p> <p><i>Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2004; 98:398-408</i></p> <p><i>J Craniomaxillofac Surg. 2005 Feb 33(1):24-9</i></p> <p><i>Journal of Emergency Medicine, 1999; 17(1):189-195</i></p> <p>*Treatment depending on severity of infection</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|--|
| | Preferred | Alternative | |
| D. Post Implant Infections (“Periimplantitis”) | | | |
| <i>Actinomyces sp.</i> <i>Eubacterium sp.</i> <i>Propionibacterium sp.</i> <i>Lactobacillus sp.</i> <i>Veillonella sp.</i> <i>P. Gingivalis</i> <i>Prevotella Intermedia</i> <i>F. Nucleatum</i> | Amoxicillin 250-500mg PO q8h* PLUS Metronidazole 200-400mg PO q8h* | Doxycycline 100mg PO q12-24h* OR Clindamycin 150-300mg PO q6h* | Bacteria associated with periimplantitis are extremely resistant to antibiotics Antibiotics are not used alone but are used as an adjunct to local mechanical and chemical debridement Also irrigation with Chlorhexidine and optimal oral hygiene by patient <i>Oral Surg Oral Med Oral Pathol Oral Radiol Endod</i> 2005; 100:550-8 <i>Periodontol</i> 2000-2002; 28:177-89 *Treatment depending on severity of infection |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|-------------|---|
| | Preferred | Alternative | |
| 2. ANTIMICROBIAL USE FOR FUNGAL INFECTIONS | | | |
| A. Oral Candidiasis | | | |
| Acute Pseudomembranous Candidiasis <i>Candida sp.</i> | Nystatin (topical) 500,000 units q6h for up to 4 weeks Systemic antifungal for severe infections, severely immunocompromised patients and for infections resistant to topical antifungal: Fluconazole 50-100mg PO/IV q24h for 2 weeks OR Itraconazole 100mg PO q24h for 2 weeks | | Use chlorhexidine mouthwash as adjunct <i>J Prosthetic Dent.</i> 1989; 61:699 <i>J Biol Buccale</i> 1992; 20:45 <i>Oral Surg. Oral Med. Oral Pathol.</i> 1992; 73 (6):682-689 <i>Crit. Rev. Oral Biol. Med.</i> 2000; 11:172-198 <i>Clin. Infect. Dis.</i> 1994; 18(3):298-304 |
| Hyperplastic Candidiasis (Candidal Leukoplakia) | Nystatin (topical) 500,000 units q6h for up to 4 weeks <u>Systemic antifungal for infections resistant to topical antifungal:</u> Fluconazole 50-100mg PO/IV q24h for 2 weeks OR Itraconazole 100mg PO q24h for 2 weeks | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|-------------|---|
| | Preferred | Alternative | |
| Candida-associated denture stomatitis with or without angular cheilitis | Local measures first Consider antifungal if local measures fail Nystatin (topical) 500,000 units q6h for up to 4 weeks | | |
| 3. ANTIMICROBIAL USE FOR VIRAL INFECTIONS | | | |
| Primary Herpes Simplex Infection (Primary herpetic gingivostomatitis) <i>Herpes Simplex Virus</i> | Symptomatic treatment only in most cases For severe infections may consider: For adult & healthy patients Acyclovir 200-400mg PO 5 times daily for 5-7 days For immunocompromised patients: Acyclovir 250mg/m ² IV q8h | | <i>J Am Acad Dermatol</i> 1988 January; 18 (1 Part 2):176-179 <i>Drug Intell Clin Pharm</i> 1985 July-August; 19 (7-8):518-524 |
| Secondary Herpes Simplex Infection <i>Herpes Simplex Virus</i> | Acyclovir 5% cream to be applied q6h For external use only | | <i>J Infect Dis</i> 1990; 161 (2):185-190 <i>JAMA</i> 1988; 260 (11):1597-1599 <i>Ann Intern Med</i> 1993; 118:268-272 |

RESPIRATORY INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| A. UPPER RESPIRATORY TRACT INFECTIONS | | | |
| 1. Throat And Upper Respiratory | | | |
| Acute Tonsillitis Acute Pharyngitis <i>Strep. Pyogenes, Group A Beta Hemolytic Streptococcus</i> | Phenoxymethylpenicillin 250-500mg PO q8h for 10 days OR (in penicillin allergic patients) EES 400mg PO q12h for 10 days | | Antibiotics should be prescribed in suspected/proven bacterial infections, only as sore throats are common viral in origin. In severe cases, start with parenteral penicillin In infections of the throat and tonsil due to mononucleosis, Ampicillin/ Amoxicillin frequently precipitates a non-allergic rash (this is not an indication of Penicillin hypersensitivity) <i>Practice Guidelines for the Diagnosis and Management of Group A Streptococcal Pharyngitis. Clinical Infectious Diseases 2002</i> |
| Acute Peritonsillar Abscess <i>Streptococcus Pyogenes, Fusobacterium</i> | Benzylpenicillin 2-4 mega units IV q6h followed by Phenoxymethylpenicillin 500mg PO q6h for 10 days PLUS/MINUS Metronidazole 500mg IV q8h followed by Metronidazole 400mg PO q8h | β -lactam/ β -lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV q8h followed by Amoxicillin/Clavulanate 625mg PO q12h for 10 days OR Ampicillin/Sulbactam 1.5g IV q8h followed by Ampicillin/Sulbactam 375mg PO q12h for 10 days | Abscess to be drained |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Diphtheria <i>Corynebacterium Diphtheriae</i> | Benzylpenicillin 50,000 units/kg/24h IV for 5 days followed by Phenoxymethylpenicillin 50mg/kg/24h PO for 5 days | | Antitoxin and supportive treatment are critical in management. Antibiotic is not the mainstay of treatment |
| Acute Epiglottitis <i>Haemophilus Influenzae Type b,</i> <i>Streptococcus Pneumoniae</i> | <i>2nd or 3rd gen. Cephalosporins, e.g.</i> Cefuroxime 750mg IV q8h, may be followed by Cefuroxime 250mg PO q12h for total of 14 days OR Ceftriaxone 1g IV q24h | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h; may be followed by Amoxicillin/Clavulanate 625mg PO q12h for 14 days OR Chloramphenicol 500mg-1g IV q6h, may be followed by 250-500mg PO q12h for 14 days | Urgent hospitalisation. May present with life threatening upper airway obstruction, especially in paediatrics |
| Deep Neck Abscess <i>Polymicrobial,</i> <i>S. Aureus, Strep. sp.,</i> <i>Bacteroides sp.</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h; OR Cefuroxime 750mg IV q8h PLUS Metronidazole 500mg IV q8h for at least 7 days | <i>2nd or 3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV q24h PLUS Metronidazole 500mg IV q8h for at least 7 days | Abscess needs to be drained |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| 2. Rhinology | | | |
| Acute Bacterial Rhinosinusitis (ABRS) <i>Streptococcus Pneumoniae,</i> <i>Haemophilus Influenzae, Moraxella Catarrhalis</i> | Amoxicillin 500mg PO q8h for 7-14 days OR (<i>in penicillin allergic patients</i>) EES 400mg PO q12h for 7-14 days | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 625mg PO q12h for 7-14 days OR (<i>in penicillin allergic patients</i>) Cefuroxime 500mg PO q12h for 7-10 days OR <i>Macrolides, e.g.</i> Azithromycin 500mg PO q24h for 3 days | The Cochrane Database of Systematic Reviews 2004, Issue 1 |
| Subperiosteal Abscess Secondary to ABRS <i>S. Pneumoniae,</i> <i>S. Pyogenes,</i> <i>H. Influenzae</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h OR Ampicillin/Sulbactam 1.5g IV q8h for 10-14 days OR Cefuroxime 750mg IV q8h for 10-14 days | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV q24h for at least 10 days | Abscesses must be drained |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| 3. Otology | | | |
| Acute Otitis Media <i>Streptococcus Pneumoniae,</i> <i>Haemophilus Influenzae</i> | Amoxicillin 500mg PO q8h for 7 days OR (in penicillin allergic patients) EES 400mg PO q12h for 7 days | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 625mg PO q12h for 7 days | Myringotomy may be required in cases of impending rupture of tympanic membrane |
| Malignant Otitis Externa/ Necrotizing Otitis Externa <i>Pseudomonas Aeruginosa</i> | Ciprofloxacin 400mg IV q12h followed by Ciprofloxacin 500-750mg PO q12h for 6 weeks | | Aural toileting required. Surgical debridement normally required |
| Acute Mastoiditis/ Mastoid Abscess <i>S. Pneumoniae, S. Pyogenes, Coag.-negative Staph, S. Aureus, Proteus and Bacteroides sp.</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h followed by Amoxicillin/Clavulanate 625mg PO q12h for 7-14 days OR Ampicillin/Sulbactam 1.5g IV q8h followed by Ampicillin/Sulbactam 375mg PO q12h OR Cefuroxime 750mg IV q8h followed by Cefuroxime 250mg PO q12h | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IV q24h for 7-14 days | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|---|
| | Preferred | Alternative | |
| Acute Diffuse Otitis Externa <i>P. aeruginosa and Staph Aureus</i> | Framycetin Sulphate 0.5%, Dexamethasone 0.05% & Gramicidin 0.005% ear drop 2-3 drops 3-4 times/day for 7 days | Ofloxacin 0.3% otic solution 6-10 drops q12h for 10 days | Aural toileting required in discharging ears The dosage should be reduced appropriately for children |
| Chronic Suppurative Otitis Media <i>P. aeruginosa, Staph Aureus and Epidermidis, Proteus sp.</i> | Ofloxacin 0.3% otic solution 6-10 drops twice a day for 10 days OR Framycetin Sulphate 0.5%, Dexamethasone 0.05% & Gramicidin 0.005% ear drop 2-3 drops 3-4 times/day for 7 days | | Aural toileting required in discharging ears The dosage should be reduced appropriately for children |
| Otomycosis <i>Aspergillus sp.</i> | Kenacomb Otic Drops (Triamcinolone Acetonide 0.9mg/ml, Neomycin base 2.25mg/ml, Nystatin 90,000 units/ml and Gramicidin 0.225mg/ml) 2-3 drops 2-3 times/day for 2 weeks | | Aural toileting required and tympanic membrane needs to be inspected prior to administration In paediatric patient, medication should be monitored, least amount and shortest duration compatible with effective therapeutic regimen |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|--|
| | Preferred | Alternative | |
| B. LOWER RESPIRATORY TRACT INFECTIONS | | | |
| 1. Community Acquired Penumonia (CAP) | | | |
| Mild CAP (out-patient) | | | |
| a. No comorbidity <i>Streptococcus Pneumonia</i> <i>Mycoplasma Pneumoniae</i> | No recent antibiotic therapy EES 800mg PO q12h for 1 week OR Amoxycillin 500mg PO q8h for 1 week | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxycillin/Clavulanate 625mg PO q12h for 1 week OR Ampicillin/Sulbactam 375mg PO q12h for 1 week OR Doxycycline 100mg PO q12h for 1 week | |
| b. Presence of comorbidity or History of recent antibiotic therapy (2 months) <i>Streptococcus Pneumoniae</i> <i>Mycoplasma Pneumoniae</i> <i>Haemophilus Influenzae</i> | Azithromycin 500mg PO q24h for 3 days OR EES 800mg PO q12h for 1 week PLUS <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxycillin/Clavulanate 625mg PO q12h for 1 week | Levofloxacin 500mg PO q24h for 1 week | Conservative use of quinolone is recommended to minimise resistant pathogen. Use when patients failed first line regimens or allergic to alternative |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| <p>Moderate & Severe CAP (not requiring mechanical ventilation) <i>Streptococcus Pneumoniae</i> <i>Mycoplasma Pneumoniae</i> <i>Haemophilus Influenzae</i> <i>Klebsiella Pneumoniae</i> <i>Legionella</i> <i>Staphylococcus Aureus</i> Other Gram Negative Bacilli - <i>Enterobacter</i> - <i>Escherichia Coli</i></p> | <p>Azithromycin 500mg IV/PO q24h OR Erythromycin 500mg IV q6h/EES 800mg PO q12h</p> <p>PLUS <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1-2g IV q24h OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> (Amoxicillin/Clavulanate OR Ampicillin/Sulbactam)</p> <p>Duration: 1 week</p> | <p>Levofloxacin 500mg IV/PO q24h for 1 week</p> | <p>Empirical therapy for melioidosis should be considered if patient has diabetes mellitus</p> <p>Conservative use of quinolone is recommended to minimise resistant pathogen. Use when patients failed first line regimens or allergic to alternative</p> |
| <p><i>Pseudomonas Infection</i></p> | <p>Piperacillin/Tazobactam 4.5g IV q8h for 1 week OR Cefepime 2g IV q12h for 1 week</p> <p>PLUS Gentamicin¹ 5mg/kg IV q24h PLUS Azithromycin 500mg IV q24h for 1 week</p> <p>For severe CAP Requiring Mechanical Ventilation. Refer to Page 68 (Infections In Intensive Care Units)</p> | <p>Piperacillin/Tazobactam 4.5g IV q8h for 1 week OR Cefepime 2g IV q12h for 1 week</p> <p>PLUS Ciprofloxacin 500mg IV q12h for 1 week</p> | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|----------|
| | Preferred | Alternative | |
| <p>2. Lung Abscess Organisms likely to be involved are anaerobes (34%), Gram positive cocci (26%), <i>Klebsiella Pneumoniae</i> (25%), <i>S. Milleri</i> (16%), <i>Nocardia</i> (3%).</p> <p>If suspect melioidosis</p> <p><i>Staphylococcus Aureus</i> (e.g. among IVDU)</p> | <p>3rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV q24h</p> <p>PLUS Metronidazole 500mg IV q8h followed by 400mg PO q8h for 4-6 weeks</p> <p>Ceftazidime 2g IV q8h for 10-14 days</p> <p>Cloxacillin 2g IV q4-6h for 2-4 weeks</p> | <p>Piperacillin/Tazobactam 4.5g IV q8h for 4-6 weeks</p> | |
| <p>3. Empyema Always investigate as per pleural effusion. Drainage via chest tube required. Tuberculosis must be excluded</p> | | | |
| <p>Empyema</p> <p>If Anaerobes isolated/suspected: <i>Strep Milleri</i> <i>Enterobacteriaceae</i> <i>Bacteroides sp.</i></p> <p>If <i>Staphylococcus Aureus</i> Isolated</p> | <p>3rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV q24h</p> <p>OR Cefotaxime 1g IV q8h</p> <p>3rd gen. Cephalosporins, e.g. Ceftriaxone 2g IV q24h</p> <p>OR Cefotaxime 1g IV q8h</p> <p>PLUS Metronidazole 500mg IV q8h</p> <p>Cloxacillin 2g IV q4h</p> | <p>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV q8h</p> <p>OR Ampicillin/Sulbactam 1.5g IV q8h</p> <p>Vancomycin 1g IV q12h (if MRSA suspected)</p> | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| 4. Acute Exacerbation of Chronic Bronchitis (AECB) - Chronic bronchitis - presence of both cough & sputum production on most days for at least 3 months each year for 2 consecutive years. Exacerbations are recurrent episodes of worsening respiratory symptoms. For classification of AECB please refer to Anthonisen et al. (Ann Int Med 1987; 106:196-204) and Seemungal et al (AJRCCM 1998; 157:1418-1422) - 40-50% AECB are caused by bacteria, usually H. Influenzae, S. Pneumoniae & M. Catarrhalis and 40% are due to viruses (influenzae A or B, rhinovirus, parainfluenzae, coronavirus) | | | |
| Acute tracheobronchitis - usually viral | None unless symptoms persist > 7 days | EES 800mg PO q12h for 1 week OR Doxycycline 100mg PO q12h for 1 week | Symptoms & risk factors: Cough & sputum without previous pulmonary disease |
| Chronic bronchitis without risk factors (simple) <i>H. Influenzae</i> <i>Haemophilus spp</i> <i>M. Catarrhalis</i> <i>S. Pneumoniae</i> <i>Atypical Respiratory Pathogens</i> | Azithromycin 500mg PO q24h for 1 week OR <i>2nd or 3rd gen. Cephalosporins (except ceftazidime)</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 625mg PO q12h for 1 week OR Doxycycline 100mg PO q12h for 1 week | Symptoms & risk factors: Increased cough & sputum, purulent sputum, and increased dyspnoea |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---------------------------------------|---|
| | Preferred | Alternative | |
| Chronic bronchitis with risk factors (complicated) <i>H. Influenzae</i> <i>M. Catarrhalis</i> <i>S. Pneumoniae</i> <i>Atypical Respiratory Pathogens</i> <i>Klebsiella sp</i> <i>Other gram negatives</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 625mg PO q12h for 1 week</i> OR <i>Ampicillin/Sulbactam 375mg PO q12h for 1 week</i> | Levofloxacin 500mg PO q24h for 1 week | Symptoms & risk factors: As in <i>chronic bronchitis without risk factors</i> plus (≥ 1 of): FEV1 <50%, > 4 exacerbations/year, > 65 years, significant co-morbidity (especially heart disease), use of home oxygen, chronic oral corticosteroid use, antibiotic use in the past 3 months |
| Chronic suppurative bronchitis <i>H. Influenzae</i> <i>M. Catarrhalis</i> <i>S. Pneumoniae</i> <i>Atypical respiratory pathogens</i> <i>Klebsiella sp</i> <i>Other gram negatives</i> <i>Pseudomonas Aeruginosa</i> <i>Multi-resistant Enterobacteriaceae</i> | Ambulatory patients: Tailor treatment to airway pathogen <i>Pseudomonas aeruginosa</i> common (Ciprofloxacin 500mg PO q12h) Hospitalised patients: parenteral therapy usually required | | Symptoms & risk factors: As in chronic bronchitis with risk factors with constant purulent sputum, some have bronchiectasis, FEV1 usually < 35%, or multiple risk factors (e.g. frequent exacerbations & FEV1 < 50%) |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

References:

- Gleason PP, Meehan TP, Fine JM, Galusha DH, Fine MJ. Associations between initial antimicrobial therapy and medical outcomes for hospitalized elderly patients with pneumonia. *Arch Intern Med* 1999; 159:2562-72
- Houck PM, et al. *Chest* 2001; 119:1420-6
- Gleason PP et al. *JAMA* 1997; 278:32-9
- Gordon GS et al. *Chest* 1996; 110:55S
- Stahl JE et al. *Arch Intern Med* 1999; 159:2576-80)
- CID* 40:915 & 923, 2005
- Gilbert DN, Moellering Jr RC, Eliopoulos GM, Sande MA. *The Sanford Guide To Antimicrobial Therapy* 2006.
- Anzueto AR, Schaberg. *Clinician's Manual On Acute Exacerbations Of Chronic Bronchitis*. 2003, Science Press Ltd

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Secondary Syphilis Incubation period: 6-8 weeks | As above | As above | Contact tracing |
| Early Latent Syphilis <i>Syphilis infection of less than 2 years duration.</i> Positive serology without symptoms and signs. | As above | As above | Contact tracing |
| Late Latent Syphilis Syphilis infection of more than 2 years duration | Procaine Penicillin 600,000 units IM q24h for 17 days OR Benzathine Penicillin 2.4 mega units IM weekly for 3 weeks | If allergic to penicillin: Doxycycline 100mg PO q12h for 28 days OR Tetracycline 500mg PO q6h for 28 days OR EES 800mg PO q12h for 28 days OR *Amoxicillin 2g PO q8h PLUS Probenecid 500mg PO q6h for 28 days | Contact tracing *Reference: British Association of Sexual Health and HIV Clinical Effectiveness Guidelines 2006 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|--|
| | Preferred | Alternative | |
| Neurosyphilis | Benzylpenicillin 3-4 mega units IV q4h for 14 days OR Procaine Penicillin 2.4 mega units IM q24h PLUS Probenecid 500mg PO q6h for 17 days | If allergic to penicillin: *Doxycycline 200mg PO q12h for 28 days OR *Amoxicillin 2g PO q8h PLUS Probenecid 500mg PO q6h for 28 days | Repeat CSF examinations every 6 months. Consider retreatment if cell count is not decreased in 6 months or CSF is not entirely normal in 2 years (Ref: MMWR 1998; 47, RR-1) All patients with neurosyphilis should be considered for corticosteroid cover at the start of the therapy to prevent the Jarisch-Herxheimer reaction (Prednisolone 10-20mg PO q8h for 3 days commencing one day prior to syphilis treatment) <u>*Reference:</u> British Association of Sexual Health and HIV Clinical Effectiveness Guidelines 2006 |
| Syphilis in HIV Primary, secondary, early and late latent, and of unknown duration | Treat as for non-HIV patients with neurosyphilis | Treat as for non-HIV patients with neurosyphilis | CSF examination should be done |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|---|--|---|
| | Preferred | Alternative | |
| Syphilis in Pregnancy | As in non-pregnant patients with syphilis | Use Erythromycin as in non-pregnant patients with syphilis | Tetracycline and Doxycycline are contraindicated in pregnancy Erythromycin can be used, but has a high risk of failure to cure the infection in infants. Therefore, all infants should be treated at birth |
| Congenital Syphilis | Benzylicillin 100,000-150,000 units/kg/day, administered as 50,000 units/kg/dose IV q12h during the first 7 days of life and q8h thereafter for a total of 10 days OR Procaine Penicillin 50,000 units/kg/dose IM q24h for 10 days | If allergic to penicillin: No proven alternative therapy. Penicillin desensitisation may be required | If a non-penicillin agent is used, close serologic and CSF follow-up are indicated |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|---|
| | Preferred | Alternative | |
| Gonorrhoea <i>Neisseria Gonorrhoeae</i> | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 250mg IM stat OR Spectinomycin 2g IM stat | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 500mg IM stat PLUS Probenecid 1g PO stat OR Cefuroxime 1.5g IM stat PLUS Probenecid 1g PO stat OR Norfloxacin 800mg PO stat OR Ciprofloxacin 500mg PO stat OR Ofloxacin 400mg PO stat OR Azithromycin 1g PO stat (covers NSU as well) | Contact tracing Also treat for non-specific urethritis (NSU) in view of high incidence of coexisting NSU in patients with gonorrhoea |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Gonococcal Epididymitis/ Epididymo-orchitis | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 500mg IM q24h for 5-7 days | Spectinomycin 2g IM q24h for 5-7 days PLUS Doxycycline 100mg PO q12h for 14 days OR Spectinomycin 2g IM q24h for 5-7 days PLUS EES 800mg PO q12h for 14 days | Contact tracing |
| Disseminated Gonorrhoea | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 1g IM/IV q24h continued for 24-48 hours after improvement begins, then switch to: Ciprofloxacin 500mg PO q12h OR Ofloxacin 400mg PO q12h | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 1g IV q8h OR Spectinomycin 2g IM q12h OR Ciprofloxacin 400mg IV q12h OR Ofloxacin 400mg IV q12h | Admit patient Contact tracing Duration of treatment depends on clinical response |
| Chlamydial/Non-Specific Urethritis (NSU)/Non-Specific Genital Infection in Women (NSGI) | Doxycycline 100mg PO q12h for 7 days | EES 800mg PO q12h for 7 days OR Ofloxacin 200mg PO q12h for 7 days OR Azithromycin 1g PO stat | Contact tracing Doxycycline and Ofloxacin are contraindicated in pregnancy |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|---|
| | Preferred | Alternative | |
| Chancroid <i>Haemophilus Ducreyi</i> | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 250mg IM stat OR Ciprofloxacin 500mg PO q12h for 3 days | EES 800mg PO q12h for 7 days OR Azithromycin 1g PO stat | Contact tracing |
| Lymphogranuloma Venereum <i>Chlamydia Trachomatis</i> Serovar L1, 2, 3 | Doxycycline 100mg PO q12h for 21 days OR Tetracycline 500mg PO q6h for 21 days | Minocycline 100mg PO q12h for 21 days OR EES 800mg PO q12h for 21 days OR Azithromycin 1g PO weekly for 3 weeks | Contact tracing Final duration depends on clinical response |
| Granuloma Inguinale <i>Klebsiella Granulomatis</i> | Doxycycline 100mg PO q12h for 3 weeks OR Tetracycline 500mg PO q6h for 3 weeks | Minocycline 100mg PO q12h for 3 weeks OR Trimethoprim/Sulfamethoxazole 160/800mg PO q12h for 3 weeks OR EES 800mg PO q12h for 3 weeks OR Ciprofloxacin 750mg PO q12h for 3 weeks OR Azithromycin 1g PO weekly for 3 weeks or 500mg PO q24h for 7 days | Contact tracing Add Gentamicin ¹ 1.5mg/kg IM/IV q8h in patients whose lesions do not respond in the first few days to other agents Duration of treatment should be until lesions have healed. Healing times vary greatly between patients. A minimum of 3 weeks treatment is recommended |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|-------------|----------|
| | Preferred | Alternative | |
| Trichomoniasis <i>Trichomonas Vaginalis</i> | Refer to Page 71 (Obstetrics & Gynaecology Infections) | | |
| Bacterial vaginosis <i>Gardnerella Vaginalis, Anaerobes</i> | Refer to Page 71 (Obstetrics & Gynaecology Infections) | | |
| Herpes Genitalis <i>Herpes Simplex Virus 1 and 2</i> | <p><u>First episodic:</u> Acyclovir 200mg PO 5 times a day for 5 days</p> <p><u>Recurrent - episodic:</u> Acyclovir 200mg PO 5 times a day for 5 days</p> <p><u>Suppressive therapy:</u> (may be indicated if ≥ 6 recurrences per year) Acyclovir 400mg PO q12h or 200mg PO 4 times a day for up to 1 year, then reassess</p> | | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

References:

1. British Association of Sexual Health and HIV Clinical Effectiveness Guidelines 2006
2. Center for Disease Control and Prevention, Sexually Transmitted Diseases Treatment Guidelines 2006. MMWR 2006 Aug; Vol. 55, RR-11
3. European STD Guidelines. Int J STD AIDS 2001 Oct. 12 Suppl 3:2-3

SKIN AND SOFT TISSUE INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Bacterial Infections | | | |
| Impetigo/Ecthyma <i>S. Aureus</i> <i>S. Pyogenes</i> | Cloxacillin 500mg PO q6h for 5-7 days | EES 800mg PO q12h for 5-7 days OR Cephalexin 500mg PO q6h for 5-7 days OR Azithromycin 500mg PO q24h for 3-5 days | <u>References:</u> 1. Australian Medicines Handbook 2006 (revised July 2006) 2. Cambridgeshire GP antibiotic Guidelines from NHS Primary Care Trust. Reviewed: Sept 2006 3. Practice guidelines for the diagnosis and management of skin and soft-tissue infections. Clinical Infectious Diseases 2005; 41:1373-1406 |
| Boils/Carbuncles <i>S. Aureus</i> | Cloxacillin 500mg PO q6h for 7-10 days | EES 800mg PO q12h for 7-10 days OR Cefuroxime 500mg PO q12h for 7-10 days OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 625mg PO q12h for 7-10 days | Surgical drainage is important in the management <u>Reference:</u> Australian Medicines Handbook 2006 (revised July 2006) |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| Cellulitis/Erysipelas <i>Strep Pyogenes</i> <i>Staph Aureus</i> | Cloxacillin 1g IV q6h Change to oral (Cloxacillin 1-2g q6h) once condition improves | Cefazolin 1g IV q8h OR EES 800mg PO q12h OR Cephalexin 500mg PO q6h <i>Change to oral once condition improves</i> | <u>References:</u> 1. Australian Medicines Handbook 2006 (revised July 2006) 2. Cambridgeshire GP antibiotic Guidelines from NHS Primary Care Trust. Reviewed: Sept 2006 |
| Diabetic Foot Infections | Refer to Page 123 (Bone & Joint Infections) | | |
| Gas Gangrene/Myonecrosis/ Necrotizing Fasciitis <i>Streptococci</i> <i>Clostridium sp.</i> <i>Polymicrobial</i> | Refer to Page 123 (Bone & Joint Infections) | | |
| Yaws <i>Treponema Pertenu</i> | Benzathine Penicillin 2.4 mega units IM single dose If allergic to penicillin: Tetracycline 500mg PO q6h for 15 days OR EES 800mg PO q12h for 15 days | Doxycycline 100mg PO q12h for 15 days | <u>Reference:</u> Fitzpatrick's Dermatology in General Medicine Vol II Sixth Edition |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|--|
| | Preferred | Alternative | |
| Mycobacterial Infections | | | |
| Hansen's Disease (Leprosy) Mycobacterium Leprae | Sg. Buloh Augmented Regime <i>Paucibacillary</i> Rifampicin 600mg PO monthly (supervised) PLUS Dapsone 100mg PO q24h PLUS Clofazimine 50-100mg PO q24h Duration: 1 year Surveillance: BI/MI annually for 5 years | WHO Regime <i>Paucibacillary</i> (1-5 skin lesions) Rifampicin 600mg PO monthly PLUS Dapsone 100mg PO q24h Duration: 6 months | <u>References:</u> 1. Guidelines for M.D.T. 1991 by Dr. T. Ganesapillai 2. World Health Organisation health guidelines |
| | <i>Multibacillary</i> <u>Intensive phase:</u> Rifampicin 600mg PO q24h PLUS Dapsone 100mg PO q24h PLUS Clofazimine 100mg PO q24h Duration: 3 weeks (or till MI=0) | <i>Multibacillary</i> (>5 skin lesions) Rifampicin 600mg PO monthly PLUS Dapsone 100mg PO q24h PLUS Clofazimine 300mg PO monthly and 50mg q24h Duration: 1 to 2 years | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|--|----------|
| | Preferred | Alternative | |
| | <p><u>Maintenance phase:</u> Rifampicin 600mg PO monthly PLUS Dapsone 100mg PO q24h PLUS Clofazimine 300mg PO monthly and 50-100mg q24h Duration: 3 years For those with BI>3, treat till smear negative Surveillance: BI/MI annually for 10 years</p> | <p>Single skin lesion paucibacillary leprosy Single dose of: Rifampicin 600mg PO PLUS Ofloxacin 400mg PO PLUS Minocycline 100mg PO</p> <p>Bacterial resistance or hypersensitivity to first line Can be substituted with one of the following: Minocycline 100mg PO q24h Ofloxacin 400mg PO q24h Clarithromycin 500mg PO q24h Ethionamide 250mg PO q24h</p> | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| Atypical Mycobacterial Infections <i>Mycobacterium Marinum</i> | Clarithromycin 500mg PO q12h PLUS Minocycline/Doxycycline 100mg PO q12h OR Trimethoprim/Sulphamethoxazole 160/800mg PO q12h For 4-6 months, and continue for at least 1 month after lesions have been cleared | Rifampicin 600mg PO q24h PLUS Ethambutol 15mg/kg PO q24h for 4-6 months, and continue for at least 1 month after lesions have been cleared | No available consensus guidelines Only case reports |
| <i>Mycobacterium Kansasi</i> | Isoniazid 300mg PO q24h PLUS Rifampicin 600mg PO q24h PLUS Ethambutol 15mg/kg PO q24h for 18 months | | |
| <i>Mycobacterium Ulcerans</i> | Amikacin ¹ 15mg/kg IV q24h PLUS Clarithromycin 500mg PO q12h | | Wide surgical excision and debridement are important |
| <i>Mycobacterium Fortuitum/Chelonei</i> | Doxycycline/Minocycline 100mg PO q12h PLUS Clarithromycin 500mg PO q12h | | Surgical debridement is important |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| | <p>OR Amikacin¹ 15mg/kg IV q24h PLUS Clarithromycin 500mg PO q12h</p> <p>For 4-6 months, and continue for at least 1 month after lesions have been cleared</p> | | |
| Fungal Infections | | | |
| <p>Tinea Capitis / Tinea Barbae <i>Trichophyton, Microsporum</i></p> | <p>Griseofulvin 10-15mg/kg/24h PO OR 500mg q12h or q24h for 6 weeks</p> | <p>Terbinafine 250mg PO q24h OR Itraconazole 200mg PO q24h for 2-6 weeks</p> | <p><u>Reference:</u> Australian Medicines Handbook 2006 (revised July 2006)</p> |
| <p>Tinea Corporis / Tinea Cruris / Tinea Faciei <i>Trichophyton, Microsporum, Epidermophyton</i></p> | <p><u>Mild infections:</u> Topical imidazole cream: Clotrimazole 1% OR Miconazole 2% OR Tioconazole 1% Duration: 4 weeks</p> <p><u>Extensive infections:</u> Griseofulvin 500mg PO q12h or q24h for 4-6 weeks</p> | <p>Terbinafine 250mg PO q24h for 2 weeks OR Itraconazole 200mg PO q24h for 2 weeks</p> | <p><u>Reference:</u> Australian Medicines Handbook 2006 (revised July 2006)</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Tinea Manuum/ Tinea Pedis <i>Trichophyton, Microsporum, Epidermophyton</i> | Griseofulvin 500mg PO q12h for 6-12 weeks OR Itraconazole 200mg PO q24h for 2-4 weeks | Terbinafine 250mg PO q24h for 2-4 weeks | |
| Tinea Unguium <i>Trichophyton, Microsporum, Epidermophyton</i> | Terbinafine 250mg PO q24h For 6 weeks (finger nails) For 12 weeks (toe nails) OR Pulse Itraconazole 200mg PO q12h for 1 week per month For 2 months (finger nails) For 3 months (toe nails) | Griseofulvin 500mg PO q12h For 6 months (finger nails) For 12 months (toe nails) OR Amorolfine 5% Nail Lacquer weekly application For 6 months (finger nails) For 12 months (toe nails) | <u>Reference:</u> Australian Medicines Handbook 2006 (revised July 2006) |
| Tinea Versicolor <i>Malassezia Furfur Pityrosporum Orbiculare</i> | Selenium Sulphide 2% shampoo apply to affected areas 20-30 minutes before bathing OR Dilute to 1:1 with water, apply and leave overnight (treat for 1-2 weeks) <u>For face:</u> Topical Imidazole for 4-6 weeks e.g. Miconazole 2% cream, Clotrimazole 1% cream, Tioconazole 1% cream | Itraconazole 200mg PO q24h for 1 week OR Ketoconazole 200mg PO q24h for 1 week | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Subcutaneous Fungal Infections 1. Sporotrichosis | Itraconazole 200mg PO q12h for 4-6 months and continue for at least 1 month after recovery | Terbinafine 250mg PO q24h for 4-6 months and continue for at least 1 month after recovery OR Potassium iodide (saturated solution 50mg/drop) PO 500-1500mg/day, increase to 4000-6000mg/day in 3 divided doses for 6-10 weeks | In some immunocompromised condition such as AIDS, longer treatment maybe necessary. Refer to Page 53 (Opportunistic Infections In HIV Patients) |
| 2. Chromomycosis, Eumycetoma | Itraconazole 200mg PO q12h for 4-6 months and continue for at least 1 month after recovery | | |
| 3. Cryptococcosis | Fluconazole 200-400mg IV/PO q24h for 2 weeks (in ill patients initial therapy with IV Amphotericin B is preferred) | Amphotericin B IV 0.6-1mg/kg q 24h | |
| 4. Histoplasmosis, Penicilliosis, etc. | Itraconazole 200mg PO q12h for 2-4 months or till lesions healed, then 200mg q24h for 1-2 months (in ill patients initial therapy with IV Amphotericin B is preferred) | Amphotericin B IV 0.6-1mg/kg q24h | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|--|
| | Preferred | Alternative | |
| Viral Infections | | | |
| Herpes Simplex Infections | <p>Oral: Primary: Acyclovir 200-400mg PO 5 times daily for 5 days</p> <p>Recurrent: Regular normal saline dabs/gargle</p> <p>In immunocompromised patients. Refer to Page 53 (Human Immunodeficiency Virus)</p> <p>Genitalia: (Refer to Page 100 Sexually Transmitted Infections)</p> <p>Eczema herpeticum: Acyclovir 200mg PO 5 times daily for 7-10 days</p> | <p>Severe cases: Acyclovir 5mg/kg IV q8h for 5 days or until able to take orally, then change to oral</p> | |
| Chickenpox <i>Varicella Zoster</i> | <p><u>Immunocompetent:</u> Acyclovir 800mg PO 5 times daily for 1 week</p> <p><u>Immunocompromised/disseminated:</u> Acyclovir 10mg/kg IV q8h for 1 week (change to oral once there is an improvement)</p> | | <p>Advisable to start treatment early within 48 hours</p> <p><u>Reference:</u> Infectious Diseases Society of America Guidelines 2005</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|---|
| | Preferred | Alternative | |
| Herpes Zoster <i>Varicella Zoster</i> | Acyclovir 800mg PO 5 times daily for 1 week* | | *Only indicated in immunocompromised patients, herpes zoster ophthalmicus, Ramsay-Hunt syndrome and the elderly Advisable to start treatment early within 48 hours |
| Parasitic Infestations | | | |
| Scabies <i>Sarcoptes Scabiei</i> | Benzyl Benzoate emulsion 25% (EBB) apply from neck down and leave for 24 hours for 2 days | Gamma Benzene Hexachloride 1% (Lindane) apply and leave for 8 hours (not to be repeated in less than a week) OR Permethrin 5% cream apply and leave for 8 hours Pregnant women: Sulphur 6% in calamine lotion apply q12h OR Crotamiton (Eurax) cream apply q12h for 2-3 weeks OR Permethrin 5% cream apply and leave for 8 hours | References: 1. Centers for Disease Control and Prevention. Sexually transmitted diseases treatment guidelines 2006 2. David Flinders. American Academy of Family Physicians 2003 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|----------------------|----------|
| | Preferred | Alternative | |
| Head Lice <i>Pediculus Humanus Capitis</i> | Gamma Benzene Hexachloride 0.1% (Lindane) apply and leave for 8 hours | Malathion 1% shampoo | |
| Body Lice/pubic Lice <i>Pediculus Humanus</i> | As for Head Lice | | |

*Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

SURGICAL INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| A. GENERAL SURGERY | | | |
| Appendicitis <i>Enterobacteriaceae Enterococci, Bacteroides</i> | Ampicillin 500mg IV q4-6h PLUS Gentamicin ¹ 5mg/kg IV q24h PLUS Metronidazole 500mg IV q8h | <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g IV q6-8h</i> OR Amoxicillin/Clavulanate 1.2g IV q8h | Start upon diagnosis, discontinue after surgery |
| Perforated Appendix, Appendicular Mass | Metronidazole 500mg IV q8h PLUS <i>3rd gen. Cephalosporins, e.g. Cefoperazone 2-4g/day IV in divided doses q12h</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g IV q6-8h</i> OR Amoxicillin/Clavulanate 1.2g IV q8h | Duration 5-7 days |
| Perforated Viscus Peritonitis | Ampicillin 500mg IV q6h PLUS Metronidazole 500mg IV q8h PLUS Gentamicin ¹ 5mg/kg IV q24h OR <i>3rd gen. Cephalosporins, e.g. Cefoperazone 2-4g/day IV in divided dose q12h</i> PLUS Metronidazole 500mg IV q8h | Cefoperazone/Sulbactam 1-2g q12h, up to maximum 8g/day OR <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g IV q6-8h</i> OR Amoxicillin/Clavulanate 1.2g IV q8h | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| Abdominal trauma Suspected bowel or solid organ injury <i>Gram negative enteric aerobes and anaerobes</i> | Cefuroxime 1.5g IV q8h OR <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 1g IV q8h OR Cefoperazone 1g IV q12h | Cefoperazone/Sulbactam 1g IV q12h PLUS Metronidazole 500mg IV q8h OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Ampicillin/Sulbactam 1.5g IV q8h OR Amoxicillin/Clavulanate 1.2g IV q8h | Duration - min 5 days |
| <i>Breast Abscess</i> <i>Staph Aureus</i> | Cloxacillin 1g IV q6h | | Drainage may be required |
| VASCULAR | | | |
| Mycotic Pseudoaneurysm in IVDU | Cloxacillin 2g IV q6h | Based on C&S | Initial therapy is high dose IV followed by oral therapy once debridement and ligation done. The duration will depend on clinical response |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| Prosthetic Graft Infection Non-MRSA MRSA | 3 rd gen. Cephalosporins, e.g. Cefotaxime 1g q8h OR Cefoperazone 2-4g/24h IV in two divided doses Vancomycin ¹ 1g IV q12h | Based on C&S Linezolid 600mg IV q12h | Duration may need to be prolonged if graft salvage considered Vancomycin levels need to be monitored. Graft may need to be explanted |
| Ischaemic Ulcers | β -lactam/ β -lactamase inhibitors, e.g. Amoxicillin/Clavulanate 625mg PO q12h IV OR Ampicillin/Sulbactam 375mg PO q12h | Based on C&S | Given IV if diabetes present Refer Page 123 (Bone & Joint Infections) |
| BITES (penetrating injuries) | | | |
| Animal bite <i>S. Aureus, Strep., Gram -ve Bacilli, Anaerobes</i> | β -lactam/ β -lactamase inhibitors, e.g. Amoxicillin/Clavulanate 625mg PO q12h | If severe, Cefuroxime 750mg IV q8h | Consider IV for severe cases Duration 3-5 days. If infected: 10 days |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|---|
| | Preferred | Alternative | |
| Human bite <i>S. Aureus</i> , <i>Anaerobes</i> , <i>Eikenella</i> | <i>β-lactam/β-lactamase inhibitors</i> , e.g. Amoxicillin/Clavulanate 625mg PO q12h | If allergic to Penicillin, Clindamycin 300mg PO q6h PLUS Ciprofloxacin 500-750mg PO q12h OR Trimethoprim/Sulphamethoxazole 160/800mg PO q12h | Duration 3-5 days Delay or do not suture |
| B. BONE AND JOINT INFECTIONS | | | |
| <i>Septic Arthritis</i> <i>Staph. Aureus</i> | Cloxacillin 1-2 g IV q6h | If Penicillin allergy (immediate hypersensitive type) Clindamycin 300-600mg IV q8h followed by oral therapy (same dose) | Drainage, debridement and washout of infected joint is important to limit further damage Empirical therapy wherever possible should be directed by the result of the Gram stain of the joint aspirate If initial gram stain is gram positive cocci use: ♦ Cloxacillin If initial gram stain is gram negative bacilli use: ♦ 3 rd gen. <i>Cephalosporins</i> , e.g. Ceftriaxone 2g IV daily |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| OSTEOMYELITIS | | | |
| Acute Osteomyelitis <i>S. Aureus</i> (80%), <i>Group A Strep Pyogenes</i> , rarely gram negative Bacilli | Cloxacillin 1-2g IV q6h PLUS <i>3rd gen. Cephalosporins, e.g. Ceftriaxone</i> 1-2g IV q24h if gram negative bacilli on gram stain | If Penicillin allergy (immediate hypersensitive type) Clindamycin 300-600mg IV q8h followed by oral therapy (same dose) | Duration: Initial IV therapy for 2-4 weeks followed by oral therapy. Minimum 6 weeks. Modify according to clinical response |
| Chronic Osteomyelitis (after 3 months of appropriate antibiotic therapy or presence of dead bone on x-ray) Commonest <i>S. Aureus</i> | Empirical treatment is not indicated Thorough Surgical debridement required (Removal of dead bone/ orthopaedic hardware) Choice of antibiotic depends on C&S result from tissue/bone | | Surgical debridement if necessary Minimum length 6 weeks but usually > 3 months Treat until inflammatory parameters are normal |
| Diabetic Foot Infections Antibiotics should not be used unless there are local or systemic symptoms of infection. Local treatment including surgical debridement is important. Antibiotic selection should be based on the most recent culture and sensitivity report. | | | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| <p>Mild Infections:</p> <p><i>Presence of > 2 markers of inflammation (purulence or erythema, pain, tenderness, warmth, or induration) with any cellulitis/erythema extending less than 2 cm around the ulcer; infection is limited to the skin or superficial subcutaneous tissues; no systemic toxicity</i></p> | <p>Cloxacillin 500mg PO q6h</p> <p>OR</p> <p><i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 625mg PO q12h</i></p> | <p>Cephalexin 500mg PO q6h</p> <p>OR</p> <p>Clindamycin 300-450mg PO q6</p> | <p>Duration of treatment: 1-2 weeks</p> |
| <p>Moderate Infections:</p> <p>Features of mild infection, no systemic toxicity or metabolic instability and > 1 of the following: cellulitis extending more than 2 cm around an ulcer, lymphangitic streaking, spread beneath the superficial fascia, deep tissue abscess, gangrene, or involvement of muscle, tendon, joint, or bone</p> | <p><i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5-3g IV q8h</i></p> <p>OR</p> <p><i>2nd or 3rd gen. Cephalosporins, e.g. Cefuroxime 750mg-1.5g IV q8h</i></p> <p>OR</p> <p>Ceftriaxone 1-2g q24h</p> <p>PLUS/MINUS</p> <p>Metronidazole 500mg IV q8h</p> | <p>Ciprofloxacin 500-750mg PO q12h</p> <p>OR</p> <p>Clindamycin 300-450mg PO q6h</p> <p>If antibiotic-resistant organisms are likely, treat as severe infection</p> | <p>Duration of treatment: usually 2-4 weeks. Modify according to clinical response</p> <p>If proven osteomyelitis: at least 4-6 weeks. However, a shorter duration (3 weeks) is sufficient if the entire infected bone is removed</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| <p>Severe Infections:</p> <p><i>Infection plus systemic toxicity or metabolic instability (e.g. fever, chills, tachycardia, hypotension, confusion, vomiting, leukocytosis, metabolic acidosis, severe hyperglycemia, or azotemia above baseline)</i></p> | Piperacillin/Tazobactam 4.5g IV q6-8h OR <i>3rd gen. Cephalosporins, e.g. Ceftazidime 2g IV q8h</i> PLUS Metronidazole 500mg IV q6h | Imipenem/Cilastatin 500mg IV q6h | Add Vancomycin ¹ 1g IV q12h, if high risk for MRSA Duration of treatment: as in moderate infection Necrotizing fasciitis |
| Necrotizing Fasciitis | | | |
| <p>Type 1</p> <p>Polymicrobial infection. Primarily occurs in patients who are immunocompromised or have certain chronic diseases such as diabetes</p> | Cloxacillin 2g IV q4-6h PLUS Metronidazole 500mg IV q8h PLUS Gentamicin ¹ 5mg/kg IV q24h | <i>3rd gen. Cephalosporins</i> PLUS Metronidazole 500mg IV q8h OR <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g IV q8h</i> OR Amoxicillin/Clavulanate 1.2g IV q8h PLUS/MINUS Gentamicin ¹ 5mg/kg IV q24h | Early aggressive surgical debridement essential |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Type 2 <i>Group A strep</i> | Benzympenicillin 2-4 mega units IV q4h PLUS Clindamycin 600mg IV q8h | | Suspect Group A Strep if Gram stain shows Gram positive cocci in chains Early aggressive surgical debridement essential |
| Soft Tissue Infection Secondary To Gas Producing Organism | | | |
| <i>e.g. Clostridium spp,</i> <i>Gram -ve org</i> | *Benzympenicillin 2-4 mega units IV q4h PLUS Metronidazole 500mg IV q8h PLUS/MINUS Gentamicin ¹ 5mg/kg IV q24h | <i>3rd gen. Cephalosporins</i> PLUS Gentamicin ¹ 5mg/kg IV q24h Depends on culture & sensitivity | *For Clostridium sp.: Benzympenicillin 4 mega units q6h is preferred Early aggressive surgical debridement essential |
| Suppurative Wound Infections, Surgical Or Traumatic | | | |
| Suppurative wound infections, surgical or traumatic | If there is surrounding cellulitis and/or systemic symptoms are present: Cloxacillin 500mg PO/IV q6h If gram negative organisms suspected or known to be involved: Gentamicin ¹ 5mg/kg IV q24h OR As a monotherapy: Cefuroxime 1.5g IV q8h | Change antibiotics accordingly after trace culture and sensitivity result | Topical antibiotics are not recommended for treatment of wound infections as it may result in the emergence of resistant organisms Patient tetanus immunisation status should be assessed in all cases |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Muscular, Skeletal and Soft Tissue Trauma, Crush Injuries and Stab Wounds | | | |
| Muscular, skeletal and soft tissue trauma, crush injuries and stab wounds | Cloxacillin 2g IV q6h PLUS Gentamicin ¹ 5mg/kg IV q24h PLUS Metronidazole 500mg IV q8h Duration: Not less than 5 days | Cefuroxime 1.5g as a loading dose, followed by 750mg IV q8h PLUS Metronidazole 500mg IV q8h Duration: Not less than 5 days | Thorough surgical debridement, soft tissue and fracture stabilisation For severe penetrating injuries, especially those involving joints and/or tendons, antibiotics must be given for at least 5 days |
| Compound Fractures | | | |
| Compound fractures | Cloxacillin 1g IV q6h OR Cefuroxime 1.5g IV q8h If wound soiling or tissue damage is severe and/or devitalized tissue is present: PLUS Gentamicin ¹ 5mg/kg IV q24h PLUS Metronidazole 500mg IV q8h Duration: 5-10 days | | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| C. UROLOGY | | | |
| Pyonephrosis/Perinephric Abscess <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV q8h</i> PLUS Gentamicin ¹ 5mg/kg IV q24h OR <i>3rd gen. Cephalosporins, e.g. Cefoperazone 1g IV q12h</i> | Ciprofloxacin 200-400mg IV q12h | PLUS Drainage followed by definitive surgery |
| Renal Abscess <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas, Staph Aureus</i> | <i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 1.5g IV q8h followed by 375mg PO q12h</i> OR Cefuroxime 750mg IV q8h followed by 250mg PO q12h PLUS/MINUS Gentamicin ¹ 5mg/kg IV q24h Minimum of 2 weeks | <i>3rd gen. Cephalosporins, e.g. Ceftriaxone 1-2g IV q24h</i> | Drainage may be required. Commence oral after temperature settled |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| Acute Prostatitis <i>E. Coli</i> <i>Staph Saprophyticus</i> <i>Enterococcus</i> <i>Enterobacteriaceae</i> <i>Proteus</i> | If ill and hospitalised Ciprofloxacin 200mg IV q12h PLUS/MINUS Gentamicin ¹ 5mg/kg IV q24h Less Severe infection: Ciprofloxacin 500mg PO q12h | <i>3rd gen. Cephalosporins, e.g.</i> Cefoperazone 1g IV q12h Trimethoprim/Sulfamethoxazole 160/800mg PO q12h OR Trimethoprim 300mg PO q24h | Treatment for 4 weeks |
| Chronic Bacterial Prostatitis (CPPS NIH Type II) <i>Mostly culture negative</i> | Ciprofloxacin 500mg PO q12h for 2 weeks Then reassess, if beneficial to continue for 4-6 weeks | Trimethoprim/Sulfamethoxazole 160/800mg PO q24h for 2 weeks Then reassess, if beneficial to continue for 4-6 weeks | Pending positive culture on prostatic secretion |
| Prostatic Abscess <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | Ciprofloxacin 200-400mg IV q12h followed by 500mg PO q12h minimum of 2-4 weeks | <i>3rd gen. Cephalosporins, e.g.</i> Cefoperazone 1g IV q12h followed by, Cefuroxime 500mg PO q12h minimum of 2-4 weeks | Drainage mandatory |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|--|
| | Preferred | Alternative | |
| Non Gonococcal Urethritis | | | Refer to Page 100 (Sexually Transmitted Infections) |
| Epididymo-orchitis <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | Doxycycline 100mg PO q12h minimum of 2 weeks | Ciprofloxacin 500mg PO q12h minimum of 2 weeks | |
| Testicular Abscess <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h OR Ampicillin/Sulbactam 1.5g IV q8h | <i>3rd gen. Cephalosporins, e.g.</i> Cefoperazone 1g IV q12h | PLUS drainage |
| Fournier's Gangrene <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas, Anaerobes</i> | <i>3rd gen. Cephalosporins, e.g.</i> Cefoperazone 1g IV q12h PLUS Metronidazole 500mg IV q8h | Cefoperazone/Sulbactam 1g IV q12h PLUS Metronidazole 500mg IV q8h | PLUS debridement |
| Urosepsis (Septicaemia post urological instrumentation or urological infections) <i>E. Coli, Klebsiella, Proteus, Enterococcus, Pseudomonas, MRSA</i> | Cefepime 1g IV q12h OR Imipenem/Cilastatin 500mg IV q8h | Cefoperazone/Sulbactam 1g IV q12h | Choice of antibiotics should be adapted based upon culture results |

*Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| D. NEUROSURGERY | | | |
| Brain Abscess | | | |
| <i>Contiguous source of infection</i> Paranasal sinuses Orogenic infection | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q12h PLUS Metronidazole 500mg IV q8h | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 2g IV q6h PLUS Metronidazole 500mg IV q8h | Usual treatment for uncomplicated infection is 7-14 days, for complicated is 6-8 weeks |
| Postoperative | Cloxacillin 2g IV q4h | Vancomycin ¹ 1g IV q12h (MRSA) PLUS <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q12h | |
| Post-traumatic | Cloxacillin 2g IV q4h PLUS <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q12h | | |
| Source of infection unknown | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV q12h PLUS Metronidazole 500mg IV q8h PLUS/MINUS Cloxacillin 2g IV q4h | Vancomycin ¹ 1g IV q12h (MRSA) PLUS Metronidazole 500mg IV q8h | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|------------|
| | Preferred | Alternative | |
| Penetrating craniocerebral injuries (PCCI) and depressed fractures including base of skull fracture | Cefuroxime 1.5g IV stat dose followed by 750mg IV q8h PLUS Metronidazole 500mg IV q8h OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g q8h IV/625mg PO q12h | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 2g IV stat followed by 1g IV q12h PLUS Metronidazole 500mg IV q8h | For 5 days |
| Open scalp laceration | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h/625mg PO q12h | Cloxacillin 1-2g IV q6h | For 5 days |

*Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

TROPICAL INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| 1. Management of Typhoid Fever | | | |
| Stable Case Fully sensitive | Pefloxacin 400mg PO q12h for 5-7 days OR Ciprofloxacin 750mg PO q12h for 5-7 days OR Levofloxacin 500mg PO q24h for 5-7 days | Ampicillin 500mg PO q6h for 14 days OR Chloramphenicol 500mg PO q6h for 14 days OR Trimethoprim/Sulphamethoxazole 160/800mg PO q12h for 14 days | WHO, 2003 Fever clearance is faster with Quinolones |
| Stable Case Multidrug resistance (Resistance to CMC, Ampicillin and TMP-SMX) | Ciprofloxacin 500mg PO q12h for 5-7 days | Azithromycin 500mg PO q24h for 7 days | WHO, 2003 |
| Quinolone resistance | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 3g IV q24h for 10-14 days OR Azithromycin 500mg PO q24h for 7 days | | WHO, 2003 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| Unstable or complicated cases | <p><i>3rd gen. Cephalosporins, e.g. Ceftriaxone 3g/24h IV for 7-10 days</i></p> <p>OR</p> <p>Ciprofloxacin 200mg IV q12h for 7-10 days</p> | | <p>Indication of Dexamethasone (discuss with physician)</p> <p>i) Thyphoid psychosis ii) Sepsis with shock</p> <p>Dose: 3mg/kg loading. Followed by 1mg/kg q6h for 2 days WHO, 2003 Paed. Inf. Dis J,1988</p> |
| 2. Management of Cholera | | | |
| <p><i>Non Tetracycline resistance</i></p> <p><i>Tetracycline resistance</i></p> | <p>Doxycycline 300mg PO stat (once patient can take orally)</p> <p>EES 400mg PO q12h for 3 days (The only option in pregnancy)</p> | <p>Ciprofloxacin 1g PO stat</p> <p>Ciprofloxacin 1g PO stat</p> | <p>Principle of Treatment:</p> <p>i) Rehydration ORS if tolerating orally ii) Monitor urine output iii) Avoid anti-diarrhoea agents - Diphenoxylate HCL/Atropine Sulphate (Lomotil) or Loperamide HCL (Imodium)</p> <p>WHO Global Task on Cholera Control 2004</p> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| 3. Management of Scrub Typhus | | | |
| Scrub Typhus (Orientia tsutsugamushi) | | Chloramphenicol 500mg PO q6h for 3-7 days OR Azithromycin 500mg PO stat (mild scrub typhus) | Pregnancy: Azithromycin 500mg PO stat CID 2004 Nov 1; 39(9):1329-35 |
| Tetracycline sensitive | Doxycycline 200mg PO q24h for 3-7 days | Rifampicin 900mg PO q24h for 7 days | |
| Reduced susceptibility to Tetracycline | Azithromycin 500mg PO stat (mild scrub typhus) | | |
| 4. Management of Brucellosis | | | |
| <i>Brucellosis</i> <i>B. Melitensis, B. Abortus, B. Suis</i> <i>and B. Canis</i> | Doxycycline 100mg PO q12h PLUS Rifampicin 600-900mg (15mg/kg) PO q24h for 6 weeks; OR Doxycycline 100mg PO q12h for 6 weeks PLUS Gentamicin ¹ 1.5mg/kg IV q8h for 7 days | Ofloxacin 400mg PO q24h PLUS Rifampicin 600-900mg PO q24h for 6 weeks; OR Rifampicin 900mg PO q24h PLUS Trimethoprim/Sulphamethoxazole 160/800mg PO q12h for 6 weeks | Pregnancy: Rifampicin 900mg PO q24h CID 42:10752006 NEJM 352; 2005 |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|--|
| | Preferred | Alternative | |
| 5. Management of Leptospirosis | | | |
| Severe disease (Leptospiral pulmonary syndrome, multiorgan involvement, sepsis) | Benzylpenicillin 2.4 mega units IV q6h for 7 days; OR 3 rd gen. Cephalosporins, e.g. Ceftriaxone 1g IV q24h for 7 days | 3 rd gen. Cephalosporins, e.g. Cefotaxime 1g IV q8h for 7 days | Clin Infect Dis 2003; 36:1507-1513 Clin Infect Dis 2004; 39:1417-1424 |
| Mild to Moderate disease | Benzylpenicillin 2.4 mega units IV q6h for 7 days | Doxycycline 100mg PO q12h for 7 days OR Azithromycin 500mg PO q24h for 7 days | <u>Reference:</u> Clin Infect Dis 2003; 36:1514-1515 |
| 6. Management of Tetanus | | | |
| Clostridium Tetani | Metronidazole 500mg IV q6h for 7-10 days | Erythromycin 1g IV q6h OR Clindamycin 600mg IV q6h for 10 days | (Penicillin, a GABA antagonist, may aggravate the spasms) |
| Toxin neutralisation (if visible point of entry) | Human Tetanus Immunoglobulin 3000 to 6000 iu IM | | A single 500-iu dose of human immunoglobulin may be as effective |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|---|---|
| | Preferred | Alternative | |
| 7. Management of Melioidosis | | | |
| Melioidosis <i>Burkholderia Pseudomallei</i> | <p>Initial Therapy 3rd gen. Cephalosporins, e.g. Ceftazidime 120mg/kg/24h IV q6-8h PLUS/MINUS Trimethoprim/Sulphamethoxazole 8/40mg/kg/24h IV for 2-3 weeks</p> <p>Maintenance Therapy Trimethoprim/Sulphamethoxazole 10/50mg/kg/24h PO PLUS Doxycycline 100mg PO q12h Duration minimum 20 weeks</p> | <p>Cefoperazone/Sulbactam 2g IV q8h PLUS/MINUS Trimethoprim/Sulphamethoxazole 8/40mg/kg/24h IV for 2-3 weeks</p> <p>OR Imipenem 500-750mg IV q6h for 2-3 weeks</p> <p><i>β-lactam/β-lactamase inhibitors, e.g.</i> * Amoxicillin/Clavulanate 1250mg (2 tablets of 625mg) PO q8h</p> <p>OR Trimethoprim/Sulphamethoxazole 8/40mg/kg/24h Duration minimum 20 weeks</p> | <p><u>Reference:</u> Clinical Microbiology Reviews, Apr 2005, p. 383-416</p> <p>Look for source of infection</p> <p>Antimicrobial Agents and Chemo, Oct 2005, 4020-4025</p> <p>*Well tolerated and has better adverse effect profile than the conventional regimen (Doxycycline & Trimethoprim/Sulphamethoxazole) but it is associated with a higher relapse rate</p> |

8. Malaria (Ref: 1) WHO malaria guidelines 2006
2) CDC: Malaria (Prescription drugs for Malaria updated Feb 2007)

WHO recommended combination therapies on the basis of the available safety and efficacy data

Risk group:

Pregnancy
Children < 5 years old
Severe vomiting, headache
BFMP: parasites >100,000/ul or BFMP ++++

Features of severe/complicated Malaria includes at least one of the following:

Clinical manifestation:

Prostration
Impaired consciousness -GCS <15
Respiratory distress (acidotic breathing)
Multiple convulsions
Pulmonary oedema (radiological)
Abnormal bleeding
Jaundice
Shock/Algid malaria
Haemoglobinuria- coffee coloured urine

Laboratory test:

Acute Renal Failure (Sr creatinine >265umol/l)
Metabolic acidosis- HCO₃ <15mmol/l
Hyperlactatemia; serum lactate >5mmol/l
Hepatic dysfunction
Hyperparasitemia
Hypoglycaemia
Severe anaemia
DIVC

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| Malaria | | | |
| Plasmodium Falciparum a) Non Complicated i) New Infection | Adult (>35kg) D1-D3: (Artequin) Artesunate 200mg/day Mefloquine 500mg/day Adult (<35kg) D1-D3: (Artequiner®) Artesunate 100mg q24h Mefloquine 250mg q24h OR Riamet® (1 tablet: 20mg artemether/120mg lumefantrine) Adult (>35kg) D1: 4 tablets stat then again 4 tablets at 8 hours later D2-3: 4 tablets q12h (am, pm) (total course =24 tablets) Adult (<35kg) D1: 3 tablets stat then again 3 tablets at 8 hours later D2-3: 3 tablets q12h (am, pm) (total course = 18 tablets) | Quinine 10mg/kg PO q8h PLUS/MINUS Doxycycline 100mg PO q12h for 7 days | The choice of drug should be governed by drug availability and safety. Artemisinin derivatives are contraindicated in pregnancy; use quinine If gametocytes continue to be present at D7 onwards, Primaquine 30mg as a single dose may be given (check G6PD status before use). Patient may be discharged home |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| ii) Treatment Failure | Artemether/Lumefantrine (as above) PLUS Doxycycline 100mg PO q12h for 7 days | Quinine 10mg/kg PO q8h PLUS Doxycycline 100mg PO q12h for 7-10 days | Mefloquine should not be taken for a second time within 28 days (neuropsychiatric side effects) In pregnancy: Quinine 10mg/kg PO q8h PLUS Clindamycin 600mg PO q12h for 7-10 days |
| b) Complicated (see definition above) | D1: Artesunate 2.4mg/kg IV stat then second dose 1.2mg/kg at 12 hours D2-D7: Artesunate 1.2mg/kg IV q24h OR D1: Quinine 7mg/kg IV in 100ml N/S over 1 hour then 10mg/kg in 250-500ml D5% over 4 hours Then: Quinine 10mg/kg IV q8h (can give orally if tolerated) PLUS Doxycycline 100mg PO q12h for 7 days | D1: Loading dose Quinine IV 20mg/kg over 4 hours in D5% Then: Quinine 10mg/kg IV q8h (can give orally if tolerated) PLUS Doxycycline 100mg PO q12h for 7 days | Patient should be managed in an intensive care facility. Monitor patient's blood glucose and ECG while on IV quinine In pregnancy: Use Quinine IV regime and Clindamycin 600mg q12h as a substitute to Doxycycline In renal failure: Use 1/2-1/3 of the dose of Quinine. May maintain normal dose if patient receives dialysis. Watch out for toxicity |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|--|---|
| | Preferred | Alternative | |
| Plasmodium Vivax or Ovale | Chloroquine 10mg/kg (max 600mg) stat then 5mg/kg (max 300mg) 6 hours later, D2 and D3 PLUS Primaquine 15mg/day PO for 14 days | Treatment failure: Repeat Chloroquine as first line PLUS Primaquine 15mg PO q12h for 14 days | Usually benign presentation. Check G6PD before starting Primaquine as it may cause haemolysis in G6PD deficient |
| Plasmodium Malariae/Knowlesi | Chloroquine 10mg/kg (max 600mg) stat then 5mg/kg 6 hours later, D2 and D3 | Severe cases: Treat as complicated Plasmodium Falciparum | |
| Mixed Infection | Treat as Plasmodium Falciparum (see above) | | |
| Chemoprophylaxis | Mefloquine 250mg weekly (up to 1 year) | Doxycycline 100mg q24h (up to 3 months) | To start 1 week before and continued till 4 weeks after leaving the area |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

MANAGEMENT OF TUBERCULOSIS

(Adapted from *Practice Guidelines For The Management of Tuberculosis, Ministry of Health Malaysia, 2nd edition 2002*)

1. Drugs

Five drugs are considered essential (1st line) for the treatment of tuberculosis. These are Isoniazid (H), Rifampicin (R), Pyrazinamide (Z), Streptomycin (S) and Ethambutol (E).

- * Isoniazid (H),
 - * Rifampicin (R),
 - * Pyrazinadine (Z),
 - * Streptomycin (S) &
 - * Ethambutol (E).
- Essential 1st line drugs**

2. Treatment regimens

Treatment regimens are divided into:

- (i) Initial or intensive phase.
- (ii) Continuation or maintenance phase.

During the intensive phase, three or four drugs are given daily. This leads to rapid sputum conversion and amelioration of clinical symptoms. During the continuation phase, two or three drugs are usually given intermittently. The sterilising effect of the therapy eliminates remaining bacilli and reduces drastically the chances of subsequent relapse.

Category I: New Case

- (i) Intensive phase: 2SHRZ or 2EHRZ or 2HRZ (2 months of daily doses).
- (ii) Continuation phase: 4H₂R₂ or 4S₂H₂R₂ or 4HR or 4H₃R₃ or 4S₃H₃R₃ (Duration may be extended for severe forms of extra pulmonary tuberculosis and immunocompromised patients).

*The number preceding the treatment regimen refers to the treatment duration in months.

**The subscript below the drug symbol refers to the frequency of doses per week.

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Category II: Relapse, Treatment failure, Treatment after interruption

- (i) Send *Mycobacterium tuberculosis* culture and sensitivity (MTB C&S) (Rapid culture method if available).
- (ii) Do not initiate standard therapy.
- (iii) Refer to chest physician or physician in charge of chest clinic.
- (iv) Subsequent drug regimen based on sensitivity results and clinical response.

Category III: Chronic Case

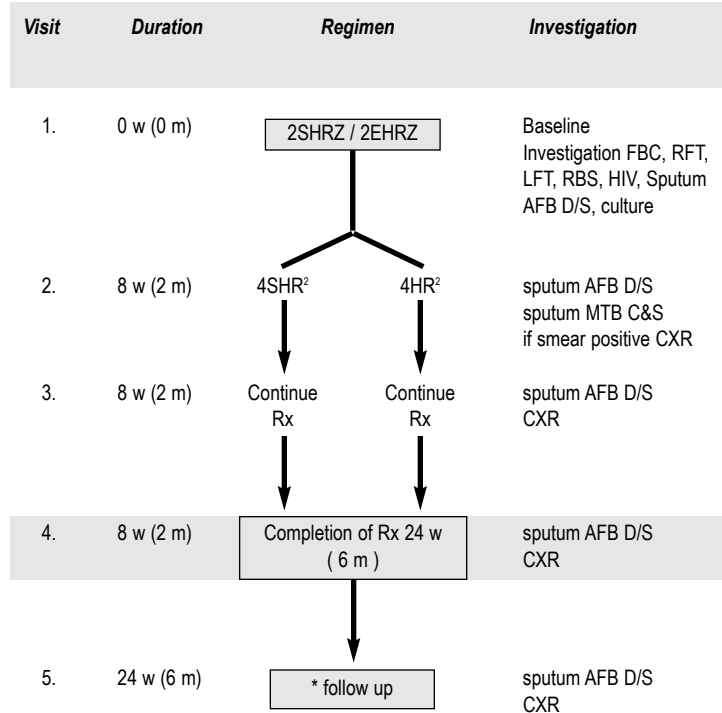
- (i) Send *Mycobacterium tuberculosis* culture and sensitivity (MTB C&S) (Rapid culture method if available).
- (ii) Refer to chest physician or physician in charge of chest clinic.

3. Anti-tuberculosis drugs (1st line) and the recommended dosages

| 1st line drug | Daily dosage | | Biweekly dosage | |
|------------------|--------------|----------|-----------------|----------|
| | mg/kg | max (mg) | mg/kg | max (mg) |
| Isoniazid (H) | 5 - 8 | 300 | 15 - 20 | 1200 |
| Rifampicin (R) | 10 - 15 | 600 | 15 - 20 | 600 |
| Pyrazinamide (Z) | 20 - 40 | 1500 | 50 | 2000 |
| Ethambutol (E) | 15 - 25 | 1200 | 50 | 2000 |
| Streptomycin (S) | 15 - 20 | 1000 | 15 - 20 | 1000 |

Note: For patients more than 65 years of age, the dose of streptomycin should not exceed 750 mg.

4. Flow chart for recommended 24 weeks (w) / 6 months (m) treatment regimen (adult)



| | | |
|-------------------------|----------------------------------|---------------------------------|
| E = Ethambutol | FBC = Full blood count | RBS = random blood sugar |
| H = Isoniazid | LFT = Liver function test | HIV = anti-HIV antibody |
| R = Rifampicin | RFT = Renal function test | (for screening) |
| S = Streptomycin | D/S = Direct smear | MTB = Mycobacterium |
| Z = Pyrazinamide | Rx = Treatment | tuberculosis |
| W = week | | C&S = culture and |
| M = month | | sensitivity test |

Note: (*) Recommended to be done where facilities are available

5. Management of Tuberculosis in Special Situations

A. Tuberculosis during pregnancy and lactation

Untreated tuberculosis presents a much greater risk to a pregnant woman and her foetus than does the treatment of the disease. Standard treatment using Isoniazid, Rifampicin, Pyrazinamide and Ethambutol is used. Doses of anti-tuberculosis drugs given in pregnancy are similar to that in a non-pregnant patient. Streptomycin is best avoided because of the risk of ototoxicity to the foetus. Normal recommended dosages of Rifampicin are safe in pregnant patients.

Tuberculosis treatment in lactating mothers is safe as the amount of drug ingested by nursing infant is minimal. If the mother at the time of delivery is smear-positive, the newborn should be separated from the mother at least for a period of two weeks.

Breast-feeding is best avoided during these two weeks and expressed milk should be given to the child. BCG should be given as scheduled and Isoniazid prophylaxis should be given for 6 months followed by Mantoux test at the end of 6 months. In the event of absence of scar, BCG vaccination should be repeated. When there is doubt about the presence of active tuberculosis, the child should be treated.

Congenital tuberculosis, although rare should be suspected if an infant born to a tuberculous mother fails to thrive, has non-specific symptoms such as fever, respiratory distress, poor feeding and vomiting, or has suggestive signs such as hepatosplenomegaly.

B. Tuberculosis treatment for women taking the oral contraceptive pill

Rifampicin interacts with the oral contraceptive pill, with a risk of decreased protective efficacy against pregnancy. A woman who usually takes the oral contraceptive pill may choose between an oral contraceptive pill containing a higher dose of oestrogen (50mcg) or use another form of contraception after consultation with a doctor.

C. Tuberculosis in patients with liver impairment

Patients with no evidence of chronic liver disease (e.g. hepatitis virus carrier, past history of acute hepatitis and alcoholics) can receive the usual short-course chemotherapy regimens but therapy should be modified in patients with established chronic liver disease and acute hepatitis. These cases are best referred to specialists for management.

i) Established chronic liver disease

The following regimens are recommended:

- (i) 2SHRE/7H₂R₂
- (ii) 2SHE/10HE
- (iii) 2SH/12S₂H₂

ii) Acute hepatitis (e.g. acute viral hepatitis)

It is a rare eventuality that a patient has tuberculosis and also at the same time acute hepatitis unrelated to tuberculosis or anti-tuberculosis treatment. Clinical judgement is necessary. In some cases it is possible to defer tuberculosis treatment until the acute hepatitis has resolved. In other cases when it is necessary to treat tuberculosis during acute hepatitis, the safest regimen is 3SE/6HR.

D. Tuberculosis in patients with renal impairment

Isoniazid, Rifampicin and Pyrazinamide are either eliminated almost entirely by biliary excretion or metabolised into non-toxic compounds. These drugs can, therefore, be given in normal dosage to patients with renal failure. Streptomycin and Ethambutol are excreted by kidney. Where facilities are available to monitor renal function closely it may be possible to give Streptomycin and Ethambutol in reduced doses. The safest regimen to be administered in patients with renal failure is 2HRZ/6HR.

E. Extra pulmonary tuberculosis

The regimen of treatment is similar as for pulmonary tuberculosis but the duration may be extended and it varies from 6 months to 12 months or longer depending on the clinical response of the individual patient, for example in tuberculosis meningitis, it is advisable to treat the patient for at least 12 months.

Steroids should be given in tuberculous meningitis, genitourinary tract tuberculosis and may also be considered in miliary tuberculosis.

F. Tuberculosis in patients with HIV infection

Recommended treatment regimens for patients who have tuberculosis with HIV infections (The recommendations are based on those of the CDC, Davidson and The American Thoracic Society-modified)

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Clinical presentation of TB in HIV/AIDS (from chemotherapy guideline 1994)

| Clinical situation | Treatment |
|---|--|
| Initial therapy <ul style="list-style-type: none"> ◆ No suspicion of drug resistance ◆ Possible drug resistance | Isoniazid, Rifampicin, Pyrazinamide daily Isoniazid, Rifampicin, Pyrazinamide, Ethambutol daily |
| Long-term therapy <ul style="list-style-type: none"> ◆ Drug-susceptible organisms ◆ Isoniazid resistance or intolerance ◆ Rifampicin resistance or intolerance | Isoniazid, Rifampicin, Pyrazinamide for 2 months daily followed by Isoniazid, Rifampicin for 7 months biweekly or for 6 months after cultures are negative, whichever is longer. Avoid protease inhibitor if regimen contains Rifampicin. Rifampicin, Ethambutol and Pyrazinamide daily for 2 months followed by Rifampicin and Ethambutol daily for 12-16 months or 12 months after cultures are negative, whichever is longer. Isoniazid, Pyrazinamide, Ethambutol daily for 18 months to 24 months, or for 12 months after cultures are negative whichever is longer. |

URINARY TRACT INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|-----------------------------------|
| | Preferred | Alternative | |
| Acute Uncomplicated Cystitis <i>E. Coli</i> <i>Enterobacteriaceae:</i> <i>Klebsiella</i> <i>Proteus</i> <i>Enterobacter species</i> <i>Staphylococcus - saprophyticus</i> <i>Enterococcus</i> | Trimethoprim 300mg PO q24h for 7 days | Cefuroxime 250mg PO q12h for 7 days OR Nitrofurantoin 50mg PO q6h for 7 days OR *Trimethoprim/Sulphamethoxazole 160/800mg PO q12h for 3 days | *Avoid sulfonamides in pregnancy |
| Acute Cystitis in Pregnancy | Cefuroxime 250mg PO q12h for 7 days | Nitrofurantoin 50mg PO q6h for 7 days OR Cephalexin 500mg PO q12h for 7 days OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 625mg PO q12h for 7 days | Modify treatment based on culture |
| Recurrent Urinary Tract Infections: > 3 episodes/year | Trimethoprim/Sulphamethoxazole 80/400mg PO ON for 3-12 months | Nitrofurantoin 50mg PO ON for 3-12 months OR Cephalexin 250mg PO ON for 3-12 months OR Trimethoprim 100mg PO ON for 3-12 months | As Prophylaxis |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| Acute Uncomplicated Pyelonephritis <i>E. Coli</i> <i>Enterobacter</i> <i>Proteus</i> <i>Pseudomonas</i> | If ill, hospitalised Cefuroxime 750mg IV q8h for 2 weeks PLUS/MINUS Gentamicin ¹ 5mg/kg IV q24h for 2 weeks <i>(If use of aminoglycosides deemed undesirable, consider 3rd generation Cephalosporins)</i> | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 1-2g IV q24h for 2 weeks OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h for 2 weeks OR Ciprofloxacin 500-750mg PO q12h | Adjust according to culture & sensitivity May step down to oral antibiotic following clinical improvement (afebrile for 48 hours) |
| Acute Complicated Pyelonephritis Calculi especially struvite stones Urethral stricture or tumour Papillary necrosis Congenital abnormalities Neuropathic bladder Previous genito-urinary surgery predisposing to obstruction Polycystic kidneys <i>E. Coli</i> <i>Proteus sp.</i> <i>Klebsiella</i> <i>Pseudomonas</i> <i>Serratia</i> <i>Enterococci</i> | If ill, hospitalised Cefuroxime 750mg IV q8h PLUS Gentamicin ¹ 5mg/kg IV q24h for 2 weeks If Enterococci Ampicillin 500mg IV q6h PLUS Gentamicin ¹ 5mg/kg IV q24h for 2 weeks | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 1-2g IV q24h for 2 weeks OR <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate 1.2g IV q8h OR Piperacillin/Tazobactam 4.5g IV q8h for 2 weeks OR Ciprofloxacin 200mg IV q12h for 2 weeks | Adjust according to culture sensitivity |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|---------------------------------------|---|---|
| | Preferred | Alternative | |
| Acute Pyelonephritis in Pregnancy | Cefuroxime 750mg IV q8h for 2 weeks | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 1.2g IV q8h for 2 weeks</i> OR <i>3rd gen. Cephalosporins, e.g. Ceftriaxone 1-2g IV q24h for 2 weeks</i> | |
| Asymptomatic Bacteriuria <i>E. Coli</i> in 75% of elderly patients <i>Proteus</i> <i>Klebsiella</i> <i>Enterobacter</i> <i>Pseudomonas</i> | Trimethoprim 300mg PO q24h for 7 days | Cefuroxime 250mg PO q12h for 7 days OR Nitrofurantoin 50mg PO q6h for 7 days OR *Trimethoprim/Sulphamethoxazole 160/800mg PO q12h for 3 days | Recommendation for treatment is only for the following conditions:- a) Pregnant women if test results are positive b) Patients who undergo traumatic urologic interventions with mucosal bleeding, and such patients should be treated prior to such interventions c) Before transurethral resection of the prostate *Avoid sulfonamides in pregnancy |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---------------------------------------|
| | Preferred | Alternative | |
| Asymptomatic Bacteriuria in Pregnancy | Cefuroxime 250mg PO q12h for 7 days | Nitrofurantoin 50mg PO q6h for 7 days OR <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin/Clavulanate 625mg PO q12h for 7 days</i> | Avoid Quinolones |
| Catheter Related Bacteriuria | Antibiotics not recommended for asymptomatic bacteriuria | | Remove or change catheter if possible |
| Acute Prostatitis | Refer to Page 129 (Urology) | | |
| Chronic Prostatitis | Refer to Page 129 (Urology) | | |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

References:

1. The Management of Urinary and Male Genital Tract Infections. European Association of Urology 2006
2. Antibiotic Guidelines 2000/2001, Hospital Kuala Lumpur
3. Use of Antibiotics in Adults: CPG Guidelines, Ministry of Health, Singapore, 2006
4. MIMS Antimicrobial Guide: Malaysia 2005/2006 3rd Edition

SECTION B: PAEDIATRICS



CARDIOVASCULAR INFECTIONS

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| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| 1. Acute Myocarditis | | | |
| Commonly caused by viruses | Treatment mainly supportive | | Reference: 1, 2 |
| 2. Acute pericarditis | | | |
| Viral (commonest cause) | Treatment mainly supportive | | Consider surgical drainage if pericardial empyema detected |
| Bacterial: <i>Staphylococcus aureus</i> | Cloxacillin 200mg/kg/24h IV in 4-6 divided doses for 6 weeks PLUS/MINUS Gentamicin ¹ 1mg/kg IV/IM q8h for 3 - 5 days | Penicillin allergic: Cefazolin 100mg/kg/24h IV in 3 equally divided doses OR Vancomycin ¹ 40mg/kg/24h IV in 2-4 divided doses | Reference: 3, 4 |
| 3. Infective Endocarditis | | | |
| Empirical Therapy for Infective Endocarditis | Benzylpenicillin 200,000 units/kg/24h IV in 4-6 equally divided doses for 4 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks | Vancomycin ¹ 15mg/kg q12h IV for 4-6 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks | Reference: 3, 4 |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--|
| | Preferred | Alternative | |
| Infective Endocarditis caused by <i>Streptococcus Viridans</i> | Benzylpenicillin 200,000 units/kg/24h IV in 4-6 equally divided doses for 4 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone 100mg/kg IV/IM q24h for 4 weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks For patients allergic to Penicillin or Ceftriaxone: Vancomycin ¹ 40mg/kg/24h IV in 2-3 equally divided doses for 4 weeks | Dosages suggested are for patients with normal renal and hepatic function. Maximum dosages per 24 hours: Penicillin 18 million units; Ampicillin 12g; Ceftriaxone 4g, Gentamicin 240 mg. <i>Reference: 8, 9</i> |

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| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Infective Endocarditis caused by Enterococcus | Benzylpenicillin 300,000 units/kg/24h IV in 4-6 equally divided doses OR Ampicillin 300mg/kg/24h IV in 4-6 divided doses for 4-6weeks PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 4-6 weeks | Penicillin allergic: Vancomycin ¹ 40mg/kg/day IV in 2-3 equally divided doses PLUS Gentamicin ¹ 1mg/kg IV/IM q8h for 2 weeks for 6 weeks | Reference: 8, 9 |
| Infective Endocarditis Caused by <i>Staphylococcus</i> a) Methicillin sensitive | Cloxacillin 200mg/kg/24h IV in 4-6 divided doses for 6 weeks PLUS/MINUS Gentamicin ¹ 1mg/kg IV/IM q8h for 3-5 days | | Clinical benefit of aminoglycosides has not been established. Cefazolin or other first-generation cephalosporin in equivalent dosages may be used in patients who do not have a history of immediate type hypersensitivity (urticaria, angioedema, anaphylaxis) to penicillin or ampicillin. |
| | b) Penicillin allergic | Cefazolin 100mg/kg/24h IV in 3 equally divided doses for 6 weeks | |

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| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|-------------|--|
| | Preferred | Alternative | |
| c) Methicillin Resistant | Vancomycin ¹ 40mg/kg/24h IV in 2-4 divided doses for 6 weeks | | Reference: 4, 8, 9 |
| Culture-Negative Endocarditis | <p><i>β-lactam/β-lactamase inhibitors, e.g. Ampicillin/Sulbactam 300mg/kg/24h IV in 4-6 equally divided doses for 4-6 weeks</i></p> <p>PLUS Gentamicin¹ 1mg/kg IV/IM q8h for 4-6 weeks</p> | | <p>Patients with culture-negative endocarditis should be treated in consultation with an ID specialist</p> <p>Reference: 4, 8, 9</p> |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

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CENTRAL NERVOUS SYSTEM INFECTIONS

NATIONAL ANTI-BIOTIC GUIDELINE 2008

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|---|---|
| | Preferred | Alternative | |
| Meningitis empirical treatment | Benzylpenicillin 50mg/kg IV q4-6h PLUS <i>3rd gen. Cephalosporins, e.g.</i> *Cefotaxime OR *Ceftriaxone IV for 10-14 days | Vancomycin ¹ 15mg/kg IV q6h PLUS <i>3rd gen. Cephalosporins, e.g.</i> *Cefotaxime OR *Ceftriaxone for 10-14 days | Reference: 1, 2, 5 |
| H. influenza | <i>3rd gen. Cephalosporins, e.g.;</i> *Cefotaxime OR *Ceftriaxone IV for 10-14 days | Chloramphenicol 40mg/kg IV stat then 25mg/kg q6h for 10-14 days; | Prophylaxis for all household contacts if there are unimmunised or partially immunised children < 4 years old (Red Book 2006) |
| Strep Pneumoniae** | if MIC < 0.1 mg/L: Benzylpenicillin 50mg/kg IV q4-6h for 10-14 days if MIC 0.1- to < 2mg/L <i>3rd gen. Cephalosporins, e.g.</i> *Cefotaxime OR *Ceftriaxone for 10-14 days If MIC > 2mg/L Vancomycin ¹ PLUS <i>3rd gen. Cephalosporins</i> for 10-14 days | OR Cefepime 50mg/kg IV q8h for 10-14 days | |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|---|--|--|
| | Preferred | Alternative | |
| Neisseria meningitidis** | Benzylicillin 50mg/kg IV q4-6h for 7 days | 3 rd gen. Cephalosporins, e.g. *Cefotaxime OR *Ceftriaxone IV for 7 days; OR Chloramphenicol 40mg/kg stat then 25mg/kg IVq6h | Prophylaxis for all household contacts and Health care workers involved in intubation and suctioning of airway |
| Herpes Simplex encephalitis | Acyclovir: 12 weeks-12 years old: 500mg/m ² q8h If > 12 years olds: 10mg/kg IV q8h Duration: for 14-21 days | | Reference: 3, 4 |
| Brain Abscess | 3 rd gen. Cephalosporins, e.g. *Cefotaxime OR *Ceftriaxone PLUS Metronidazole 15mg/kg IV stat then 7.5mg/kg IV q8h (duration of antibiotic would depends on response by neuroimaging; 4-8 weeks may be needed) | Add Cloxacillin if secondary to trauma | Surgical drainage may be indicated if appropriate Reference: 4 |

*Cefotaxime 50mg/kg q4-6h (severe infection)

*Ceftriaxone 50mg/kg q12h (severe infection)

** Duration of antibiotic may need to be extended as a result of complications subdural empyema or brain abscess

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CHEMOPROPHYLAXIS

A. NON-SURGICAL

| Condition/Infection & Likely Organism | Prophylactic Regimen | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Rheumatic fever (Secondary prevention) | Benzathine Penicillin IM 1.2 mega units (>25kg); 0.6 mega units (<25 kg) every 3-4 weeks Duration <u>With carditis:</u> 10 years or until 25 years of age <u>Without carditis:</u> 5 years or until 18 years of age OR Cephalexin 50mg/kg PO 1 hour prior to procedure | Gentamicin ¹ 1.5mg/kg IV within Phenoxymethylpenicillin 250mg PO q12h <u>Penicillin allergy</u> EES 400mg PO q12h | <i>Reference: 1</i> |
| Infective Endocarditis | Dental, oral, respiratory or esophageal procedures: Amoxicillin 50mg/kg PO 1 hour before procedure | <u>Penicillin allergy</u> Clindamycin 20mg/kg PO 1 hour before procedure OR Azithromycin/Clarithromycin: >10 years old = 500mg >5 and <10 yrs = 300mg <5 yrs = 200mg OR 15mg/kg 1 hour before procedure OR Cephalexin 50mg/kg PO 1 hour prior to procedure | Prophylaxis recommended for high risk and moderate risk categories and for specific procedures (as described in AHA Recommendations reference 2, 3, 4) <i>Reference: 2</i> |

| Condition/Infection & Likely Organism | Prophylactic Regimen | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| | <p>Genitourinary or gastrointestinal procedures:</p> <p><i>High risk:</i> Ampicillin 50mg/kg IV PLUS Gentamicin¹ 1.5mg/kg IV within 30 minutes prior to procedure Followed by: (Repeat Ampicillin 25mg/kg PO 6 hours later)</p> <p><i>Moderate risk:</i> Amoxicillin 50mg/kg PO 1 hour before procedure</p> | | |
| <p>Post-splenectomy</p> <p><i>At risk for pneumococcus, meningococcus, Haemophilus</i></p> | <p>Phenoxymethylpenicillin: < 5 yrs: 125mg PO q12h > 5yrs: 250mg PO q12h</p> <p><u>Duration:</u></p> <ul style="list-style-type: none"> ◆ Children up to the age of 16 years ◆ Post-splenectomy for at least 2-3 years ◆ Indefinitely for patients with an underlying immunocompromised state and asplenia <p><i>(Require ongoing surveillance for resistant pneumococci)</i></p> | <p>Amoxicillin 20mg/kg/24h PO</p> <p><u>Penicillin allergy:</u> EES < 2 yrs: 200mg PO q24h > 2 yrs: 400mg PO q24h</p> | <p>Risk of sepsis is lifelong, but especially the first 2 years after splenectomy</p> <p>Important adjunct: Immunisation against <i>pneumococcus, haemophilus, meningococcus</i> prior to splenectomy</p> <p>To seek immediate medical attention when febrile</p> <p><i>Reference: 5, 6, 16</i></p> |

| Condition/Infection & Likely Organism | Prophylactic Regimen | | Comments |
|---------------------------------------|---|-------------|--|
| | Preferred | Alternative | |
| H. influenza B exposure | Rifampicin PO <u>Children:</u> 20mg/kg q24h x 4 days <u>Infants:</u> 10mg/kg q24h x4 days | | <p><u>Household contacts</u> If there is one unvaccinated contact ≤ 4 years old in the household, RIF recommended for all household contacts except pregnant women</p> <p><u>Nursery Contact</u></p> <ul style="list-style-type: none"> ◆ With 1 case, if attended by unvaccinated children ≤ 2 yrs, consider prophylaxis + vaccinate susceptibles ◆ If all contacts > 2 yrs: no prophylaxis ◆ If ≥ 2 cases in 60 days and unvaccinated children attend, prophylaxis recommended for children and personnel ◆ Give chemoprophylaxis to index case if treated with regimens other than cefotaxime or ceftriaxone ◆ Contacts < 2 years not immunised: complete immunisation <p><i>Reference: 7</i></p> |

| Condition/Infection & Likely Organism | Prophylactic Regimen | | Comments |
|---------------------------------------|--|--|---|
| | Preferred | Alternative | |
| Meningococcal exposure | Rifampicin PO <u>Children:</u> <1 month: 5mg/kg q12h for 2 days >1 month: 10mg/kg (max 600mg) q12h for 2 days | <i>3rd gen. Cephalosporins, e.g.</i> Ceftriaxone IM <15 yrs: 125mg stat >15 yrs: 250mg stat Ciprofloxacin PO >18 yrs 500mg single dose | CLOSE contact: All household, child care and nursery contacts. <u>Others</u> <ul style="list-style-type: none"> ◆ Close contact for at least 4 hours during the week before illness onset ◆ Exposure to index's nasopharyngeal secretions (eg kissing, sharing of toothbrushes, eating utensils) ◆ Airline flights lasting >8 hours: directly next to case <u>Healthcare staff</u> Routine prophylaxis not recommended, unless exposure to secretions such as unprotected mouth to mouth resuscitation, intubation or suctioning <i>Reference: 8</i> |
| UTI prophylaxis | Refer to Page 202 (Urinary Tract Infections) | | |

| Condition/Infection & Likely Organism | Prophylactic Regimen | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| Neonatal Group B Strep (GBS) Infection <i>Treat during labour if previously delivered infant with invasive GBS, GBS bacteriuria or screening swabs positive OR if</i> <ul style="list-style-type: none"> ◆ Preterm <37 weeks ◆ PROM >18 hours ◆ Intrapartum temp >38°C | Intrapartum maternal prophylaxis till delivery Benzylpenicillin 5 mega units IV load then 2.5 mega units q6h | Ampicillin 2g IV load then 1g q6h <u>Penicillin allergy:</u> Erythromycin 500mg IV q6h (according to susceptibility) | <i>Reference: 12</i> |
| Malaria prophylaxis | Mefloquine 5mg/kg PO once a week To start one week before and continued till 4 weeks after leaving the area | Doxycycline 2mg/kg PO q24h (max 100mg/day) in children >8 years old OR Clindamycin 10mg/kg q12h in children < 8 years and in pregnancy To start one week before and continued till 4 weeks after leaving the area | <i>Reference: 13</i> |
| Pertussis (Post-exposure prophylaxis) | EES 20mg/kg PO q12h (max.400mg/day) for 10-14 days | | Prophylaxis for all household and close contacts irrespective of age and immunization status Complete immunization for close contact ≤ 7 years of age <i>Reference: 14</i> |

| Condition/Infection & Likely Organism | Prophylactic Regimen | | Comments |
|---|---|-------------|--|
| | Preferred | Alternative | |
| Chicken pox (<i>Post-exposure prophylaxis</i>) | *Varicella-Zoster Immune Globulin (VZIG) (125 units/10kg, max 625 units) OR Intravenous Immunoglobulin (IVIG) (400mg/kg) within 96 hours <i>Post-exposure</i> varicella vaccine may have some benefit | | Susceptible hosts include: <ul style="list-style-type: none"> ◆ Neonate where maternal varicella develops 5 days before and 2 days after delivery ◆ Immunocompromised hosts ◆ Hospitalized premature infants: <ul style="list-style-type: none"> - <28 weeks regardless of maternal history of varicella - >28 weeks: whose mothers lack reliable history of varicella *Requires DG approval <i>Reference: 13, 15, 16</i> |
| Tuberculosis | <5yrs Isoniazid 5mg/kg/24h for 6 months | | Newborns: BCG after 6 months of prophylaxis Follow-up every 2 months If child confirmed positive, treat Prophylaxis > 5 years not recommended If child HIV positive, suggest prophylaxis irrespective of age <i>Reference: 17</i> |

¹Refer Appendix 1(Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

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GASTROINTESTINAL INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Acute Gastroenteritis <i>Usually viruses eg rotavirus</i> | Antibiotics not recommended | | Oral rehydration is the cornerstone of treatment Antibiotic therapy may prolong carriage state of salmonellosis <i>Reference: 1</i> |
| Dysentery <i>Shigella, E. coli, Campylobacter</i> | Most mild infections resolved spontaneously without antibiotics Trimethoprim/Sulphamethoxazole (TMP: 5-8mg/kg/24h) PO in 2 divided doses for 5-7 days OR Ampicillin 100mg/kg/24h PO in 4 divided doses for 5-7 days | If severe: <i>3rd gen. Cephalosporins, e.g. Cefotaxime 150-200mg/kg/24h IV in 4 divided doses for 7 days</i> | <i>Reference: 2</i> |
| Dysentery <i>Amoebiasis</i> | Metronidazole 30-50mg/kg/24h PO in 3 divided doses for 5 days (10 days for severe infection) | | <i>Reference: 2</i> |
| Giardiasis | Metronidazole 15mg/kg/24h PO in 3 divided doses for 5 days | | <i>Reference: 2</i> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| Cholera | Trimethoprim/Sulphamethoxazole 8-10mg (TMP)/kg/24h PO in 2 divided doses for 3 days OR Tetracycline 50mg/kg/24h PO q6h for 3 days (children > 8 years) OR Doxycycline 6mg/kg (max. 300mg) PO q24h (children > 8 years) | Erythromycin 50mg/kg/24h PO in 4 divided doses for 3 days (for strains resistant to tetracyclines) <i>Single dose Azithromycin or Ciprofloxacin may be considered in special circumstances (e.g. during major outbreaks)</i> | Oral rehydration is the cornerstone of treatment. Antibiotics therapy reduces the volume and duration of diarrhoea Avoid using Tetracycline or Doxycycline for young children as they can cause staining of the teeth <i>Reference: 3, 4, 5, 6, 7</i> |
| Liver abscess (amoebic) <i>Entamoeba histolytica</i> | Metronidazole 35-50mg/kg/24h IV in 3 divided doses for 10-14 days | | Amoebic abscess tend to be solitary lesion. Consider surgical drainage if needed <i>Reference: 11, 12</i> |
| Liver abscess (pyogenic) <i>Gram-ve, Anaerobic, S. aureus</i> | Ampicillin 150-200mg/kg/24h IV in 4 divided doses PLUS Gentamicin ¹ 5mg/kg IV q24h PLUS Metronidazole 10mg/kg IV q8h | <i>3rd gen. Cephalosporins, e.g. Cefotaxime 50mg/kg IV q6h</i> PLUS Metronidazole 35-50mg/kg/24h IV in 3 divided doses | Surgical drainage is needed in most cases <i>Reference: 11, 12</i> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|--------------------------|
| | Preferred | Alternative | |
| | <p><i>If S. aureus:</i> Cloxacillin 150-200mg/kg/24h IV in 4-6 divided doses PLUS Gentamicin¹ 5mg/kg IV q24h for 4-6 weeks</p> | | |
| <p>Acute cholangitis Gram negative, anaerobes, gram positive</p> | <p>Ampicillin 150-200mg/kg/24h IV in 4 divided doses PLUS Gentamicin¹ 5mg/kg IV q24h PLUS Metronidazole 10mg/kg IV q8h for 7 days</p> | <p><i>3rd gen. Cephalosporins, e.g.</i> Cefoperazone 50mg/kg IV q8h PLUS Metronidazole 10mg/kg IV q8h</p> | <p>Reference: 11, 12</p> |
| <p>Peritonitis (Primary) Strep. Pneumoniae, gram-neg organisms</p> | <p>Ampicillin 150-200mg/kg/24h IV in 4 divided doses PLUS Gentamicin¹ 5mg/kg IV q24h for 7 days</p> | <p><i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 150-200mg/kg/24h IV in 4 divided doses</p> | <p>Reference: 11, 12</p> |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

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INFECTIONS IN IMMUNUCOMPROMISED PATIENTS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|---|
| | Preferred | Alternative | |
| First Line Febrile neutropenia Fever >38°C Neutrophil<500mm ³ <i>Klebsiella sp,</i> <i>E.coli, Pseudomonas</i> | Cefepime 100-150mg/kg/24h IV in 3 divided doses | Piperacillin/Tazobactam 300-360mg/kg/24h IV in 3-4 divided doses | Meta analysis has shown that there is no clinical advantage with β lactam-aminoglycoside combination therapy ¹ |
| Second line Persistent fever > 72 hours <i>MRSA</i> <i>coagulase -ve staph</i> | Imipenem 20mg/kg IV q8h PLUS/MINUS Vancomycin ¹ 15mg/kg IV q6h | Meropenem 20mg/kg IV q8h PLUS/MINUS Vancomycin ¹ 15mg/kg IV q6h | Consider adding Vancomycin in suspected catheter related infections, positive blood culture for gram +ve cocci, hypotension patients and patients who are known to be colonised with MRSA |
| Third Line Fever > 5 days <i>Candida sp</i> <i>Aspergillus sp</i> | Imipenem 20mg/kg IV q8h PLUS Amphotericin B 0.5mg/kg IV and gradually escalate by 0.25 to 1mg/kg q24h | Meropenem 20mg/kg IV q8h PLUS Amphotericin B 0.5mg/kg IV and gradually escalate by 0.25 to 1mg/kg q24h | 1/3 of febrile neutropenia patients with persistent fever >1 week have systemic fungal infections ² |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

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NEONATAL INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|---|
| | Preferred | Alternative | |
| Congenital Infections | | | |
| Congenital Syphilis <i>T pallidum</i> | Benzympenicillin 50,000 units/kg IV q12h for the first 7 days of life and q8h thereafter for 10-14 days | Procaine Benzympenicillin 50,000 units/kg IM q24h in a single dose for 10-14 days | <ul style="list-style-type: none"> ◆ Isolate till non-infectious (at least 24 hours of treatment) ◆ Screen for other STDs and HIV ◆ Investigate and treat parents <p><i>Follow-up</i> Nontreponemal serologic tests at 3, 6, 12 and 24 months. (Should become -ve by 6 months)</p> <p>For those with abnormal CSF - recommended to repeat CSF FEME and VDRL at 6 months intervals. Persistent +VDRL of CSF requires reevaluation and possible re-treatment</p> <p><i>Reference: 1, 2</i></p> |

NATIONAL ANTI-BIOTIC GUIDELINE 2008

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Congenital Toxoplasmosis <i>T. gondii</i> | *Pyrimethamine Initial loading dose of 2mg/kg PO q24h for 2 days followed by 1mg/kg PO q24h (<i>maximum 25mg</i>) for 6 months, then 3x/wk for subsequent 6 months PLUS Sulfadiazine 50mg/kg PO q12h (<i>maximum 4g</i>) for 1 year PLUS Folinic Acid 10mg PO 3 times/wk for 1 year <i>(IV formulation of Folinic Acid may be considered for oral use)</i> | *Pyrimethamine 1.25mg/kg PO every 15 days for 24 months PLUS Folinic acid 5mg/week PO | Drug regimen not definitively established. Clinical trials ongoing Prednisone (1mg/kg/day) can be used when active chorioretinitis involves the macula or otherwise threatens vision *Fansidar (Sulfadoxine/Pyrimethamine) contains 25mg Pyrimethamine <i>Reference: 4, 5, 6</i> |
| Herpes Simplex | Acyclovir 20mg/kg IV q8h Duration: Skin, eyes, mouth: 14 days CNS/Disseminated: 21 days | | <ul style="list-style-type: none"> ◆ Isolate ◆ Ocular involvement requires topical antiviral ◆ Screen for other STDs ◆ For CNS disease repeat LP at end of therapy for HSV PCR and treat till negative ◆ Investigate and treat parents <i>Reference: 7, 8</i> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|---|--|--|
| | Preferred | Alternative | |
| Tetanus neonatorum | <p>Metronidazole 5-30mg/kg/24h PO in 2-3 divided doses for 7 days, not to exceed 2g/24h</p> <p>Weight-based dosing: Body weight <2000g 0-7 days: 7.5mg PO/IV q24h 8-28 days: 7.5mg PO/IV q12h</p> <p>Body weight >2000g 0-7 days: 7.5mg PO/IV q12h 8-28 days: 15mg PO/IV q12h</p> <p>Duration: Metronidazole PO/IV for 10 days</p> | <p>Benzylpenicillin 100,000 units/kg IV q12h for 1st wk of life and q6h after 1st wk for 10 days</p> | <ul style="list-style-type: none"> ◆ Debridement ◆ Human Tetanus IG IM; optimum dose for IM human TIG yet to be established ◆ Traditional recommendations: single dose of 3000-6000 units ◆ Limited data suggests doses as low as 500 units as effective <p>Penicillin - GABA antagonist are associated with seizures Metronidazole recommended as choice</p> <p>Check maternal immunisation</p> <p><i>Reference: 9, 10</i></p> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|--|---|
| | Preferred | Alternative | |
| Gonococcal Ophthalmitis | <p>Immediate and frequent saline eye irrigation</p> <p>Non-disseminated disease: 3rd gen. <i>Cephalosporins</i>, e.g. Ceftriaxone 25-50mg/kg IV (max 125mg) once</p> <p>Disseminated disease: 3rd gen. <i>Cephalosporins</i>, e.g; Ceftriaxone 50mg/kg IV q24h 1st week of life, then q12h for 7 days (Cefotaxime for neonates with hyperbilirubinemia)</p> | | <p>Prophylaxis for infants born to mothers with gonococcal infections: topical Silver Nitrate 1%</p> <ul style="list-style-type: none"> ◆ Screen mother and baby for Chlamydial Infection ◆ Screen for other STDs ◆ Investigate and treat parents <p><i>Reference: 11,12</i></p> |
| <p>Conjunctivitis <i>Chlamydia trachomatis</i></p> | <p>EES 50mg/kg/24h PO in 4 divided doses for 14 days</p> <p>(Topical therapy not necessary if systemic treatment given)</p> | <p>Azithromycin 20mg/kg PO q24h for 3 days</p> | <p>Diagnosis by tissue culture, antigen detection (IFA, EIA) or NAAT Eye swab from conjunctiva of everted eyelid with Dacron tipped swab or swab from test kit Test also for gonococcus Treat mother & sexual partner</p> <p>Efficacy of treatment 80%, follow-up necessary. Second course of therapy may be required</p> <p><i>Reference: 17, 18</i></p> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Early onset sepsis (<48 hrs) Sepsis/pneumonia/meningitis Group B Strep (GB) Gram -ve bacteria (GNB) | Benzylpenicillin IV OR Ampicillin IV PLUS Gentamicin ¹ IV (Till C&S results) <u>Duration:</u> Sepsis: 7-10 days G+ve meningitis: 2 weeks G-ve meningitis: 3 weeks | Ampicillin PLUS 3 rd gen. Cephalosporins, e.g. Cefotaxime (Refer Drug Dosages - Frank Shann) | Suspect in maternal chorioamnionitis, sepsis, PROM (>18 hours) Do full septic workup, CXR No evidence from randomised trials to suggest that any antibiotic regimen may be better than any other in the treatment of presumed early neonatal sepsis <i>Reference: 13</i> |
| Group B Strep (GBS) Infection <i>Streptococcus agalactiae</i> | Benzylpenicillin IV OR Ampicillin IV PLUS Gentamicin ¹ IV <u>Duration</u> Sepsis: 10 days Meningitis: 14 days Osteomyelitis: 4 weeks | | <i>Reference: 14</i> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|--|--|---|
| | Preferred | Alternative | |
| Postnatal Infections | | | |
| Community Acquired Infections (Late onset sepsis >48 hrs) Pneumonia, Sepsis Group B Strep <i>E coli</i> <i>Klebsiella</i> <i>Enterobacter</i> , <i>S aureus</i> Possible <i>Listeria</i> | Ampicillin OR Penicillin PLUS Gentamicin ¹ (Refer Drug Dosages - Frank Shann) | Penicillin PLUS 3 rd gen. <i>Cephalosporins</i> , e.g. Cefotaxime (Refer Drug Dosages - Frank Shann) | Inadequate evidence from randomised trials in favour of any particular antibiotic regimen for the treatment of suspected late onset neonatal sepsis Discontinue antibiotics after 72 hours if culture negative or course does not support diagnosis Reference: 15 |
| Hospital Acquired Infection (Pneumonia, sepsis, meningitis) Based on predominant flora and susceptibility <i>Coagulase-negative staphylococci</i> , <i>Staphylococcus aureus</i> , <i>E coli</i> , <i>Klebsiella</i> , <i>Pseudomonas</i> , <i>Enterobacter</i> , <i>Candida</i> , <i>GBS</i> , <i>Serratia</i> , <i>Acinetobacter</i> | Cloxacillin IV PLUS Gentamicin ¹ /Amikacin ¹ IV (Use Cloxacillin if <i>S.aureus</i> is a problem in the respective nursery Otherwise replace Cloxacillin with any other antibiotic appropriate for the predominant flora) | 3 rd gen. <i>Cephalosporins</i> , e.g. Cefotaxime IV PLUS Gentamicin ¹ OR Vancomycin ¹ IV if MRSA strongly suspected | Antibiotics used should be according to the microorganisms prevalent in NICU |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| <i>Necrotising Enterocolitis</i> <i>Klebsiella, E. Coli, Clostridia,</i> <i>Coagulase-negative Staphylococcus</i> (CoNS), <i>Enterococci, Bacteroides</i> | Ampicillin IV PLUS Gentamicin ¹ IV PLUS Metronidazole IV For 10-14 days (<i>Vancomycin</i> ¹ if CoNS suspected) | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate PLUS Gentamicin ¹ | There is insufficient evidence on benefit or risk regarding choice of antibiotic regimens or duration of antibiotic treatment of NEC Note: Decisions regarding antibiotic choice and duration might best be guided by culture results & antibiotic resistance patterns present within nurseries <i>Reference: 15</i> |

¹Refer to Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

²Refer to Appendix 3 (Antibiotic Dosages For Neonates)

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OCULAR INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|---|--|
| | Preferred | Alternative | |
| Preseptal cellulitis <i>Strep pneumoniae, Staph aureus, Streptococcus sp</i> | Cloxacillin 50mg/kg PO q6h for 5 days | 3 months and older and under 40kg, Amoxicillin 25-45mg/kg/24h PO in 3 divided doses | Consider corresponding intravenous antibiotics: ♦ in severe infections ♦ if secondary to sinusitis |
| Orbital cellulitis/abscess <i>H. influenzae</i> | <i>3rd gen. Cephalosporins, e.g. Ceftriaxone</i> 20-80mg/kg IV q24h for 7 to 14 days | <u>Less than 20kg:</u> Cloxacillin 25-50mg/kg/24h IV in 4 divided doses <u>Over 20kg:</u> Cloxacillin 250-500mg IV q6h OR <u>0 to 1 week of age</u> <i>3rd gen. Cephalosporins, e.g. Cefotaxime</i> 50mg/kg IV q12h <u>1 to 4 weeks of age</u> <i>3rd gen. Cephalosporins, e.g. Cefotaxime</i> 50 mg/kg IV q8h <u>1 month to 12 years AND under 50kg</u> <i>3rd gen. Cephalosporins, e.g. Cefotaxime</i> 50-180mg/kg/24h IV/IM in 2-4 divided doses | Treat underlying cause (e.g. sinusitis) In orbital abscess, surgical drainage is often necessary References: 1. Medical and Surgical Management of Orbital Cellulitis Michael T. Yen, M.D. Contemporary Ophthalmology, June 2005, Vol 4, No.11, Page 1-6 2. Role of Inflammation in Orbital Cellulitis Carolyn E. Kloek, MD Peter A.D. Rubin, MD Manuscript on Role of Inflammation in Orbital Cellulitis Page 57-68 |

NATIONAL ANTI-BIOTIC GUIDELINE 2008

RESPIRATORY TRACT INFECTIONS

A. UPPER RESPIRATORY TRACT INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|---|--|---|
| | Preferred | Alternative | |
| Tonsillitis/Pharyngitis | Phenoxymethylpenicillin 10mg/kg PO q6h for 10 days | If allergic to penicillin, EES 20mg/kg PO q12h for 10 days (max 1gm/day) | Antibiotic required if: <ul style="list-style-type: none"> ◆ Streptococcus suspected ◆ fever >38°C ◆ tender cervical lymphadenopathy ◆ tonsillar swelling exudates ◆ NO cough Reference: 1, 11 |
| Rhinosinusitis | Mainly viral, therefore antibiotic not recommended | | Reference: 1, 5, 11 |
| Otitis media Sinusitis | Amoxicillin 80-90mg/kg/24h PO in 3 divided doses for 5-7 days | If resistance suspected to Amoxicillin, β -lactam/ β -lactamase inhibitors, e.g. Amoxicillin (90mg/kg/24h)/ Clavulanate PO in 2 divided doses for 5-7 days | Reference: 6 |

B. LOWER RESPIRATORY TRACT INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|---|
| | Preferred | Alternative | |
| 1. Community Acquired Pneumonia (Outpatient) | | | |
| Less than 5 years <i>Empirical therapy</i> | Amoxicillin 30-75mg/kg/24h PO in 3 divided doses for 5-7 days | <i>β-lactam/β-lactamase inhibitors, e.g. Amoxicillin (30-75mg/kg/24h)/ Clavulanate PO in 2 divided doses for 5-7 days</i> OR EES 20mg/kg PO q12h | <i>Reference: 2, 3, 5, 7, 8</i> |
| Age more than 5 years | EES 20mg/kg PO q12h for 7 days OR Azithromycin 15mg/kg (day 1) PO q24h then 7.5 mg/kg (day 2-5) PO q24h | Amoxicillin 30-75mg/kg/24h PO in 3 divided doses for 5-7 days | |
| 2. Community Acquired Pneumonia (Inpatient) | | | |
| Pneumonia inpatient | Benzylopicillin 30-60mg/kg IV q6h for 7 days | Benzylopicillin 30-60mg/kg IV q6h PLUS Gentamicin ¹ 5mg/kg IV q24h for 7 days | Cloxacillin if Staphylococcus aureus <i>Reference: 3</i> |

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|--|
| | Preferred | Alternative | |
| 3. Severe Community Acquired Pneumonia | | | |
| Severe community acquired | 3 rd gen. Cephalosporins, e.g. Cefotaxime 50mg/kg q4-6h OR Ceftriaxone 50mg/kg q12h OR Cefuroxime 50mg/kg IV q8h PLUS Erythromycin 15-25mg/kg IV q6h for 7 days | Benzylpenicillin 30-60mg/kg IV q6h PLUS Gentamicin ¹ 5mg/kg IV q24h PLUS Erythromycin 15-25mg/kg IV q6h for 7 days | Cloxacillin if Staphylococcus <i>Reference: 8, 10</i> |

¹Refer Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

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SKIN AND SOFT TISSUE INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|---|
| | Preferred | Alternative | |
| Abscess <i>Staphylococcus aureus</i> | Cloxacillin 50-100mg/kg/24h PO/IV in 4 divided doses for 7-10 days | | Incision & drainage if indicated. Pus for culture. Parenteral mode for severe infections |
| Animal bites <i>Pasteurella multocida, Staphy. Spp, Streptococcus spp</i> | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin (30-75mg/kg/24h)/ Clavulanate PO in 2 divided doses for 7 days | Amoxicillin (30-75mg/kg/24h)/ Clavulanate PO in 2 divided doses | Consider rabies prophylaxis according to local epidemiology |
| Cellulitis <i>Staphylococcus aureus</i> <i>Streptococcus pyogenes</i> | Cloxacillin 50-100mg/kg/24h PO/IV in 4 divided doses for 7-10 days | | Parenteral mode for extensive lesions |
| Impetigo <i>Staphylococcus aureus,</i> <i>Streptococcus pyogenes</i> | Cloxacillin 50mg/kg/24h PO in 4 divided doses for 7 days | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin (30-75mg/kg/24h)/ Clavulanate PO in 2 divided doses for 7 days OR Cephalexin 50-75mg/kg/24h PO in 3 divided doses for 7 days | Localised lesions: Use Mupirocin topical q8h |
| Necrotizing fasciitis | Benzylpenicillin 50,000 units/kg IV q4h PLUS Gentamicin ¹ 5 mg/kg IV q24h | | Aggressive surgical debridement; consider combination of Penicillin and Clindamycin and IVIG to bind toxin for streptococcal infection with toxic shock |

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| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|----------|
| | Preferred | Alternative | |
| Polymicrobial: Gram +ve cocci, Anerobic Gram-ve rods, Anerobes | PLUS Metronidazole 10mg/kg IV q8h for 10 days | | |
| Scalded skin syndrome <i>Staphylococcus aureus</i> | Cloxacillin 150mg/kg/24h IV in 4 divided doses <u>then</u> , step down to 50mg/kg/24h PO in 4 divided doses for 7 days OR Cephalexin 50-75mg/kg/24h PO in 3 divided doses for 7 days | | |
| Scabies <i>Sarcoptes scabiei</i> | For children > 2 years and <12: Benzyl Benzoate emulsion (EBB) 12.5% apply from neck down and leave for 24 hours for 2 days | Gamma Benzene Hexachloride 0.5% (Lindane) apply and leave for 8 hours (not to be repeated in less than a week) Babies: Sulphur 6% in calamine lotion q12h OR Crotamiton (Eurax) cream q12h for 2-3 weeks OR Permethrin 5% cream apply and leave for 8 hours (not for babies less than 2 months) | |

¹Refer to Appendix 1 (Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

SURGICAL INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|-----------------------------------|
| | Preferred | Alternative | |
| A. General Surgery | | | |
| Empyema thoracis <i>Staph aureus</i> | Cloxacillin 25-50mg/kg/24h IV in 4 divided doses | Based on C&S | |
| Enterocolitis <i>Enterobacteriaceae</i> <i>enterococci, Bacteroides</i> | Metronidazole 500mg IV q8h PLUS <i>2nd or 3rd gen Cephalosporins e.g.</i> Cefuroxime 750mg IV q6-8h or 1.5g IV q6-8h for severe infection OR Cefoperazone 100-150mg/kg/24h IV in 2-3 divided doses | | |
| B. Bone & Joints Infections | | | |
| Septic Arthritis <i>Staph. Aureus</i> <i>Haemophilus Influenza</i> | Cloxacillin 200mg/kg/24h IV in 4 divided doses for 14 days followed by oral for 14 days, longer if necessary | <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin/Clavulanate IV for 14 days followed by oral for 14 days, longer if necessary Depends on C&S | Surgical debridement if necessary |

TROPICAL INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|--|---|---|
| | Preferred | Alternative | |
| MALARIA | | | |
| Uncomplicated malaria (Symptomatic infection with malaria parasitaemia without signs of severity or evidence of vital organ dysfunction) Plasmodium falciparum | **Artesunate/Mefloquine (Artequine®) (Refer Notes 1*) D1-3: Artesunate 4mg/kg PO q24h D1-3: Mefloquine 25mg/kg PO over 2 days OR 8.3mg/kg PO q24h <u>Dosage according to body wt</u> <10kg : Artesunate 25mg q24h for 3 days Mefloquine 125mg single dose 10-20kg: Artesunate 50mg q24h for 3 days Mefloquine 125mg q24h for 3 days 20-40kg: Artesunate: 100mg q24h for 3 days Mefloquine 250mg q24h for 3 days (Artequine® 300/750) OR Artemether/Lumefantrine(Riamet®) (Refer Notes 2*) | Quinine D1-7: Quinine 10mg salt/kg PO q8h PLUS Doxycycline 3.5mg/kg PO q24h OR Clindamycin 10mg/kg PO q12h Either drug to be given for 7 days Doxycycline for children >8 years Clindamycin for children <8 years | Check G6PD before giving primaquine Add Primaquine 0.75mg/kg single dose q24h if gametocyte is present at any time during treatment ** Not available in Ministry of Health National Formulary (Artesunate/ Mefloquine available in 3 formulations: Artequine Paediatric in pellets form for small children < 20kg, Artequine 300/750 for those between 20-40kg & Artequine 600/1500 for > 40kg) <u>Notes 1*:</u> - Do not use AS/MQ in pregnancy - AS/MQ may cause seizure in children with epilepsy - AS/MQ interact with Quinine, Chloroquine and Halofantrine and may cause arrhythmia |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|--|---|---|
| | Preferred | Alternative | |
| | <p><u>Dosage according to body wt</u> 5-14kg: D1: 1 tablet stat then 1 tablet again after 8 hours D2-3: 1 tablet q12h 15-24kg: D1: 2 tablets stat then 2 tablets again after 8 hours D2-3: 2 tablets q12h 25-35kg: D1: 3 tablets stat then 3 tablet again after 8 hours D2-3: 3 tablets q12h</p> | | <p>GIT symptoms such as abdominal pain, nausea, vomiting and diarrhoea are the most common side effects. Other symptoms include headache, dizziness and insomnia, convulsions and other symptoms</p> <p><u>Notes 2*:</u> Artemether/Lumefantrine is available as co-formulated tablets containing 20mg of artemether and 120 mg of lumefantrine. Lumefantrine absorption is enhanced by co-administration with fat containing food or milk</p> |
| <p>Complicated malaria</p> <ul style="list-style-type: none"> ♦ almost always due to P. falciparum ♦ always suspect mixed infections if vivax / malariae malaria appear more severe than usual <p>a) <i>Plasmodium falciparum</i></p> | <p>D1: **Artesunate 2.4mg/kg IV on admission, then repeat again at 12h</p> | <p>D1: Quinine loading 7mg/kg IV over 1 hour followed by infusion Quinine 10mg/kg over 4 hours then 10mg/kg q8h</p> | <p>Dilute Quinine in 250ml of D5% over 4 hours. Change to oral if able to tolerate. Quinine: Maximum 600mg.</p> |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---------------------------------------|--|---|---|
| | Preferred | Alternative | |
| | D2-7: **Artesunate 1.2mg/kg IV q24h | OR Loading 20mg/kg IV over 4 hours then IV 10mg/kg IV q8h D2-7: Quinine 10mg/kg IV q8h PLUS Doxycycline 3.5mg/kg PO q24h OR Clindamycin 10mg/kg/dose q12h Both drugs to be given for 7 days | ** Not available in Ministry of Health National Formulary |
| b) <i>Plasmodium vivax</i> | Total Chloroquine 25mg base/kg divided over 3 days as below: D1: 10mg base/kg stat then 5mg base/kg 6 hours later D2: 5mg base/kg q24h D3: 5mg base/kg q24h PLUS Primaquine 0.25mg base/kg PO q24h for 14 days | Repeat Chloroquine and Primaquine | Check G6PD status before giving Primaquine Primaquine 0.75mg base/kg once a week for 8 weeks |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|--------------------|
| | Preferred | Alternative | |
| c) <i>Plasmodium knowlesi/malariae</i> | Total Chloroquine 25mg base/kg divided over 3 days, as below: D1: 10mg base/kg PO stat then 5mg base/kg 6 hours later D2: 5mg base/kg PO q24 D3: 5mg base/kg PO q24h | Treat as complicated <i>Plasmodium falciparum</i> | |
| Mixed infection | Treat as <i>Plasmodium falciparum</i> | | |
| LEPTOSPIROSIS | | | |
| Leptospirosis <i>L. ictero-haemorrhagiae</i> , <i>L. canicola</i> | Benzympenicillin 50,000 units/kg IV q6h for 7 days | 3 rd gen. Cephalosporins, e.g. Ceftriaxone 60-80mg/kg IV q24h OR Cefotaxime 150-200mg/kg/24h IV in 4 divided doses for 7 days | Reference: 2, 3, 4 |

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|---|---|--|---|
| | Preferred | Alternative | |
| MELIOIDOSIS | | | |
| Melioidosis <i>Burkholderia Pseudomallei</i> | Initial therapy: 3 rd gen. Cephalosporins, e.g. Ceftazidime 150mg/kg/24h IV in 3 divided doses for 10-14 days Maintenance: <i>β-lactam/β-lactamase inhibitors, e.g.</i> Amoxicillin (60/mg/kg/24h)/ Clavulanate PO in 3 divided doses for total treatment duration of 20 weeks | Initial therapy: Imipenem 75-100mg/kg/24h IV in 3-4 divided doses | Parenteral treatment should be used for at least 10 days or until clear improvement is noted <i>Reference: 5, 6</i> |
| SCRUB TYPHUS | | | |
| Scrub typhus <i>Rickettsia tsutsugamushi</i> | Chloramphenicol 50-75mg/kg/24h PO in 4 divided doses for 5-7 days | For children > 8 years, Doxycycline 2-4mg/kg/24h in 1-2 divided doses for 5-7 days | Avoid using Tetracycline or Doxycycline for young children as they can cause staining of the teeth <i>Reference: 7</i> |

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TUBERCULOSIS CHEMOTHERAPY IN CHILDREN

Treatment of TB disease

- Treatments have 2 phases, an initial intensive phase and a second continuation phase.
- Directly observed therapy is recommended for treatment of active disease
- In either phase, treatment can be given daily or three times weekly. Table 1 shows the first line (or essential) anti-TB drugs and their recommended doses

Table 1: Recommended doses of first-line anti-TB drugs for children

| Drug | Dose | | Intermittent Dose (thrice weekly) | |
|------------------|---------------------------|-------------------------|--------------------------------------|-----------------|
| | Daily Dose (mg/kg/day) | Maximum Dose (mg) | mg/kg/day | Maximum (mg) |
| Isoniazid (H) | 5 (4-6) | 300 | 10 (8-12) | |
| Rifampicin (R) | 10 (8-12) | 600 | 10 (8-12) | 600 |
| Pyrazinamide (Z) | 25 (20-30) | | 35 (30-40) | - |
| Ethambutol (E) | 20 (15-25) _b | | 30 (25-35) | |
| Streptomycin (S) | 15 (12-18) | | 15 (12-18) | |

a. Source: *Treatment of tuberculosis: guidelines for national programmes*

- b. The recommended daily dose of Ethambutol is higher in children (20 mg/kg) than in adults (15 mg/kg), because the pharmacokinetics are different (peak serum Ethambutol concentration is lower in children than in adults receiving the same mg/kg dose). Although ethambutol was frequently omitted from treatment regimens for children in the past, due in part to concern about the difficulty of monitoring for toxicity (particularly for optic neuritis) in young children, a literature review indicates that it is safe in children at a dose of 20 mg/kg (range 15-25 mg/kg) daily (3)
- c. *Streptomycin should be avoided when possible in children because the injection is painful and irreversible auditory nerve damage may occur. The use of Streptomycin in children is mainly reserved for the first 2 months of treatment of TB meningitis*

Table 1: Recommended treatment regimens for children in each TB diagnostic category

| TB Diagnostic category | TB cases | Regimena | |
|------------------------|--|---|----------------------------|
| | | Intensive phase - daily | Continuation phase - daily |
| III | <ul style="list-style-type: none"> ◆ New smear-negative pulmonary TB (other than in category I) ◆ Less severe forms of extrapulmonary TB | 2HRZ _b | 4HR or 6HE |
| I | <ul style="list-style-type: none"> ◆ New smear-positive pulmonary TB ◆ New smear-negative pulmonary TB with extensive parenchyma involvement ◆ Severe forms of extrapulmonary TB (other than TB meningitis see below) ◆ Severe concomitant HIV disease | 2HRZE | 4HR or 6HE _c |
| I | <ul style="list-style-type: none"> ◆ TB meningitis | 2RHZS _d | 4HR |
| II | <ul style="list-style-type: none"> ◆ Previously treated smear-positive pulmonary TB <ul style="list-style-type: none"> - relapse - treatment after interruption - treatment failure | 2HRZES/1HRZE | 5HRE |
| IV | <ul style="list-style-type: none"> ◆ Chronic and MDR-TB | Specially designed standardised or individualised regimens refer ID paediatrician | |

E, Ethambutol; H, Isoniazid; R, Rifampicin; S, Streptomycin; Z, Pyrazinamide

- a. Direct observation of drug administration is recommended during the initial phase of treatment and whenever the continuation phase contains Rifampicin
- b. In comparison with the treatment regimen for patients in diagnostic category I, Ethambutol may be omitted during the initial phase of treatment for patients with non-cavitary, smear-negative pulmonary TB who are known to be HIV-negative, patients known to be infected with fully drug-susceptible bacilli and young children with primary TB
- c. This regimen (2HRZE/6HE) may be associated with a higher rate of treatment failure and relapse compared with the 6-month regimen with Rifampicin in the continuation phase
- d. In comparison with the treatment regimen for patients in diagnostic category I, Streptomycin replaces Ethambutol in the treatment of TB meningitis

Corticosteroids

- ◆ May be used for the management of some complicated forms of TB, e.g. TB meningitis, complications of airway obstruction by TB lymph glands, and pericardial TB
- ◆ Recommended in all cases of TB meningitis

Prednisolone

- ♦ dosage of 2mg/kg daily
- ♦ increased up to 4mg/kg daily in more seriously ill children
- ♦ maximum dosage of 60mg/day for 4 weeks
- ♦ dose should then be gradually reduced over 1-2 weeks before stopping

Reference: Guidance for national tuberculosis programmes on the management of tuberculosis in children WHO/HTM/TB/2006.371 WHO/FCH/CAH/2006.7

URINARY TRACT INFECTIONS

| Condition/Infection & Likely Organism | Suggested Treatment | | Comments |
|--|---|--|--|
| | Preferred | Alternative | |
| Acute cystitis <i>E. Coli</i> <i>Proteus spp</i> | Trimethoprim 4mg/kg PO q12h (max 300mg daily) for 1 week | Trimethoprim(4mg/kg)/ Sulphamethoxazole PO q12h for 1 week | Cephalexin and Cefuroxime can also be used for UTI especially in children who had prior antibiotics Note: single dose of antibiotic therapy not recommended |
| Acute pyelonephritis Organisms: <i>E. Coli</i> <i>Proteus spp</i> | <i>3rd gen. Cephalosporins, e.g.</i> Cefotaxime 100mg/kg/24h IV in 3 divided doses for 10-14 days | Cefuroxime 100mg/kg/day IV q8h; OR Gentamicin ¹ 5mg/kg IV q24h | Culture should be repeated within 48hours. Antibiotic may need to be changed according to sensitivity Suggest to continue intravenous antibiotic until child is afebrile for 2-3 days and then switch to appropriate oral therapy after culture results e.g. Cefuroxime, for total of 10-14 days if susceptible |
| Prophylaxis for UTI | Trimethoprim 1-2mg/kg PO ON | Nitrofurantoin 1-2mg/kg PO ON | Antibiotic prophylaxis should not be routinely recommended in children with UTI Prophylactic antibiotics should be given for 3 days with MCUG (Micturating Cystourethrogram) taking place on the second day ¹ |

¹Refer Appendix 1(Clinical Pharmacokinetic Guidelines: Aminoglycosides & Vancomycin)

1 The Cochrane Database of Systematic Reviews

2. The Cochrane Library, Copyright 2006, The Cochrane Collaboration Volume (4), 2006

3. Stanley Hellerstein, MD. E-medicine, Urinary Tract infection Nov 2006

4. NICE Guidelines: Urinary tract infection: diagnosis, treatment and long term management of urinary tract infection in children 2007

VASCULAR INFECTIONS

| Infection/Condition & Likely Organism | Suggested Treatment | | Comments |
|---|--|-------------|---|
| | Preferred | Alternative | |
| IV line temporary/semi-permanent/tunnel type | | | |
| <i>S. epidermidis</i> <i>S. aureus</i> | Vancomycin ¹ 40mg/kg/24h IV in 3 divided doses (CoNS/MRSA) Cloxacillin 100mg/kg/24h IV in 4 divided doses (MSSA) | | ♦ <i>S. epid</i> : can try to save catheter 80% cure rate after 7-10 days of treatment ♦ <i>S. aureus</i> : remove catheter |
| <i>Candida sp*</i> <i>C. albicans</i> | Fluconazole 10mg/kg IV infusion stat, then 3-6mg/kg IV q24h | | *Immunocompromised - Amphotericin B efficacy limited - treat +ve blood cultures - remove catheter <i>Reference: 3</i> |
| <i>Non-C. albicans</i> | Amphotericin B 0.5-1mg/kg IV infusion over 4 hours q24h | | Fungal & Staph : Antibiotic therapy is usually given 2 weeks after catheter line removal <i>Reference: 1</i> |
| Septic thrombophlebitis | | | |
| <i>S. aureus</i> MSSA MRSA | Cloxacillin 100mg/kg/24h IV in 4 divided doses (MSSA) Vancomycin ¹ 40mg/kg/24h IV in 3 divided doses (MRSA) | | Gram-ve: Antibiotic therapy is given for additional 1 week after catheter removal |

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APPENDICES

Appendix 1

CLINICAL PHARMACOKINETIC GUIDELINES

AMINOGLYCOSIDES AND VANCOMYCIN

1. AMINOGLYCOSIDES

- A. Single Daily Dosing**
- B. Extended Interval Dosing**
- C. Conventional Dosing**

A. SINGLE DAILY DOSING (SDD)

Definition;

Is an approach of administering aminoglycosides for otherwise healthy individuals in a single daily dose by slow infusion (30 minutes).

The pharmacodynamic rationale for SDD is based on the following concepts¹:

- Aminoglycosides display concentration-dependent bactericidal action—that is, higher dose and serum concentrations result in more rapid bacterial killing.
- Aminoglycosides exhibit a long post-antibiotic effect, resulting in persistent bacterial suppression even when serum concentrations decline large, single daily doses result in prolonged periods with negligible serum concentrations, potentially reducing renal cortical and auditory accumulation of the drug.
- SDD has the potential of reducing costs associated with drug administration and monitoring; patient convenience and outpatient administration are also facilitated by SDD.
- Below the MIC and thereby allowing less frequent drug administration.

Exclusion criteria;

SDD administration of aminoglycosides is reasonable in most patients, with the following exceptions²:

- Diagnosed with enterococcal endocarditis, for which multiple(conventional) dosing regimens have been found superior in experimental animals
- Pregnant patients;
- Children;
- Patients with severe renal insufficiency; and
- Patients with neutropenia, unless the aminoglycoside is used in combination with a β -lactam antibiotic agent.

Conventional multiple daily dosing regimens should also be considered for the treatment of serious *P. aeruginosa* infections (other than those confined to the urinary tract) because publish studies have included relatively few of these cases.

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| TABLE 1: RECOMMENDATIONS FOR SINGLE DAILY DOSING OF AMINOGLYCOSIDES | | | |
|---|--------------------------|----------|-------------------|
| Estimated creatinine clearance (mL/min)* | Dose (mg/kg) | | Dose interval (h) |
| | Gentamicin or Tobramycin | Amikacin | |
| >80 | 5.0 | 15.0 | 24 |
| 60-79 | 5.0 | 12.0 | 24 |
| 50 | 3.5 | 7.5 | 24 |
| 40 | 2.5 | 4.0 | 24 |
| <30 | Use conventional dosing | | |

Monitoring:

- Suspected unstable renal function- Post 2 hours and Post 7 hours
- Suggested monitoring: assess 18-hours serum concentration after second dose.
- Suggested "trough" levels:
 - 0.6 to 2.0 µg/mL for Gentamicin or Tobramycin;
 - 2.5 to 5.0 µg/mL for Amikacin.

Data from Gilbert.³

B. EXTENDED INTERVAL DOSING

Definition:

Is an approach of giving standard dosing over 30 minutes at an extended interval (24 hourly, 36 hourly or more). The theoretical benefits of high-dose, extended-interval dosing are to⁴:

- Optimise concentration-dependent bacterial killing by achieving a high peak (>10x MIC).
- Minimize nephrotoxicity by administering larger, less frequent doses and potentially decreasing renal cortical aminoglycoside concentrations.
- Utilize the post-antibiotic effect (PAE), defined as a recovery period before organisms can resume growth after drug removal.
- Minimize the development of adaptive resistance by allowing a recovery period during the dosing interval.

Patient's criteria:

| Inclusion criteria ⁵ | Exclusion criteria |
|---|--|
| <ul style="list-style-type: none"> • Concurrently receiving nephrotoxic agents such as amphotericin, cyclosporin or vancomycin • Exposed to contrast media • Quadriplegics or amputees • In the intensive care unit • More than 60 years of age • Continue on the once a day dose for more or equal than 5 days whose drug random concentration should be determined once a week thereafter | <ul style="list-style-type: none"> • Elderly (>65 yrs) • Creatinine clearance less than 30ml/min • Dialysis • Pregnancy • Endocarditis • Cystic fibrosis • Ascites • >20% burns • History of hearing loss or vestibular dysfunction • Gram positive infections (when AMG is used for synergy) • Mycobacterial infection |

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Dose adjusted to Creatinine Clearance⁶

| Drug | Dose (mg/kg) | CrCl : >60ml/min | CrCl : 40-59ml/min | CrCl : 20-39 ml/min | CrCl : <20ml/min |
|------------|--------------|------------------|--------------------|---------------------|------------------|
| Amikacin | 15 | Q24 hours | Q36 hours | Q48 hours | NR |
| Gentamicin | 5-7 | Q24 hours | Q 36 hours | Q48 hours | NR |
| Netilmicin | 5-7 | Q24 hours | Q36 hours | Q48 hours | NR |
| Tobramycin | 5-7 | Q24 hours | Q 36 hours | Q48 hours | NR |

NR-Not recommended

Monitoring:

At the second dose.

1. Trough level (1 hour before the next dose): <1mg/L or less
 - If >1mg/L extension of dosing interval necessary
2. Post levels (7-14 hours post dose): varies with dose and renal function
 - Determining new dosing interval by plotting to normograms eg. Hartford Hospital monogram

C. CONVENTIONAL DOSING**Definition:**

Is an approach of administrating in slow bolus dosing (50mg/minute) of Aminoglycosides in 8 hourly dosing.

Inclusion Criteria:

- Patients (especially when immunosuppressed) are receiving for life threatening infections
- Patients expected to require prolonged therapy (whose drug concentrations should be determined within 48 hours of therapy initiation and monitored at least once a week)
- Patients not responding to treatment or have suspected aminoglycoside- related toxicity but continuation of therapy is desirable.

TABLE 2: RECOMMENDED* DOSAGES AND SERUM CONCENTRATIONS OF THE AMINOGLYCOSIDES: CONVENTIONAL MULTIPLE DAILY DOSING

| Drug | Route | Daily dosage* | | Serum concentration† (µg/mL) | |
|------------|----------|---------------|--------------------------|------------------------------|--------|
| | | Total (mg/kg) | Divided into doses given | Peak‡ | Trough |
| Gentamicin | IV or IM | 3-5§ | Every 8 h | 4-6 | 1-2 |
| Tobramycin | IV or IM | 3-5 | Every 8 h | 4-6 | 1-2 |
| Netilmicin | IV or IM | 3-5 | Every 8 h | 4-6 | 1-2 |
| Amikacin | IV or IM | 15 | Every 8 h | 20-30 | 5-10 |

*Recommendations

- based on normal renal function.
- Adjustments of dosage based on age and impaired renal function

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- ♦ †"Peaks" shown are expected levels.
 - Higher peak serum concentrations are desirable in the treatment of life-threatening disease (for example, endocarditis) or less susceptible organisms.
 - When aminoglycosides are used for synergistic therapy, lower serum levels are needed.
- ♦ ‡Serum specimen obtained
 - After third dose (after 24 hours)
 - Trough - 30 minutes after completion of 30-minute intravenous infusion - Post - 3 to 60 minutes after intramuscular administration.
- ♦ §For serious infections,
 - 5mg/kg should be administered. For example, endocarditis caused by *Pseudomonas aeruginosa* in a young patient who has illicitly used drugs intravenously),
- ♦ 8mg/kg per day of Gentamicin or Tobramycin has been
 - considerable toxicity affecting cranial nerve VIII has been reported with use of this high dosage.

TABLE 3. GUIDELINES FOR DESIRED SERUM CONCENTRATIONS OF AMINOGLYCOSIDES FOR MULTIPLE DAILY ADMINISTRATION^a

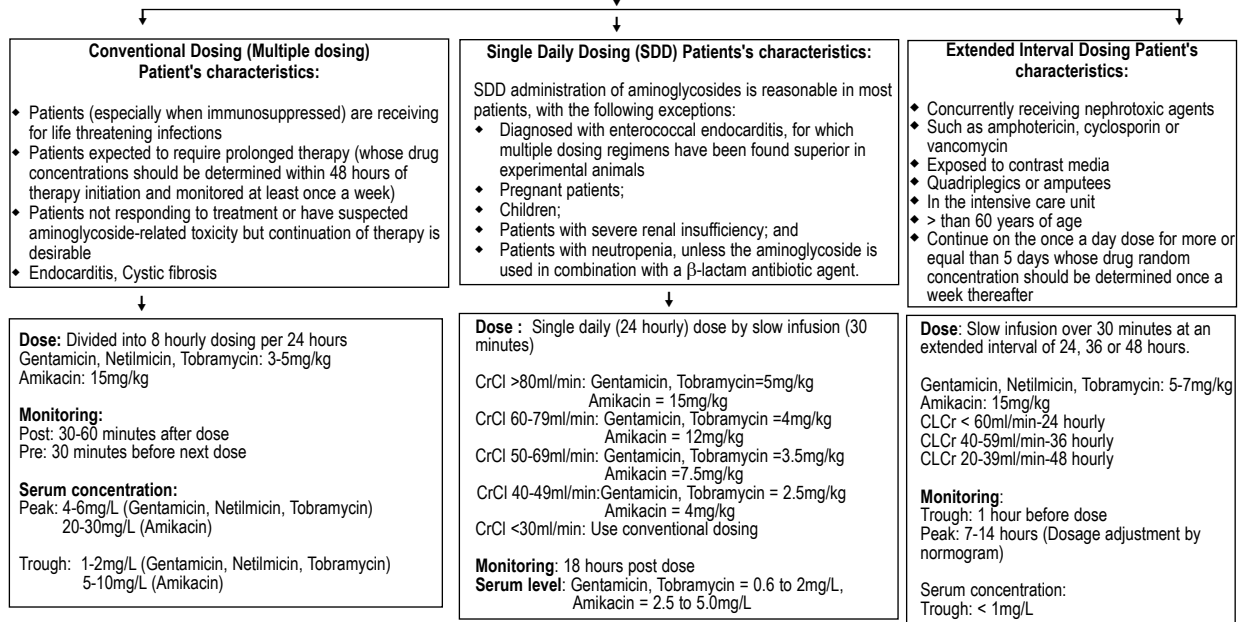
| Clinical situation | Serum concentration (mg/L) | |
|----------------------------|--|-----------|
| | Gentamicin, Tobramycin and Netilmicin ^b | Amikacin |
| Trough: | | |
| serious infection | 0.5-1.0 | 1.0-4.0 |
| life-threatening infection | 1.0-2.0 | 4.0-8.0 |
| Peak: | | |
| serious infection | 6.0-8.0 | 20.0-25.0 |
| life-threatening infection | 8.0-10.0 | 25.0-30.0 |

^aHigher peak and trough values have also been suggested.

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- (7) Ensom MHH, Davis GA, Cropp CD, Ensom RJ. Clinical Pharmacokinetics in the 21st century. *Clinical Pharmacokinetics* 1998; 24(4): 265-279
- (8) <http://Medscape.com>. Aminoglycosides still an important option for the treatment of infections in the elderly. *Drug Therapeutic Perspective* 1998. 11(8):8-1

AMINOGLYCOSIDES DOSING AND MONITORING GUIDELINES

AMIKACIN, GENTAMICIN, NETILMICIN, TOBRAMYCIN,



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2. VANCOMYCIN

- A. Therapeutic Drug Monitoring Guidelines For Aminoglycosides**
- B. Target Therapeutic Levels For Multiple Daily Dosing Aminoglycosides**

Vancomycin has been administered to treat Gram-positive infections since the 1950s, and because of the dramatic rise in drug resistance gram-positive infections caused by *Staphylococcus*, *Streptococcus*, and *Enterococcus* organisms, its use has increased².

It is indicated to treat *Methicillin-resistant Staphylococcus aureus*, confirmed by culture and sensitivity result, unless the clinical condition and past history reckon Vancomycin to be started as soon as possible.

Vancomycin activity is considered to be time-dependent - that is, antimicrobial activity depends on the duration that the drug level exceeds the minimum inhibitory concentration (MIC) of the target organism. Thus, peak levels have not been shown to correlate with efficacy or toxicity - indeed concentration monitoring is unnecessary in most cases.

Dosing of Vancomycin is based on 10-20 mg/kg/dose every 6 hours. Some literature recommended on 1g every 12 hours. Due to its pharmacodynamic properties, giving a small dose more frequently is more advantageous, provided that the renal function is normal.¹

Vancomycin exhibit most common administration-related side effects called 'Red-man syndrome'. This side effect happens in response to histamine release due to rapid infusion. Vancomycin should be administered over 1 to 2 hours' infusion to prevent this adverse effect from happening.

Other common side effects are:

1. Nephrotoxicity
2. Ototoxicity
3. Thrombophlebitis - related to site of administration

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A. Therapeutic Drug Monitoring Guidelines For Vancomycin

| DRUGS | TIME FOR 1 ST SAMPLING | IDEAL SAMPLING TIME | COMMENTS |
|------------|---|---|---|
| Vancomycin | AFTER 24 HOURS | POST LEVEL: 1 hour after infusion ends. TROUGH LEVEL: Within 30 minutes before the next dose. | Subsequent level: ONLY TROUGH LEVEL REQUIRED. |

B. Target Therapeutic Levels For Vancomycin

| DRUGS | THERAPEUTIC RANGE (mg/L) | | | |
|------------|--------------------------|----------------------|--------------------|----------------------|
| | PEAK | | TROUGH | |
| | Mild Infections | Severe Infections | Mild Infections | Severe Infections |
| Vancomycin | 20-40 | 20-40 | 10-15 | 15-20 |

References:

1. Leader WG, Chandler MHH, Castiglia M. Pharmacokinetic optimization of vancomycin therapy. Clin Pharmacokinetic. 1995; 28(4): 327-42. - Level III
2. Christine M.Karam, Peggy S.McKinnon, Melinda M.Neuhauser, Michael J. Rybak. Outcome assessment of minimizing Vancomycin monitoring and dosing adjustments. Pharmacotherapy. 1999. 19(3):257-266. - Level III

ANTIBIOTIC DOSAGES IN ADULTS PATIENTS WITH IMPAIRED RENAL FUNCTION

Unless stated, adjusted doses are % of dose for normal renal function

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| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|--|--------------------------------|---|--------------------------------|--------------------------------|--|--|
| | | > 50-90 | 10-50 | < 10 | | |
| ANTIBACTERIAL | | | | | | |
| Aminoglycoside: Traditional multiple daily doses - adjustment for renal disease | | | | | | |
| Amikacin | 7.5mg/kg q12h | 60-90% q12h or 100% q12-24h | 30-70% q12-18h or 100% q24-48h | 20-30% q24-48h or 100% q48-72h | HEMO: Extra 1/2 of normal renal function dose AD CAPD: 15-20mg lost/L dialysate/day | High flux hemodialysis membranes lead to unpredictable aminoglycoside clearance, measure post-dialysis drug levels for efficacy and toxicity. With CAPD, pharmacokinetics highly variable - check serum levels. Usual method for CAPD: 2 liters of dialysis fluid placed qid or 8 liters/day (give 8Lx20 mg lost/L = 160 mg of Amikacin supplement IV per day). Adjust dosing weight for obesity: [ideal body weight + 0.4(actual body weight - ideal body weight)]. Where possible dosage modifications should be based on monitoring of individual pharmacokinetic parameters. Please see TDM section. |
| Gentamicin, Tobramycin | 1.5mg/kg q8h | 60-90% q8 12h or 100% q12-24h | 30-70% q12h or 100% q24-48h | 20-30% q24-48h or 100% q48-72h | HEMO: Extra 1/2 of normal renal function dose AD CAPD: 3-4mg lost/L dialysate/day | |
| Netilmicin | 2mg/kg q8h | 50-90% q8 12h or 100% q12-24h | 20-60% q12h or 100% q24-48h | 10-20% q24-48h or 100% q48-72h | HEMO: Extra 1/2 of normal renal function dose AD CAPD: 3-4mg lost/L dialysate/day | |
| Streptomycin | 15mg/kg (max. of 1g) q24h | q24h | q24-72h | q72-96h | HEMO: Extra 1/2 of normal renal function dose AD CAPD: 20-40mg lost/L dialysate/day | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|--|--------------------------------|---|----------------|----------------|--|---|
| | | > 50-90 | 10-50 | < 10 | | |
| Carbapenem | | | | | | |
| Imipenem | 250-1000mg q6h | 100% | 50% | 25% | HEMO: Dose AD CAPD: Dose for CrCl <10 | ↑ potential for seizures if recommended doses exceeded in patients with CrCl<20 ml/min. Refer package insert for patients <70 kg |
| Meropenem | 500-1000mg q6h | 500mg q6h | 250-500mg q12h | 250-500mg q24h | HEMO: Dose AD CAPD: Dose for CrCl <10 | |
| Cephalosporin: DATA ON SELECTED PARENTERAL CEPHALOSPORINS | | | | | | |
| Cefazolin | 500-1500mg q6h | q8h | q12h | q24-48h | HEMO : 0.5-1.0G AD CAPD: 0.5G q12h | |
| Cefepime | 250-2000mg q8h | q12h | q16-24h | q24-48h | HEMO: 1g AD CAPD: dose for CrCl<10 | <u>Children with impaired renal function:</u> Age 2 months months to 12 years; 50mg/kg and age 1 month to 2 months; 30mg/kg equivalent to adult 2g. Same reduction in dose and/or increase in interval as of adult with renal impairment. (Product insert). |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|----------------------------|--------------------------------|---|---------|---------|---|--|
| | | > 50-90 | 10-50 | < 10 | | |
| Cefotaxime | 2g q8h | q8-12h | q12-24h | q24h | HEMO: Extra 1g AD CAPD: 0.5-1g qd | Active metabolite of cefotaxime in ESRD. ↓ dose further for hepatic & renal failure. |
| Cefoperazone/ Sulbactam | 2g q12h | 2g q12h | 2g q12h | 1g q12h | Only sulbactam component affected by hemodialysis. Dosing scheduled following dialysis period | |
| Ceftazidime | 2g q8h | q8-12h | q24-48h | 48h | HEMO: Extra 1g AD CAPD: 0.5g qd | Volume of distribution increases with infection. |
| Cefuroxime | 0.75-1.5g q8h | q8h | q8-12h | q24h | HEMO: Dose AD CAPD: Dose for CrCl <10 | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|------------------------|---------------------------------|---|------------------------------------|---------------------------------|--|--|
| | | > 50-90 | 10-50 | < 10 | | |
| Fluoroquinolone | | | | | | |
| Ciprofloxacin | 500-750mg PO (or 400mg IV) q12h | 100% | 50-75% | 50% | HEMO: 250mg PO or 200mg IV q12h CAPD:250mg PO or 200mg IV q8h | |
| Levofloxacin | 500mg q24h | 100% | 250mg q24-48h (500mg initial dose) | 250mg q48h (500mg initial dose) | HEMO & CAPD: Dose for CrCl <10 | |
| Ofloxacin | 400mg PO/IV q12h | 100% | 200-400mg q12h | 200mg q24h | HEMO: 100-200mg AD CAPD: Dose for CrCl <10 | |
| Macrolide | | | | | | |
| Clarithromycin | 0.5-1g q12h | 100% | 75% | 50-75% | HEMO: Dose AD CAPD: None | ESRD dosing recommendations based on extrapolation |
| Erythromycin | 250-500mg q6h | 100% | 100% | 50-75% | HEMO/CAPD/CAVH: None | Ototoxicity with high doses in ESRD. Vol. of distribution increases in ESRD. |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|-------------------------------------|-------------------------------------|---|---------------|---------------|--|---|
| | | > 50-90 | 10-50 | < 10 | | |
| Miscellaneous Antibacterials | | | | | | |
| Colistin | 80-160mg q8h | 160mg q12h | 160mg q24h | 160mg q36h | HEMO: 80mg AD | |
| Linezolid | 600mg PO/IV q12h | 600mg q12h | 600mg q12h | 600mg q12h AD | HEMO: As for CrCl <10 CAPD: No data | Accumulation of 2 metabolites - risk unknown |
| Metronidazole | 7.5mg/kg q6h | 100% | 100% | 50% | HEMO: Dose AD CAPD: Dose for CrCl <10 | HEMO clears metronidazole and its metabolites |
| Nitrofurantoin | 50-100mg | 100% | Avoid | Avoid | Not applicable | |
| Sulfamethoxazole | 1g q8h | q12h | q18h | q24h | HEMO: Extra 1g AD CAPD: 1g qd | |
| Trimethoprim | 100-200mg q12h | q12h | q18h | q24h | HEMO: Dose AD CAPD: q24h | New hemodialysis membranes ↑ clear. of Vancomycin; check levels. Individualised dosage based on plasma concentration is generally preferred. Other method : Loading dose 15mg/kg followed by dose equiv. to 15 times GFR daily. In anuric patients, 1g q 7-10 days. |
| Vancomycin | 1g q12h | 1g q12h | 1g q24-96h | 1g q4-7d | HEMO/CAPD: Dose for CrCl <10 | |
| Polymyxin B | 1-1.25mg/kg q12h (1mg=10,000 iu) | 0.5-1mg/kg q12h | 0.5mg/kg q12h | 0.2mg/kg q12h | | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|-----------------------------|--------------------------------|---|--------------------------------|--------------------------------|---|--|
| | | > 50-90 | 10-50 | < 10 | | |
| Penicillins | | | | | | |
| Amoxicillin, Ampicillin | 250-500mg q8h 250mg-2g q6h | q8h q6h | q8-12h q6-12h | q24h q12-24h | HEMO: Dose AD CAPD: 250mg q12h | |
| Amoxicillin/ Clavulanate | 500/125mg q8h | 500/125mg q8h | 250-500mg AM component q12h | 250-500mg AM component q24h | HEMO: As for CrCl <10; extra dose after dialysis | |
| Ampicillin/ Sulbactam | 2g AM + 1g SB q6h | q6h | q8-12h | q24h | HEMO: Dose AD CAPD: 2g AM / 1g SB q24h | |
| Benzylpenicillin | 0.5-4 million U q4h | 100% | 75% | 20-50% | HEMO: Dose AD CAPD: Dose for CrCl <10 | 1.7 mEq potassium/mU. ↑ potential for seizures. 6mU/d upper limit dose in ESRD. |
| Piperacillin | 4g q4-6h | q4-6h | q6-8h | q8-12h | HEMO: Dose AD CAPD: Dose for CrCl <10 | 1.9 mEq sodium/g |
| Pip(P) / Tazo(T) | 4.5g q6h | 4.5g q6h | 2.25g q6h | 2.25g q8h | HEMO: Dose for CrCl <10 + 0.75g AD CAPD: Dose for CrCl <10 | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|---|---|---|---|----------------------------------|--|--|
| | | > 50-90 | 10-50 | < 10 | | |
| Tetracycline | | | | | | |
| Tetracycline | 250-500mg q6h | q8-12h | q12-24h | q24h | HEMO/CAPD: None | Avoid in ESRD |
| ANTIFUNGAL | | | | | | |
| Amphotericin B & amphotericin B lipid complex | Non-lipid: 0.4-1.0 mg/kg/d ABCC: 3-6mg/kg/d ABLC: 5mg/kg/d LAB: 3-5mg/kg/d | q24h | q24h | q24-48h | HEMO: None CAPD: Dose for CrCl <10 | For Amphotericin B, toxicity lessened by saline loading; risk amplified by concomitant nephrotoxic drugs |
| Fluconazole | 200-400mg q24h | 200-400mg q24h | 100-200mg q24h | 100-200mg q24h | HEMO: 100% of recommended dose AD CAPD: Dose for CrCl <10 | |
| Itraconazole PO | 100-200mg q12h | 100% | 100% | 100% | HEMO/CAPD: No adjustment with oral solution | |
| Flucytosine | 200mg/kg q6h | <u>>50 ml/min</u> q6h | <u>10-50 ml/min</u> q12-24h | <u><10 ml/min</u> q 24-48h | HEMO/CAPD: Dose AD | |
| Voriconazole, IV | 6mg/kg IV q12h x 2, then 4mg/kg q12h | No change | If CrCl <50 ml/min, accumulation of IV vehicle (cyclodextrin). Switch to PO or suspension (no dose adjustment). | | | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|----------------------|------------------------------------|---|----------------|------------------|---|--|
| | | > 50-90 | 10-50 | < 10 | | |
| ANTIPARASITIC | | | | | | |
| Pentamidine | 4mg/kg/d | q24h | q24h | q24-36h | HEMO/CAPD: None | |
| ANTIPARASITIC | | | | | | |
| Ethambutol | 15-25mg/kg q24h | q24h | q24-36h | q48h | HEMO: Dose AD CAPD: Dose for CrCl <10 | 25mg/kg 4-6 hrs prior to dialysis for usual 3x/wk dialysis. Streptomycin recommended in lieu of Ethambutol in renal failure. |
| Isoniazid | 5mg/kg q24h (max. 300mg) | 100% | 100% | max. 200mg daily | HEMO: Dose AD CAPD: Dose for CrCl <10 | |
| Pyrazinamide | 25mg/kg q24h (max. dose 2.5g q24h) | 25mg/kg q24h | 25mg/kg q24h | 12-25mg/kg q24h | HEMO: 25-35mg/kg after each dialysis CAPD: No reduction; CAVH: No data | |
| Rifampin | 600mg q24h | 600mg q24h | 300-600mg q24h | 300-600mg q24h | HEMO: None CAPD: Dose for CrCl <10 | Biologically active metabolite. |
| Ethionamide | 500-750mg q12-24h | 100% | 100% | 50% | No dosage adjustments | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|-------------------------------------|---|---|---|---|---|--|
| | | > 50-90 | 10-50 | < 10 | | |
| ANTIVIRAL | | | | | | |
| Acyclovir, IV | 5-10mg/kg q8h | 5-10mg/kg q8h | 5-10mg/kg q12-24h | 2.5mg/kg q24h | HEMO: Dose AD CAPD: Dose for CrCl <10 | Rapid IV infusion can cause renal failure. |
| Adefovir | 10mg PO q24h | 10mg q24h | 10mg q48-72h | No dosing recommendation | HEMO: 10mg q7d AD | |
| Ganciclovir | Induction 5mg/kg q12h IV | 2.5-5mg/kg q12h | 1.25-2.5mg/kg q24h | 1.25mg/kg 3x/wk | HEMO: 1.25mg/kg AD CAPD: Dose for CrCl <10 | |
| | Maintenance 5mg/kg q24h IV | 2.5-5.0mg/kg q24h | 0.625-1.25mg/kg q24h | 0.625mg/kg 3x/wk | HEMO: 0.625mg/kg AD CAPD: Dose for CrCl <10 | |
| Indinavir / nelfinavir / nevirapine | No data on influence of renal insufficiency. Less than 20% excreted unchanged in urine. Probably no dose reduction. | | | | | |
| Lamivudine (HIV) | 150mg q12h | 100% | 50-150mg q24h (full first dose) | 25-50mg q24h (50mg first dose) | HEMO: Dose AD CAPD: Dose for CrCl <10 | |
| Lamivudine (HepB) | 100mg PO q24h | 30-49 ml/min 100mg 1st dose, then 50mg q24h | 15-29 ml/min 100mg 1st dose, then 25mg q24h | 5-14 ml/min 35mg 1st dose, then 15mg q24h | < 5 ml/min: 35mg 1st dose, then 10mg q24h. HEMO/CAPD: No dosage adjustment or additional dose. | |
| Ritonavir & Saquinavir, SGC | Negligible renal clearance. At present, no patient data. Avoid oral solution due to propylene glycol content. | | | | | |

| ANTIMICROBIAL | DOSE FOR NORMAL RENAL FUNCTION | ADJUSTMENT FOR RENAL FAILURE Estimated creatinine clearance (CrCl), ml/min | | | SUPPLEMENT FOR HAEMODIALYSIS, CAPD | COMMENTS |
|---|--------------------------------|---|----------------------------|--------------------------------|---|----------|
| | | > 50-90 | 10-50 | < 10 | | |
| Stavudine, PO | 40mg q12h | 100% | 50% q12-24h | ≥60kg: 20mg/d <60kg: 15mg/d | HEMO: Dose as for CrCl <10 AD CAPD: No data | |
| Zidovudine | 200mg q8h or 300mg q12h | 200mg q8h or 300mg q12h | 200mg q8h or 300mg q12h | 100mg q8h | HEMO: 100mg q8h AD CAPD: Dose for CrCl <10 | |
| AD = after dialysis. "Dose AD" refers only to timing of dose with NO extra drug | | | | | | |
| D = dosage reduction, I = interval extension; ABCC = Ampho B Cholesteryl Complex (e.g. Amphocil) ; ABLC = Ampho B Lipid Complex (e.g. Abelcet); LAB = Liposomal Ampho B (e.g. AmBisome); SGC=Soft gel capsule | | | | | | |

ANTIBIOTIC DOSAGES FOR NEONATES

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| Antibiotics | Routes | Dosages (mg/kg/dose) and Intervals of Administration | | | | |
|--|--------|---|------------------------|------------------|-----------------------|-------------------|
| | | Weight < 1200g | Weight 1200-2000g | | Weight > 2000g | |
| | | Age 0-4 weeks | Age 0-7 days | >7 days | Age 0-7 days | >7 days |
| Acyclovir | IV | 20 q8h or 500mg/m ² /dose q8h | | | | |
| Amikacin | IV, IM | 7.5 q18 - 24h | 7.5 q12h | 7.5-10 q8-12h | 7.5-10 q12h | 10 q8h |
| Amphotericin B | IV | Initial dose: 0.5-1 q24h infuse 2-6h. Increment dose: Increase as tolerated by 0.25-0.5 q24h-48h. Max. 1.5 /day. Test dose: 0.1 mg/kg/dose up to max 1mg, followed by remaining initial dose. | | | | |
| Ampicillin Meningitis <i>Group B strep</i> Other diseases | IV, IM | 50 q12h | 50 q12h 200/day q8h | 50 q8h 75 q6h | 50 q8h 200/day q8h | 50 q6h 75 q6h |
| | | 25 q12h | 25 q12h | 25 q8h | 25 q8h | 25 q6h |
| Cefazolin | IV, IM | 50 q12h | 20 q12h | 20 q12h | 20 q12h | 20 q8h |
| Cefotaxime | IV, IM | 50 q12h | 50 q12h | 50 q8h | 100-150/day q8-12h | 150-200/day q6-8h |
| Ceftazidime | IV, IM | | 50 q12h | 50 q8h | 100-150/day q8-12h | 50 q8h |
| Ceftriaxone | IV, IM | | 50 q24h | 50 q24h | | 50-75 q24h |
| Cefuroxime | IV, IM | 25-50 q12h | | | | |
| Chloramphenicol | IV, PO | | 25 q24h | 25 q24h | 25 q24h | 25 q12h |

| Antibiotics | Routes | Dosages (mg/kg/dose) and Intervals of Administration | | | | |
|---------------|------------|---|----------------|------------|----------------|-----------------|
| | | Weight < 1200g | Weight < 1200g | | Weight < 1200g | |
| | | Age 0-4 weeks | Age 0-7 days | >7 days | Age 0-7 days | >7 days |
| Clindamycin | IV, IM, PO | 5 q12h | 5 q12h | 5 q8h | 5 q8h | 20-30/day q6-8h |
| Cloxacillin | IV, IM, PO | 15 q6h. Severe infection: 25-50 q12h (1 st week life), q8h (2-4 week life), q4-6h (>4 weeks) | | | | |
| EES | PO | 10 q12h | 10 q12h | 10 q8h | 10 q12h | 10 q6-8h |
| Erythromycin | IV | Slow IV (max 5mg/kg/hr) 10 q6h. Severe infection: 15-25 q6h | | | | |
| Fluconazole | IV | Premature babies: ≤29 weeks gestation: 0-14 days, 5-6 q72h. >14 days, 5-6 q48h. 30-36 weeks: 3-6 q48h. Neonates >14 days: Oropharyngeal candidiasis, 6 /day then 3/day. Oesophageal candidiasis, 6/day then 3-12 /day. Systemic candidiasis, 6-12/day Cryptococcal meningitis (acute), 12/day then 6-12/day | | | | |
| Gentamicin | IV, IM | 2.5 q18-24h (<1000g: 3.5 q24h) | 2.5 q12h | 2.5 q8-12h | 2.5 q12h | 2.5 q8h |
| Imipenem | IV, IM | 20 q18-24h | 20 q12h | 20 q12h | 20-25 q12h | 25 q8h |
| Meropenem | IV | | 20 q12h | 20 q12h | 20 q12h | 20 q8h |
| Metronidazole | IV, PO | 7.5 q48h | 7.5 q24h | 7.5 q12h | 7.5 q12h | 15 q12h |
| Netilmicin * | IV, IM | | 3 q12h | 2.5-3 q8h | 3 q12h | 2.5-3 q8h |

| Antibiotics | Routes | Dosages (mg/kg/dose) and Intervals of Administration | | | | |
|--------------------------------|--------|--|------------------------|------------------------|-----------------------------|------------------------|
| | | Weight < 1200g | Weight < 1200g | | Weight < 1200g | |
| | | Age 0-4 weeks | Age 0-7 days | > 7 days | Age 0-7 days | > 7 days |
| Benzympenicillin Meningitis | IV | 50,000 u q12h | 50,000 u q12h | 50,000 u q8h | 50,000 u q8h | 50,000 u q6h |
| Group B strep | | | | | 25,000-450,000 u/day q8h | 450,000 u/day q8h |
| Other diseases | | 25,000 u q12h | 25,000 u q12h | 25,000 u q8h | 25,000 u q8h | 25,000 u q6h |
| Penicillin G Benzathine | IM | | 50,000 u (one dose) | 50,000 u (one dose) | 50,000 u (one dose) | 50,000 u (one dose) |
| Procaine # | | | 50,000 u q24h | 50,000 u q24h | 50,000 u q24h | 50,000 u q24h |
| Vancomycin | IV | 15 q24h | 10-15 q12-18h | 10-15 q8-12h | 10-15 q8-12h | 15-20 q8h |

Adapted from:

1. Lexi-Comp's Pediatric Dosage Handbook: Including Neonatal Dosing, Drug Administration, & Extemporaneous Preparations: Carol K. Taketomo, Donna M. Kraus, Jane H. Hodding, Jane Hurlburt Hodding 2006-2007
2. Drug Doses, 13ed. Frank Shann 2005-2008
3. Product info Netromycin™ Inj. 2006

Avoid using in this age group since sterile abscesses and procaine toxicity occur more frequently with neonates than older patients

ANTIBIOTICS IN PREGNANCY AND LACTATION

| Types of Antibiotics | Pregnancy Category (Book on Drugs in Pregnancy and Lactation) |
|----------------------------------|---|
| Griseofulvin | C |
| Terbinafine HCL | B (Manufacturer) |
| Clotrimazole | B |
| Tioconazole | NA |
| Doxycycline | D (Manufacturer) |
| Tetracycline | D |
| Minocycline | D |
| Chloramphenicol | C |
| Ampicillin | B |
| Amoxicillin | B (Manufacturer) |
| Bacampicillin | B (Manufacturer) |
| Piperacillin | B (Manufacturer) |
| Benzylpenicillin | B (Manufacturer) |
| Phenoxymethyl Penicillin | B (Manufacturer) |
| Procaine Benzylpenicillin | B (Manufacturer) |
| Benzathine Penicillin | B (Manufacturer) |
| Cloxacillin | B (Manufacturer) |
| Ampicillin / Sulbactam | NA |
| Amoxicillin / Clavulanate | B (Manufacturer) |
| Piperacillin / Tazobactam | Piperacillin-B (Manufacturer) |
| Cephalexin Monohydrate | B (Manufacturer) |
| Cefuroxime Axetil | B (Manufacturer) |
| Cefuroxime Sodium | B (Manufacturer) |
| Cefaclor | B (Manufacturer) |
| Cefotaxime | B (Manufacturer) |
| Ceftazidime | B (Manufacturer) |
| Ceftriaxone | B (Manufacturer) |
| Cefepime | B (Manufacturer) |
| Cefoperazone / Sulbactam | Cefoperazone-B (Manufacturer) |
| Cefoperazone | B (Manufacturer) |
| Meropenem | B (Manufacturer) |
| Imipenem / Cilastatin | C (Manufacturer) |
| Trimethoprim | C (Manufacturer) |
| Sulphamethoxazole / Trimethoprim | Sulphamethoxazole-C (Manufacturer) D (Author) |
| Erythromycin Lactobionate | B (Manufacturer) |
| Erythromycin Ethylsuccinate | B (Manufacturer) |
| Clarithromycin | C (Manufacturer) |
| Azithromycin | B (Manufacturer) |
| Clindamycin | B (Manufacturer) |
| Streptomycin | D (Manufacturer) |
| Gentamicin | C |
| Kanamycin | D |

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| Types of Antibiotics | Pregnancy Category (Book on Drugs in Pregnancy and Lactation) |
|------------------------------------|---|
| Amikacin | C-(Author) D-Manufacturer |
| Netilmicin | NA |
| Ofloxacin | C (Manufacturer) |
| Ciprofloxacin | C (Manufacturer) |
| Pefloxacin | NA |
| Vancomycin | B (Manufacturer) |
| Fusidic Acid | NA |
| Metronidazole | B (Manufacturer) |
| Tinidazole | NA |
| Nitrofurantoin | B (Manufacturer) |
| Linezolid | C (Manufacturer) |
| Amphotericin B | B (Manufacturer) |
| Miconazole | C (Manufacturer) |
| Ketoconazole | C (Manufacturer) |
| Fluconazole | C (Manufacturer) |
| Itraconazole | C (Manufacturer) |
| Flucytosine | C (Manufacturer) |
| Cycloserine | C (Manufacturer) |
| Rifampicin | C (Manufacturer) |
| Isoniazid | C |
| Pyrazinamide | C (Manufacturer) |
| Ethambutol | B |
| Rifampicin / Dapsone / Clofazimine | C (Manufacturer) |
| Clofazimine | C (Manufacturer) |
| Dapsone | C (Manufacturer) |
| Acyclovir | B (Manufacturer) |
| Ribavirin | X (Manufacturer) |
| Ganciclovir | C (Manufacturer) |
| Indinavir | C (Manufacturer) |
| Ritonavir | B (Manufacturer) |
| Lopinavir / Ritonavir | NA |
| Zidovudine | C (Manufacturer) |
| Didanosine | B (Manufacturer) |
| Stavudine | C (Manufacturer) |
| Zalcitabine | C (Manufacturer) |
| Lamivudine | C (Manufacturer) |
| Zidovudine / Lamivudine | Both-C (Manufacturer) |
| Nevirapine | C (Manufacturer) |
| Efavirenz | C (Manufacturer) |

NA-Not Available

B/C (Manufacturer)-Manufacturer rated its product in its professional literature

GUIDE TO COLLECTION AND TRANSPORT OF CLINICAL SPECIMEN

| SPECIMEN | COLLECTION CONTAINER | TRANSPORT |
|---------------------|---|------------------------------|
| Blood | Commercial blood culture bottle | - |
| CSF | Sterile bijou bottle | Immediately |
| Ear | Swab | Amies Transport Medium |
| Eye | Swab | Amies Transport Medium |
| | Corneal Scrapping | Bacteriologic Culture Plates |
| Faeces | Clean/Sterile Container | - |
| | Selenite F broth/Alkaline Peptone Water | - |
| Genital | Swab | Amies Transport Medium |
| Nose | Swab | Amies Transport Medium |
| Sinus | Swab | Amies Transport Medium |
| Sputum | Sterile Container | - |
| Peritoneal Fluid | Sterile Container | Within 30 minutes |
| Throat | Swab | Amies Transport Medium |
| Tissue | Sterile Container | - |
| Urine | Sterile Container | Within 30 minutes |
| Wound (superficial) | Swab | Amies Transport Medium |
| Wound (deep) | Swab PUS | Amies Transport Medium |

ANTIFUNGAL ACTIVITY SPECTRUM

| DRUG | ORGANISMS INHIBITED/CLINICAL SYNDROMES |
|--|--|
| POLYENES | |
| Amphotericin B - Conventional - Ampho B lipid complex(ABLC) - Ampho B cholesteryl Complex - Liposomal Ampho B | <i>Aspergillus spp.</i> <i>Candida albicans</i> <i>Candida glabrata</i> <i>Candida parapsilosis</i> <i>Candida tropicalis</i> <i>Candida krusei</i> <i>Candida spp.</i> <i>Blastomyces dermatitidis</i> <i>Coccidioides immitis</i> <i>Cryptococcus spp.</i> <i>Fusarium spp.</i> <i>Histoplasma capsulatum</i> <i>Phycomycetes</i> <i>Penicillium marnettei</i> <i>Paracoccidioides spp.</i> <i>Sporotrichosis</i> <i>Zygomycosis</i> *** <i>Candida lusitanae</i> & <i>Candida guilliermondii</i> are resistant to Amphotericin B |
| Nystatin | <i>Aspergillus spp.</i> <i>Candida spp.</i> <i>Blastomyces spp.</i> <i>Coccidioides spp.</i> <i>Cryptococcus spp.</i> <i>Histoplasma capsulatum</i> <i>Phycomycetes</i> <i>Paracoccidioides spp.</i> <i>Sporotrichosis</i> |
| PYRAMIDINE ANALOG | |
| 5-flucytosine | <i>Cryptococcus spp.</i> <i>Candida spp.</i> (including <i>Candida glabrata</i>) <i>Chromoblastomyces</i> |

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| DRUG | ORGANISMS INHIBITED/CLINICAL SYNDROMES |
|---------------------|--|
| AZOLES | |
| Ketoconazole | <i>Dermatophytes</i> <i>Candida spp.</i> <i>Histoplasma capsulatum</i> <i>Blastomyces dermatitidis</i> <i>Coccidioides immitis</i> <i>Cryptococcus spp</i> |
| Miconazole | <i>Dermatophytes</i> <i>Candida spp.</i> <i>Pseudallescheria boydii</i> <i>Coccidioides immitis</i> <i>Cryptococcus spp</i> |
| Fluconazole | <i>Candida spp.</i> <i>Candida albicans</i> <i>Candida glabrata</i> <i>Candida parapsilosis</i> <i>Candida tropicalis</i> <i>Candida guilliermondi</i> <i>Candida lusitanae</i> <i>Cryptococcus spp.</i> <i>Blastomyces dermatitidis</i> <i>Coccidioides immitis</i> <i>Sporotrichosis</i> *** <i>Candida krusei</i> resistant to fluconazole ***Fluconazole may require dose escalation when treating <i>Candida glabrata</i> |
| Itraconazole | <i>Histoplasma capsulatum</i> <i>Blastomyces dermatitidis</i> <i>Aspergillus spp.</i> <i>Candida spp.</i> <i>Candida albicans</i> <i>Candida tropicalis</i> <i>Candida guilliermondi</i> <i>Candida lusitanae</i> <i>Coccidioides immitis</i> <i>Sporotrichosis</i> <i>Pityriasis versicolor</i> <i>Penicillium marseillei</i> <i>Onychomycosis</i> <i>Chromoblastomycosis (Cladosporium or Fonsecaea)</i> <i>Coccidioides immitis</i> <i>Cryptococcus spp</i> *** <i>Candida krusei</i> & <i>Candida glabrata</i> are resistant to itraconazole |

NATIONAL ANTIBIOTIC GUIDELINE 2008

| DRUG | ORGANISMS INHIBITED/CLINICAL SYNDROMES |
|---------------------|---|
| NEWER AZOLES | |
| Voriconazole | <i>Aspergillus</i> spp. <i>Scedosporium</i> spp. <i>Fusarium</i> spp. <i>Candida krusei</i> <i>Candida</i> spp. <i>Candida albicans</i> <i>Candida glabrata</i> <i>Candida parapsilosis</i> <i>Candida tropicalis</i> <i>Candida krusei</i> <i>Candida guilliermondi</i> <i>Candida lusitaniae</i> |
| Posaconazole | Chromoblastomycosis (<i>Cladosporium</i> or <i>Fonsecaea</i>) <i>Coccidioides immitis</i> Zygomycosis |
| ECHINOCANDIN | |
| Caspofungin | <i>Candida</i> spp. <i>Candida albicans</i> <i>Candida glabrata</i> <i>Candida parapsilosis</i> <i>Candida tropicalis</i> <i>Candida krusei</i> <i>Candida guilliermondi</i> <i>Candida lusitaniae</i> <i>Aspergillus</i> spp. |
| Micafungin | <i>Candida</i> spp. <i>Candida albicans</i> <i>Candida glabrata</i> <i>Candida parapsilosis</i> <i>Candida tropicalis</i> <i>Candida krusei</i> <i>Candida guilliermondi</i> <i>Candida lusitaniae</i> <i>Aspergillus</i> spp. |

NATIONAL ANTIBIOTIC GUIDELINE 2008

| DRUG | ORGANISMS INHIBITED/CLINICAL SYNDROMES |
|------------------------|---|
| DERMATOPHYTOSIS | |
| Terbinafine | Tinea unguium - <i>T. rubrum</i> , <i>T. mentagrophytes</i> Tinea capitis - <i>T. tonsurans</i> , <i>T. mentagrophytes</i> , <i>T. violaceum</i> - <i>M. audouinii</i> , <i>M. gypsum</i> , <i>M. canis</i> Tinea corporis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>M. canis</i> Tinea cruris - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> Tinea pedis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> |
| Itraconazole | Tinea unguium - <i>T. rubrum</i> , <i>T. mentagrophytes</i> Tinea capitis - <i>T. tonsurans</i> , <i>T. mentagrophytes</i> , <i>T. violaceum</i> - <i>M. audouinii</i> , <i>M. gypsum</i> , <i>M. canis</i> Tinea versicolor - <i>P. ovale</i> , <i>M. furfur</i> |
| Fluconazole | Tinea unguium - <i>T. rubrum</i> , <i>T. mentagrophytes</i> Tinea capitis - <i>T. tonsurans</i> , <i>T. mentagrophytes</i> , <i>T. violaceum</i> - <i>M. audouinii</i> , <i>M. gypsum</i> , <i>M. canis</i> Tinea corporis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>M. canis</i> Tinea cruris - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> Tinea pedis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> Tinea versicolor - <i>P. ovale</i> , <i>M. furfur</i> |
| Griseofulvin | Tinea capitis - <i>T. tonsurans</i> , <i>T. mentagrophytes</i> , <i>T. violaceum</i> - <i>M. audouinii</i> , <i>M. gypsum</i> , <i>M. canis</i> Tinea corporis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>M. canis</i> Tinea cruris - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> |
| Ketoconazole | Tinea corporis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>M. canis</i> Tinea cruris - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> Tinea pedis - <i>T. rubrum</i> , <i>T. mentagrophytes</i> , <i>E. floccosum</i> Tinea versicolor - <i>P. ovale</i> , <i>M. furfur</i> |

Appendix 7 (i)

PERCENTAGE OF SPECIFIC RESISTANT OF SPECIFIC BACTERIA
(2002 - 2005)

| | MRSA | | | | VRSA | | | | PPNG | | | | Spectinomycin R NG | | | | Chloramphenicol R HI | | | | Ampicillin R HI | | | | Penicillin R Strep pneumo | | | | Chloramphenicol R S.typhi | | | | Tetracycline R V. cholera | | | | Penicillin R Strep Gp A | | | | Penicillin R Strep Gp B | | | | VRE | | | |
|--------|------|-------|------|------|------|------|------|------|------|------|------|------|--------------------|------|------|------|----------------------|------|------|------|-----------------|------|------|------|---------------------------|------|------|------|---------------------------|------|------|------|---------------------------|------|------|------|-------------------------|------|------|------|-------------------------|------|------|------|------|------|------|------|
| | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 | 2002 | 2003 | 2004 | 2005 |
| HRPZII | 24.2 | 11.6 | 4.9 | - | 0 | 0 | 0 | - | - | - | 40 | - | - | - | 12.5 | - | 19.4 | 16.8 | 21.3 | - | 25.8 | 16.7 | 26.6 | - | 20 | 3 | 2.2 | - | 0 | 0 | 0 | - | - | - | 0 | - | 11.3 | 25 | 1.3 | - | 32.3 | 0 | 1.5 | - | 0 | 0 | 0 | - |
| HPP | 45.7 | 42 | 37 | 32.4 | 0 | 0 | 0 | 0 | 56 | 40 | 37.5 | 40 | 0 | 0 | 0 | 0 | 9.1 | 0 | 0 | 0 | 0 | 16.7 | 37.5 | 20 | 14.8 | 26.8 | 17 | 23 | 0 | 0 | 0 | - | - | - | - | - | 0 | 0 | 0 | 2.5 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| HKL | 38.6 | 43.9 | 44.3 | 46.4 | 0 | 0 | 0 | 0 | 37.5 | 38.7 | 0 | 33.3 | - | 0 | 0 | - | 8.3 | 3.2 | 4 | 14.5 | 19.8 | 13.8 | 12 | 10.9 | 17.3 | 19 | 16 | 35.6 | 0 | 0 | 0 | 0 | 0 | 0 | - | - | 0 | 0 | 0 | 0 | 0 | 0 | 4.5 | 0 | 1.1 | 1 | 2.2 | |
| HTAR | 35.3 | - | 20.6 | - | 0 | - | 0 | - | 0 | - | 43 | - | - | - | 0 | - | 5.1 | - | 28 | - | 35.9 | - | 32 | - | 9.1 | - | 0 | - | 0 | - | 0 | - | 0 | - | 1.1 | - | 0 | - | 15 | - | 0 | - | 0 | - | 0 | - | | |
| HSAJB | 40.2 | 30.6 | 26.4 | 26.2 | 0 | 0 | 0 | 0 | - | 0 | 31.3 | 0 | - | 0 | 0 | 0 | 12.2 | 3 | 3.1 | 0 | 14 | 12 | 12.5 | 2.7 | 29.6 | 16 | 13.7 | 1.1 | 0 | 13 | 0 | 0 | - | - | - | 0 | 1 | 0.6 | 0 | 0 | 1 | 0.6 | 0.4 | 0.3 | 0 | 0 | 0 | 0 |
| HMEL | 22.5 | 18.6 | 15.7 | 16.6 | 0 | 1.9 | 1.6 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 12 | 5.2 | 1.7 | 0 | 7 | 3.5 | 0 | | | |
| HTAA | 27.5 | 24.5 | 25.5 | 22.5 | 0 | 0 | 0 | 0 | - | - | - | - | - | - | - | 9.4 | 5.2 | 8.6 | 0 | 23.3 | 9.1 | 5.4 | 13.3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| HQE | 26.8 | 21.2 | - | 19 | 0 | 0 | - | 0 | 33.3 | - | - | - | 0 | 1 | - | 0 | 0 | - | 9.1 | 0 | 0 | - | 0 | 1.4 | 0 | - | 6 | 0 | 0 | - | 0 | 0 | - | 0 | 0 | - | 0 | 0 | - | 0 | 0 | - | 0 | 9.1 | 4.7 | - | 0 | |
| HIPH | 34 | - | 24 | - | 0 | 0 | - | - | 25 | - | 33 | - | 0 | - | 0 | - | 5.4 | - | 0 | - | 21.4 | - | 0 | - | 10 | - | 17 | - | 0 | - | 0 | - | 0 | - | 0 | - | 0 | - | 0 | - | 4.7 | - | 1 | - | 0 | - | 0 | - |
| HTJ | 28.5 | 25.2 | 28.8 | 23.3 | 0 | 0 | 0 | 0 | 100 | - | - | 0 | 0 | 1 | - | 0 | 0 | 0 | 0 | 0 | 0 | 50 | - | 100 | 0 | 3 | 0 | 9 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| HSB | 23.6 | 24.2 | 23.6 | 19.9 | 0 | 0 | 0 | 0 | 30 | - | 66.7 | 0 | 0 | - | 0 | 0 | 3.1 | 0.5 | 0 | 10.5 | 6.3 | 6.4 | 5.3 | 0 | 0 | 0 | 0 | 9.3 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| HSEL | - | 34.7 | 26.6 | 18.5 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | - | - | - | 16.7 | 10 | 5.6 | 0 | 16.7 | 40 | 5.6 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| HSNZ | - | 12.12 | 10.1 | - | 0 | 0.18 | 0 | - | - | - | - | - | - | - | - | 5.36 | 4.8 | - | 22.2 | 9.1 | - | 0 | 3 | - | 0 | 3 | - | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| HTF | - | 13.6 | 10.3 | 12 | 7 | 2.7 | 0 | - | - | - | - | - | - | - | - | - | 0 | 0 | - | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| HUS | 6.9 | - | 10.3 | 16.3 | 0 | - | 0 | 0 | - | - | - | - | - | - | - | - | 0 | 0 | - | 100 | - | - | 0 | 0 | - | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |

HPP - Hospital Pulau Pinang
 HKL - Hospital Kuala Lumpur
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 HSAJB - Hospital Sultanah Aminah
 HMEL - Hospital Melaka
 HTAA- Hospital Tengku Ampuan Afzan
 HIPH - Hospital Ipoh
 HTJ - Hospital Tuanku Jaafar
 HSB - Hospital Sultanah Bahiyah
 HSEL - Hospital Selayang
 HSNZ - Hospital Sultanah Nur Zahirah
 HTF - Hospital Tuanku Fauziah
 * - Not verified
 ND -no data

Appendix 7 (ii)

PERCENTAGE OF ANTIBIOTIC RESISTANCE OF SPECIFIC BACTERIA
2006 - 2007

| Hospital | | Staph aureus | | N.gonorrhoeae | | N.gonorrhoeae | | H.influenzae | | H.influenzae | | S.pneumoniae | | S.Typhi | | V.cholerae | | GrpA Strep | | GrpB Strep | | Enterococci | | |
|----------|------------|--------------|------|---------------|------|-----------------|------|--------------|------|--------------|------|--------------|------|-------------------|------|----------------|------|--------------|------|--------------|------|--------------|------|------|
| | | (MRSA) | | (PPNG) | | Spectinomycin R | | Chloram R | | Ampicillin R | | Penicillin R | | Chloramphenicol R | | Tetracycline R | | Penicillin R | | Penicillin R | | Vancomycin R | | |
| | | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 | 2007 | 2006 |
| HPP | %R | 36 | 37.6 | ND | 53.3 | 0 | 0 | 9.1 | 3.1 | 0 | 35.3 | 19.3 | 30 | 0 | 45.5 | ND | 100 | 0 | 0 | 0 | 2.7 | 0 | 1.1 | |
| | No. tested | 1702 | 1749 | ND | 15 | 3 | 7 | 22 | 32 | 18 | 34 | 31 | 30 | 1 | 11 | ND | 2 | 41 | 70 | 494 | 406 | 219 | 185 | |
| HKL | %R | 46.8 | 44.1 | 100 | 0 | ND | 0 | 24.1 | 33.8 | 8.4 | 20 | 0 | 1.2 | 0 | 0 | ND | 0 | 0 | 0 | 0 | 0.1 | 1.6 | 0 | |
| | No. tested | 4377 | 4280 | 3 | 0 | ND | 0 | 166 | 65 | 166 | 65 | 89 | 81 | 5 | 2 | ND | 0 | 111 | 123 | 1222 | 800 | 757 | 33 | |
| HTAR | %R | 15.6 | 13.3 | 57.1 | 75 | 0 | 0 | 13.7 | 7.1 | 19.2 | 10.7 | 37 | 0 | 33.3 | 0 | 0 | ND | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | No. tested | 1038 | 916 | 7 | 8 | 2 | 1 | 51 | 28 | 52 | 28 | 54 | 42 | 6 | 4 | 2 | ND | 126 | 108 | 573 | 579 | 218 | 24 | |
| HSAJB | %R | 27 | 26.9 | 50 | 55.6 | 0 | 0 | 0 | 0 | 17 | 24.1 | 1.4 | 23.1 | 0 | 0 | 0 | 0 | 1 | 0.5 | 2 | 0.1 | 0 | 0 | |
| | No. tested | 3258 | 3072 | 6 | 9 | 5 | 0 | 47 | 23 | 47 | 54 | 70 | 65 | 14 | 4 | 0 | 0 | 209 | 202 | 679 | 831 | 209 | 29 | |
| HMEL | %R | 28.8 | 24.7 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 30 | 21.1 | 11.1 | 0 | 0 | 0 | 0 | 11.4 | 5.9 | 27.7 | 19.9 | 0 | 0 | |
| | No. tested | 1799 | 2380 | 0 | 2 | 0 | 0 | 2 | 11 | 2 | 10 | 6.9 | 45 | 3 | 5 | 1 | 3 | 44 | 101 | 242 | 682 | 46 | 49 | |
| HTAA | %R | 21.1 | 18.3 | 0 | 100 | ND | 0 | 3.2 | 1.4 | 12.9 | 23.3 | 7.7 | 11.1 | ND | 30.8 | 0 | 0 | 2.6 | 0 | 2.2 | 2.1 | 0 | 0 | |
| | No. tested | 1376 | 971 | 0 | 1 | ND | 0 | 31 | 69 | 31 | 60 | 26 | 9 | ND | 13 | 0 | 0 | 77 | 81 | 320 | 332 | 41 | 0 | |
| HIPH | %R | ND | 24.4 | ND | ND | ND | ND | ND | 5.9 | ND | 0 | ND | 36 | ND | 45.5 | ND | 0 | ND | 0.6 | ND | 0.6 | ND | 0.8* | |
| | No. tested | ND | 2058 | ND | ND | ND | ND | ND | 17 | ND | 17 | ND | 39 | ND | 11 | ND | 0 | ND | 170 | ND | 668 | ND | 379 | |
| HTJ | %R | ND | 12.9 | ND | 0 | ND | 0 | ND | 0 | ND | 0 | ND | 0 | ND | 4 | ND | 0 | ND | 0 | ND | 0 | ND | 0 | |
| | No. tested | ND | 854 | ND | 0 | ND | 0 | ND | 3 | ND | 3 | ND | 28 | ND | 6 | ND | 0 | ND | 65 | ND | 548 | ND | 222 | |
| HSB | %R | 26.7 | 21.9 | 38.5 | 35.7 | 0 | 0 | 2.7 | 0 | 5.8 | 17.8 | 11.7 | 0 | 0 | 0 | 0 | 0 | 6.8 | 0 | 0 | 0 | 0 | 0 | |
| | No. tested | 2472 | 1639 | 13 | 14 | 13 | 14 | 259 | 129 | 259 | 129 | 60 | 44 | 3 | 6 | 79 | 0 | 51 | 132 | 792 | 968 | 476 | 424 | |
| HSEL | %R | 28.4 | 28.6 | 0 | 0 | 0 | ND | 0 | 0 | 0 | 0 | 2.4 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1.9 | 0 |
| | No. tested | 1298 | 1201 | 0 | 1 | 0 | ND | 7 | 6 | 7 | 6 | 42 | 18 | 4 | 2 | 0 | 0 | 116 | 47 | 328 | 482 | 255 | 298 | |
| HSNZ | %R | 9.8 | 6.8 | 0 | 100 | 0 | 0 | 0 | 0 | 5.8 | 40 | 27.3 | 0 | 0 | 0 | 0 | 0 | 2.9 | 11 | 2.3 | 1 | 0 | 0 | |
| | No. tested | 764 | 687 | 0 | 1 | 1 | 1 | 17 | 6 | 17 | 5 | 33 | 27 | 0 | 1 | 0 | 0 | 68 | 55 | 622 | 687 | 23 | 0 | |
| HTF | %R | 13.3 | 8.7 | 0 | 100 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 100 | 0 | 0 | 4.2 | 0 | 7.7 | 0 | 0 | 25* | |
| | No. tested | 369 | 289 | 0 | 2 | 1 | 0 | 4 | 0 | 4 | 0 | 5 | 3 | 0 | 1 | 0 | 0 | 24 | 45 | 130 | 196 | 19 | 4 | |

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HIPH - Hospital Ipoh
HTJ - Hospital Tuanku Jaafar
HSB - Hospital Sultanah Bahiyah
HSEL - Hospital Selayang
HSNZ - Hospital Sultanah Nur Zahirah
HTF - Hospital Tuanku Fauziah

* - Not verified
ND -no data

Appendix 8 (ii)

PERCENTAGE OF ANTIBIOTIC RESISTANCE AMONG GRAM NEGATIVE BACTERIA
2006

| Organism | Ambicacin | Amoxicillin/Clavulanic acid | Ampicillin | Ampicillin/ Sulbactam | Cefepime | Cefepazone | Cefepazone/ Sulbactam | Cefotaxime | Ceftazidime | Ceftiozone | Carbapenem sodium | Cephalexin | Chloramphenicol | Ciprofloxacin | Gentamicin | Imipenem | Meropenem | Netilmicin | Nitrofurantoin | Piperacillin | Piperacillin/ Tacobactam | Tetracycline | Trimethoprim/ Sulfamethoxazole | |
|-------------------------------------|----------------|-----------------------------|-----------------|-----------------------|----------------|----------------|-----------------------|-----------------|-----------------|----------------|-------------------|----------------|-----------------|-----------------|-----------------|-----------------|----------------|----------------|----------------|----------------|--------------------------|----------------|--------------------------------|---------------|
| <i>A. baumannii</i> | 23.5 (3300) | 70.6 (728) | 92.9 (911) | 43.3 (3159) | 50.7 (3041) | 73.1 (3316) | 22.9 (2631) | 74.5 (924) | 41.8 (3352) | 83.1 (803) | 90.3 (972) | | | 41.9 (3177) | 41 (3371) | 44.5 (3289) | 47 (3441) | 15.1 (2543) | 84.3 (89) | 56.4 (1582) | 49 (439) | | 40.7 (852) | |
| <i>C. freundii</i> | 8.1 (147) | 74.8 (147) | 85.6 (169) | 64 (86) | 22.6 (93) | 39.3 (122) | 10.3 (29) | 41.4 (145) | 38.2 (170) | 43.7 (126) | 60.6 (137) | 73.8 (65) | | 37 (162) | 38.8 (170) | 1.2 (168) | 4.4 (165) | 22.8 (82) | 15.2 (161) | 47.6 (21) | 28.6 (21) | | 56.1 (164) | |
| <i>Enterobacter sp.</i> | 2.6 (1348) | 65 (1705) | 88.2 (1761) | 64.3 (972) | 6.3 (1805) | 18.6 (1288) | 0.5 (843) | 18.8 (1051) | 15.2 (1757) | 19 (1682) | 37.9 (1727) | 8.4 (758) | 28.6 (42) | 9.8 (1712) | 11.5 (1762) | 0.4 (1708) | 0.6 (1239) | 27.7 (891) | 20 (441) | 17.6 (85) | 34.2 (681) | | 24.8 (1759) | |
| <i>Escherichia coli</i> | 2.3 (1010) | 21.4 (10476) | 88.6 (12470) | 44.3 (5001) | 10 (6653) | 18.2 (7399) | 4.6 (2493) | 12.5 (10253) | 10.4 (12489) | 14 (8623) | 20.2 (10431) | 18.5 (4840) | 16.6 (812) | 19.6 (10627) | 12.7 (12435) | 0.3 (11802) | 0.5 (5670) | 11 (5673) | 5.6 (5673) | 57.4 (1708) | 17.5 (1794) | 41.7 (741) | 46.3 (12448) | |
| <i>E. coli (Urine)</i> | 0.9 (3710) | 15 (3440) | 87.7 (1458) | 37.6 (442) | 5.9 (2412) | 13.9 (879) | 5.1 (219) | 5.1 (4252) | 7.2 (4455) | 11.3 (3423) | 13.6 (3869) | 25.3 (1434) | 0 (15) | 20.8 (4252) | 12 (4449) | 0.3 (245) | 0.8 (1977) | 4.9 (1412) | 4.1 (4404) | 61.4 (140) | 17.5 (140) | | 48.3 (4454) | |
| <i>E. coli (Non urine)</i> | 2.7 (3338) | 21.3 (3788) | 88.9 (14784) | 53 (2331) | 12.6 (2557) | 17.8 (4605) | 1.8 (1096) | 15.1 (3584) | 13.3 (4764) | 20.5 (3667) | 9.9 (4139) | 17.5 (1990) | 19.5 (469) | 18.5 (4347) | 0.3 (4734) | 0.5 (4318) | 13.8 (1498) | 0.5 (2844) | 16 (314) | | | 57.1 (141) | 44.5 (4776) | |
| <i>H. influenzae</i> | 3.5 (395) | 11.7 (497) | 9.3 (54) | | | | 1.2 (481) | | | | 1.6 (128) | | 11.3 (497) | 0 (22) | 0 (8) | 9.8 (41) | 10.3 (29) | | | | | | 3.8 (52) | 39.9 (316) |
| <i>H. influenzae (Invasive)</i> | 4.8 (21) | 10.7 (28) | | | | | 7.1 (28) | | | | 3.6 (28) | | 0 (5) | | | 40 (9) | 42.9 (17) | | | | | | | 14.3 (14) |
| <i>H. influenzae (Non invasive)</i> | | | | 7.8 (51) | 0 (3) | | | | | | | | 11.8 (77) | 0 (22) | 0 (36) | 5.6 (22) | 0 (2) | | | | | | 2 (50) | 41.1 (302) |
| <i>K. pneumoniae</i> | 7 (8721) | 23.1 (9872) | 98.7 (11538) | 31.8 (7044) | 25.5 (5306) | 20.1 (8680) | 20.4 (1554) | 18.5 (10357) | 18.4 (11592) | 20.9 (9017) | 28 (8800) | 24 (3390) | 13.9 (711) | 11.8 (10490) | 15 (11544) | 0.7 (10873) | 1.4 (4337) | 17.7 (6028) | 32.5 (1719) | 45.3 (2926) | 19.7 (1237) | | 22.9 (11501) | |
| <i>M. morganii</i> | 1.8 (388) | 91.4 (453) | 95.2 (528) | 59.9 (314) | 2.3 (261) | 6.2 (70) | 4.3 (47) | 6.4 (535) | 5 (400) | 6.8 (456) | 66 (458) | 84.4 (179) | 31 (49) | 9.3 (49) | 14.3 (532) | 0.8 (494) | 1.2 (161) | 5.5 (290) | 7.2 (92) | 12.2 (89) | 3.3 (50) | | 29.7 (859) | |
| <i>P. aeruginosa</i> | 8.9 (11733) | 96.7 (2421) | 97.9 (292) | 90.7 (248) | 14.2 (8419) | 18.5 (7085) | 16.2 (4494) | 60.2 (993) | 14.5 (11882) | 42 (181) | 42 (288) | 91 (42) | 4.8 (198) | 81.8 (11437) | 11.7 (11725) | 16.7 (11722) | 13.4 (6190) | 16 (6063) | 18.9 (143) | 92.3 (4688) | 15.1 (7631) | 11.5 (1353) | 80.7 (353) | |
| <i>B. pseudomallei</i> | 82.3 (192) | 11.4 (187) | 84.7 (75) | 5.7 (87) | 12.2 (82) | 6.3 (143) | 4.3 (46) | 11.8 (51) | 1.8 (28) | 27.6 (58) | 74 (73) | 6.8 (88) | 25 (204) | 95.5 (201) | 1.5 (202) | 4.7 (64) | 88.6 (132) | 4.3 (89) | 7.9 (89) | 18.8 (89) | 58.3 (118) | | 58.3 (180) | |
| <i>P. mirabilis</i> | 3 (1826) | 14.6 (2397) | 46.5 (2607) | 19.1 (1883) | 4.3 (1343) | 6.7 (2159) | 2.4 (336) | 4.6 (2205) | 2.4 (2631) | 5.1 (2178) | 17.2 (2137) | 22.4 (812) | 56.7 (90) | 9.1 (244) | 12.2 (2618) | 1 (2544) | 1.1 (965) | 8.1 (1141) | 89.7 (348) | 17 (893) | 2.5 (18) | | 40 (18) | |
| <i>Salmonella sp</i> | 0 (87) | 1.8 (57) | 19.3 (792) | 9.1 (53) | 0 (20) | 2.4 (85) | 0 (12) | 0 (84) | 1.7 (804) | 3.6 (93) | 0 (84) | 5.7 (750) | 0.5 (739) | 4.2 (96) | 0 (5) | 0 (14) | 0 (5) | 0 (14) | 19 (71) | 40 (21) | 19 (71) | | 50.8 (429) | 23.1 (785) |
| <i>S. marcescens</i> | 4.6 (240) | 88.4 (288) | 90.7 (332) | 89.5 (191) | 2.5 (157) | 6.3 (300) | 0 (48) | 5.5 (282) | 3.8 (345) | 6.4 (233) | 81.7 (312) | 73.1 (108) | 27.3 (86) | 1.1 (278) | 4.1 (341) | 0.7 (279) | 2 (96) | 6.7 (149) | 84 (25) | 7.7 (52) | 41.5 (41) | 5.1 (332) | 76.2 (21) | |
| <i>S. malthophilia</i> | 39.8 (535) | 82.3 (515) | 94.4 (248) | 81.9 (288) | 37.9 (389) | 37.8 (349) | 22.2 (182) | 75.9 (212) | 26.6 (827) | 86.7 (233) | 94.8 (246) | | 42.9 (121) | 11.8 (701) | 55 (847) | 93 (883) | 79.4 (393) | 32.6 (282) | | 65.5 (367) | 76.1 (184) | 6.8 (732) | | |

1 No. tested

Appendix 8 (iii)

PERCENTAGE OF ANTIBIOTIC RESISTANCE AMONG GRAM NEGATIVE BACTERIA
2007

| Organism | Amikacin | Amoxicillin/ Clavulanic acid | Ampicillin | Ampicillin/ Sulbactam | Cefepime | Cefoperazone | Cefoperazone/ Sulbactam | Cefotaxime | Ceftazidime | Ceftriaxone | Cefturoxime sodium | Cephalexin | Chloramphenicol | Ciprofloxacin | Gentamicin | Imipenem | Meropenem | Netilmicin | Nitrofurantoin | Piperacillin | Piperacillin/ Tazobactam | Tetracycline | Trimethoprim/ Sulfamethoxazole | |
|---------------------------------|----------------|------------------------------------|-----------------|--------------------------|-----------------|-----------------|----------------------------|-----------------|-----------------|-----------------|-----------------------|----------------|-----------------|-----------------|-----------------|-----------------|-----------------|----------------|----------------|----------------|-----------------------------|---------------|-----------------------------------|---------------|
| <i>A. baumannii</i> | 29.2 (4298) | 62.3 (871) | 94.7 (4241) | 38.3 (4302) | 53.2 (1448) | 77.8 (3236) | 14.4 (921) | 75.4 (3887) | 46.8 (802) | 83.2 (874) | 85.5 (34) | 94.1 (54) | 79.6 (3880) | 43.9 (4176) | 36.4 (4916) | 46.6 (2593) | 47.7 (2093) | 19.3 | | 53.9 (1685) | 47.5 (2333) | | 37.8 (1446) | |
| <i>C. freundii</i> | 5.9 (202) | 70.8 (212) | 80.6 (227) | 49 (98) | 25.5 (94) | 19.2 (130) | 11.3 (53) | 30.3 (142) | 29 (207) | 34.1 (182) | 36.6 (186) | 86.9 (61) | | 15.3 (203) | 20.8 (221) | 1.4* (219) | 1.9* (155) | 30.7 (75) | | 29.4 (34) | 19.7 (61) | | 39.6 (225) | |
| <i>Enterobacter sp.</i> | 3.2 (2073) | 83.6 (2082) | 93.2 (2314) | 57.5 (958) | 5.7 (871) | 15.9 (1646) | 12.8 (468) | 19.4 (1781) | 17.8 (2174) | 21.4 (1740) | 35.5 (1995) | 79 (834) | 25 (176) | 4.4 (2105) | 7.8 (2246) | 1* (2156) | 1.2* (1210) | 8.8 (873) | 25.1 (335) | 29.5 (380) | 13.7 (582) | | 22 (2266) | |
| <i>E. coli</i> (all) | 2.2 (10286) | 21.7 (11341) | 69.3 (13239) | 34.9 (4903) | 23.3 (4916) | 18.1 (6751) | 7.1 (2817) | 15.1 (10321) | 15.2 (11478) | 16.3 (9720) | 19.6 (11427) | 25.8 (4033) | 24.8 (596) | 18 (11610) | 11.7 (12760) | 0.4* (11854) | 0.2* (6753) | 6.3 (4343) | 6.5 (5448) | 51.2 (2279) | 5.8 (3230) | | 44.1 (13080) | |
| <i>E. coli</i> (Urine) | 1.9 (3927) | 17.6 (4220) | 88.4 (5438) | 29.6 (1144) | 19.9 (1698) | 17.2 (1344) | 5.9 (1093) | 12.7 (4171) | 13.8 (4584) | 15 (3429) | 15.5 (4516) | 26.8 (2144) | 21.6 (11) | 19.9 (4902) | 11.8 (5300) | 0.4* (4716) | 0.2* (2723) | 5.6 (1853) | 6.6 (5300) | 56.5 (619) | 4.5 (1114) | | 46.8 (5436) | |
| <i>H. influenzae</i> (all) | | 6.5 (306) | 20.1 (348) | 9.3 (54) | | | | 5.6 (322) | 4.7 (343) | 8.1 (137) | | | | 8.6 (326) | 9 (43) | 9.1 (11) | 9.5 (63) | 7.7 (65) | | | | | | 32.2 (255) |
| <i>H. influenzae</i> (Invasive) | | 4 (25) | 19.4 (36) | | | | | 3.6 (28) | 0 (33) | | | | 2.7 (37) | | | 0 (15) | | | | | | | | 22.2 (18) |
| <i>K. pneumoniae</i> | 6.4 (12067) | 24.8 (13741) | 98.9 (15141) | 33 (8003) | 38.4 (6457) | 21.6 (10045) | 15.3 (2912) | 23.4 (12154) | 24.4 (13890) | 25.7 (11926) | 29.6 (13628) | 27.7 (3512) | 16.5 (832) | 12.5 (13551) | 17.7 (14501) | 0.5* (13973) | 0.8* (8327) | 17.8 (5010) | 26 (2516) | 36.1 (3067) | 12.8 (4784) | | 27 (14746) | |
| <i>M. morgani</i> | 1.2 (576) | 89.4 (667) | 93.3 (716) | 65.3 (354) | 1.7 (350) | 6.8 (400) | | 9.4 (587) | 4.6 (636) | 6.1 (604) | 74.9 (654) | | | 12 (598) | 12.7 (669) | 0.3 (671) | 1.2 (576) | | | | | | | 32.9 (703) |
| <i>P. aeruginosa</i> | 8.1 (15065) | 97.9 (7986) | 94.8 (173) | 97 (1366) | 13.4 (10687) | 16.1 (7823) | | 53.5 (864) | 13.7 (14321) | 6.7 (85) | 91.5 (176) | 58.3 (24) | 78.3 (166) | 11.5 (14057) | 12.5 (14666) | 13.5 (14941) | 13.3 (10600) | 15 (6850) | 96 (149) | 11.8 (4810) | 8.9 (13556) | | 94.3 (1454) | |
| <i>B. pseudomallei</i> | 88 (325) | 5.9 (220) | 96.5 (85) | 2.9 (137) | 8.2 (159) | 3.8 (183) | 0.9 (107) | 4.2 (72) | 2.8 (358) | 16.7 (60) | 79.1 (91) | | 5.1 (138) | 20.7 (305) | 97.1 (131) | 0.3* (371) | 2.8* (121) | 90.3 (134) | | 2.3 (131) | 0.9 (215) | 10 (160) | 45 (269) | |
| <i>P. mirabilis</i> | 14 (2573) | 12.7 (3185) | 46.1 (3376) | 13.1 (2101) | 9.9 (1502) | 7.6 (2217) | | 6 (2608) | 6.2 (3135) | 8.5 (2856) | 18.1 (2943) | 31.6 (77) | 52.2 (90) | 11.3 (3103) | 11.4 (3292) | 1.8* (3241) | 1* (1928) | 6.6 (1142) | 90.6 (498) | | 1.4 (1216) | | 39.4 (3284) | |
| <i>Salmonella sp.</i> | 0 (128) | 9.1 (110) | 24.8 (1015) | 7 (57) | 0 (26) | 1.1 (94) | | 0.9 (107) | 0.9 (112) | 1.2 (914) | 9.6 (114) | 6.2 (16) | 5.8 (831) | 1.4 (858) | 1.7 (119) | 0 (110) | 0 (87) | 5.3 (19) | 25 (16) | 18.6 (43) | 4.4 (68) | 36.9 (453) | 19.9 (1011) | |
| <i>S. marcescens</i> | 15.5 (434) | 85 (472) | 97.3 (518) | 86.5 (260) | 4.9 (265) | 2.4 (330) | 8.9 (146) | 6.7 (436) | 7.6 (474) | 5.8 (413) | 80.7 (481) | 85.9 (142) | 19.6 (511) | 1 (418) | 13.6 (492) | 3.3* (481) | 3.4* (264) | 3.9 (154) | 82.8 (29) | 7.5 (93) | 2.3 (129) | 94.4 (19) | 19.6 (511) | |
| <i>S. maltophilia</i> | 40.1 (664) | 88.1 (489) | 97.8 (186) | 91.3 (436) | 48.9 (417) | 52.3 (333) | 48.8 (213) | 86.7 (181) | 35.7 (658) | 91.2 (160) | 94.1 (187) | | | 27.3 (11) | 8.7 (801) | 46.7 (788) | 89.7 (862) | 33 (565) | | 78.5 (303) | 50.8 (327) | 77.5 (510) | 7 (791) | |

† No. tested

Appendix 9 (ii)

PERCENTAGE OF ANTIBIOTIC RESISTANCE AMONG GRAM POSITIVE BACTERIA
2006

| Organisms | Amikacin | Imipenem/Cilastatin | Ampicillin | Colistin | Cefazolin | Ceftazidime | Ceftiofur | Cefuroxime sodium | Chloramphenicol | Ciprofloxacin | Clindamycin | Erythromycin | Fusidic acid | Gentamicin | Gentamicin-High | Imipenem | Methicillin | Mupirocin | Nitrofurantoin | Penicillin G | Rifampin | Ridampin | Tetracycline | Trimethoprim/Sulfamethoxazole | Vancomycin | |
|------------------------------------|--------------|---------------------|---------------|--------------|--------------|--------------|--------------|-------------------|-----------------|---------------|--------------|--------------|--------------|--------------|-----------------|--------------|-------------|------------|----------------|---------------|-------------|----------|---------------|-------------------------------|-------------|-------------|
| <i>E. faecalis</i> | 0 (1) | 3.7 (27) | 4 (998) | | | | | | 31.2 (32) | 33.9 (148) | | | | 29.4 (17) | 19.3 (54) | 9.7 (52) | | | 2.5 (10) | 16.7 (108) | | | 62.7 (127) | 17.2 (44) | 0.7 (5) | |
| <i>E. faecium</i> | | 18.2 (11) | 54.5 (146) | | | | | | 28.6 (14) | 62.1 (58) | | | | 66.7 (8) | 46.6 (13) | 60.7 (28) | | | 43.9 (57) | 56.8 (44) | | | 77.8 (5) | 53.1 (8) | 1.4 (14) | |
| <i>Enterococcus</i> sp | 7.7 (13) | 25.6 (47) | 29.6 (237) | 14.1 (89) | 32.8 (11) | 13.9 (7) | 35.7 (12) | 19.5 (8) | 25.3 (13) | 36 (107) | | | | 27.8 (5) | 27 (18) | 15.6 (16) | | | 12 (8) | 38.3 (47) | | | 77.1 (4) | 32.1 (1) | 9 (2) | |
| <i>S. aureus</i> (all isolates) | 64.3 (54) | 66.5 (60) | 66.5 (27) | | 89.6 (54) | 92.4 (52) | 25 (8) | 6.9 (1) | 31 (7) | 7.3 (1) | 31.3 (17) | 6.3 (1) | 29.5 (15) | | 15.4 (1) | 31.5 (14) | 0.2 (1) | | 7.1 (3) | 84.3 (94) | | | 5.5 (1) | 33.3 (1) | 26.7 (1) | 0.1* (1) |
| <i>S. aureus</i> (ICU isolates) | 86.5 (52) | 59 (16) | | | 88.1 (52) | 92.5 (51) | 0 (1) | 51.4 (1) | 2.4 (1) | 44.6 (1) | 7.1 (1) | 45.1 (1) | | | 41.9 (1) | 1.3 (1) | 10 (1) | | 86.6 (1) | 4.1 (1) | | | 38 (1) | 0.4* (1) | | |
| <i>S. aureus</i> (MRSA) | 66.5 (50) | 96.2 (78) | | | 93.8 (51) | 93.8 (51) | | 15.3 (1) | 72.9 (1) | 17.7 (1) | 91.7 (1) | 7.3 (1) | 91.5 (1) | | 100 (1) | 0.2 (1) | 31.3 (1) | | 99.5 (1) | 15.7 (1) | | | 81.5 (1) | 0.1* (1) | | |
| <i>Staph</i> Coag-neg | 20.2 (46) | 13.8 (8) | 55.1 (138) | | 47.6 (8) | 83.5 (48) | 79.6 (45) | 55.2 (8) | 19.4 (4) | 25.9 (4) | 18.5 (4) | 49.7 (4) | 18.6 (4) | 41.7 (4) | | 6.2 (4) | 63.5 (3) | 0.3 (1) | 18.5 (4) | 78.9 (4) | 35.9 (4) | | 14.1 (4) | 33.3 (1) | 38.1 (1) | 0.3* (1) |
| <i>S. agalactiae</i> | | | | | | | | | | | | | | | | | | | | | | | | | | |
| <i>S. pyogenes</i> | 0 (24) | 1.2 (33) | 2.9 (8) | 0 (8) | 0 (2) | 0 (2) | 0 (2) | 0 (2) | | | 5.3 (1) | 6 (1) | | | | | | | | | | | | 49.6 (1) | 5.2 (1) | 0 (1) |
| <i>Strep</i> Gp A | | 16.7 (6) | 1.4 (18) | 1.7 (5) | 1.5 (2) | 1.2 (1) | 1.2 (1) | 1.2 (1) | 6.7 (1) | 16.7 (1) | 3.5 (1) | 7.4 (1) | 30 (1) | 36 (1) | | | | | | | | | | 51.8 (1) | 18.5 (1) | 0 (1) |
| <i>Strep</i> Gp B | | 8.7 (2) | 0 (2) | 0 (2) | 0 (1) | 3.5 (1) | 1.4 (1) | 12.2 (1) | 7.4 (1) | 5.5 (1) | 27.3 (1) | 90.1 (1) | | | | | | | 6.6 (1) | 1.9 (1) | | | 61.4 (1) | 18.5 (1) | 1* (1) | |
| <i>S. pneumoniae</i> | | | | | | | | | | | | | | | | | | | | | | | | | | |
| <i>S. pneumoniae</i> (invasive) | | | | | | | | | | | | | | | | | | | | | | | | | | |
| <i>S. pneumoniae</i> (noninvasive) | | | | | | | | | | | | | | | | | | | | | | | | | | |

[] No. tested
* Not verified

Appendix 9 (iii)

NATIONAL ANTI-BIOTIC GUIDELINE 2008

PERCENTAGE OF ANTIBIOTIC RESISTANCE AMONG GRAM POSITIVE BACTERIA
2007

| Organisms | Amikacin | Amoxicillin/ Clavulanic acid | Ampicillin | Cefepime | Cefotaxime | Ceftazidime | Ceftazoxime | Cefuroxime axelam | Chloramphenicol | Ciprofloxacin | Clindamycin | Erythromycin | Fusidic acid | Gentamicin | Gentamicin- high | Imipenem | Methicillin | Nitrofurantoin | Penicillin G | Piperacillin | Rifampin | Tetracycline | Trimethoprim/ Sulfamethoxazole | Vancomycin |
|---------------------------------|---------------|---------------------------------|---------------|---------------|---------------|---------------|---------------|----------------------|-----------------|---------------|----------------|-----------------|----------------|-----------------|---------------------|--------------|-----------------|----------------|-----------------|----------------|---------------|----------------|-----------------------------------|----------------|
| <i>E. faecalis</i> | | 4.5 (179) | 6.3 (820) | 82.4 (17) | 77.3 (22) | 100 (1) | 66.9 (180) | 83.3 (6) | 32.4 (182) | 26.8 (295) | 97.1 (70) | 55.6 (266) | | 35 (452) | 22.1 (625) | 8.5 (47) | | | 36.7 (283) | | | 83.5 (297) | 33.9 (825) | 0.4* (1011) |
| <i>E. faecium</i> | | 72.1 (43) | 65.3 (239) | 100 (7) | 58.3 (12) | | 83.3 (36) | 100 (2) | 8.2 (49) | 73.1 (104) | 69 (29) | 81.4 (70) | | 63.5 (115) | 53.2 (190) | 82.6 (23) | | | 63.5 (85) | | | 85.9 (99) | 66.4 (211) | 0 (283) |
| <i>Enterococcus</i> sp | | 22 (246) | 26.4 (168) | 69.2 (19) | 67.9 (106) | 0 (8) | 78.4 (8) | | 27.7 (23) | 42.9 (99) | 84 (13) | 67.6 (139) | | 33.6 (40) | 32.3 (127) | 11.1 (9) | | | 50.8 (38) | | | 74.7 (37) | 45.9 (130) | 0.9* (176) |
| <i>S. aureus</i> (all isolates) | 78 (617) | 20 (10) | 89.8 (868) | 24.1 (29) | 50 (6) | 90.1 (627) | 98.2 (556) | | 4 (1196) | 32 (388) | 8.5 (867) | 30.4 (19927) | 7.4 (17163) | 27.6 (19922) | | 30.8 (13) | 28.8 (14986) | 3.8 (889) | 82.6 (10494) | | 4.2 (1953) | 58 (25) | 26 (11758) | 0 (19875) |
| <i>S. aureus</i> (ICU isolates) | 66.6 (62) | | 71.4 (21) | | | 96.5 (57) | 98.1 (53) | | 3.8 (52) | 34.8 (419) | 10.2 (256) | 32.9 (947) | 6.9 (685) | 28.8 (348) | | 30 (952) | 13 (23) | 83.5 (757) | | 3.5 (949) | | 30.1 (602) | 0 (855) | |
| <i>S. aureus</i> (MRSA) | 80.9 (589) | | | | | | | | 18.2 (137) | 59.1 (580) | 24.4 (245) | 95 (428) | 6.5 (365) | 93.5 (427) | | | 19 (126) | | 13.5 (429) | | | 89.3 (330) | 0 (431) | |
| Staph Coag-neg | 14.8 (539) | 16.7 (68) | 69.6 (678) | 63.6 (22) | 47 (66) | 88.4 (533) | 93.8 (514) | | 14.2 (930) | 14.2 (835) | 15.2 (3487) | 51.4 (8397) | 23.9 (7790) | 36.4 (8739) | | 15.2 (66) | | 8 (4046) | 80.2 (62) | 53.2 (8359) | 14.2 (9) | 22.2 (1770) | 37.1 (1771) | 0.3* (8855) |
| Group B Streptococcus | 4.5 (440) | 1.4 (1735) | 3.1 (519) | 2.5 (1248) | | 9.2 (222) | 2.8 (213) | | 7 (2145) | | 6.9 (6380) | 5.3 (7294) | | 74.6 (1202) | | | | 5.7 (158) | 2.3 (7361) | | | 63.8 (7194) | 27.9 (7284) | 1.2 (1897) |
| Group A Streptococcus | | 0 (47) | 0.7 (304) | 0 (35) | 2.7 (183) | | | 1.8 (183) | 11.8 (17) | 6.7 (277) | | 3.7 (1056) | 5.7 (1183) | | 1.9 (25) | | | | 2 (1181) | | | 49.9 (1137) | 30.4 (1184) | 0.8 (24) |
| <i>S. pneumoniae</i> | | 0 (28) | 12.5 (8) | 0 (24) | 0 (85) | 0 (4) | 0 (400) | 0.5 (24) | 7.3 (185) | 10.4 (77) | 22.8 (466) | | 31.6 (19) | | | | | | 15.1 (269) | | | 35.1 (465) | 38.7 (450) | 0.2 (455) |
| <i>S. pneumoniae</i> (invasive) | | 0 (12) | 0 (3) | 0 (12) | 0 (30) | | | 0 (134) | 0 (12) | 0 (52) | 20 (10) | 21.9 (146) | | 20 (10) | | | | | 15.5 (97) | | | 33.1 (133) | 36.1 (147) | 0.7 (144) |

[] No. tested
* Not verified

Appendix 10 (i)

NATIONAL ANTI-BIOTIC GUIDELINE 2008

**COMMON ISOLATES FROM INTENSIVE CARE UNIT (ICU)
2006**

| Organism | HSAJB | HKL | HKL+PAEDS | HKT | HPP | HTF | HSEL | HMLK | HTAA | HSB | All Hospital |
|----------------------------------|---------------|---------------|---------------|--------------|---------------|--------------|---------------|---------------|--------------|--------------|----------------|
| <i>Staphylococcus aureus</i> | 14.2 [407] | 15.6 [333] | 16.1 [361] | 10.5 [30] | 15.8 [109] | 7.6 [13] | 10.8 [59] | 10.7 [124] | 14.7 [66] | 17.5 [48] | 14.2 [1550] |
| <i>Pseudomonas aeruginosa</i> | 12.8 [366] | 11.6 [249] | 16.4 [367] | 10.1 [29] | 16.6 [115] | 19.4 [33] | 10.5 [57] | 12.8 [148] | 17.4 [78] | 17.5 [48] | 13.6 [1490] |
| <i>Klebsiella pneumoniae</i> | 20 [573] | | | 15.4 [44] | 9.2 [64] | 15.2 [26] | 18.6 [101] | 21.8 [252] | 10.5 [47] | 12.7 [35] | 10.5 [1142] |
| <i>Coag-negative Staph (SCN)</i> | 0.5 [143] | 14.6 [313] | 13.5 [301] | 8.4 [24] | 15.5 [107] | | 16.2 [88] | 4.3 [49] | 10.7 [48] | | 9.8 [1073] |
| <i>Acinetobacter sp.</i> | 0.4 [11] | 11.6 [249] | 13.2 [296] | 0.7 [2] | 10.5 [73] | 7.6 [13] | 14 [76] | 5 [58] | 1.8 [8] | 18.2 [50] | 7.7 [836] |
| <i>A. baumannii (anitratu)</i> | 13.4 [384] | | | 22.7 [65] | | | | 10 [116] | 17 [76] | | 6 [656] |
| <i>Escherichia coli</i> | 4.2 [120] | 7.6 [162] | [72] | 3.8 [11] | 4.3 [30] | eco [9] | 5.4 [29] | 4.5 [52] | 3.8 [17] | 4 [11] | 4.6 [513] |
| <i>Candida sp.</i> | 10.8 [308] | 2.6 [56] | 2.8 [63] | 2.4 [7] | | 2.9 [5] | | | 0.6 [3] | | 4 [442] |
| <i>Klebsiella sp.</i> | | 9.6 [206] | 9.9 [222] | 2.4 [7] | 0.1 [1] | | 0.4 [2] | | 0.2 [1] | | 4 [439] |
| <i>Enterobacter sp.</i> | 4.1 [116] | 4 [86] | 3.4 [76] | 1.7 [5] | 2.9 [20] | 3.5 [6] | 5.2 [28] | 3.2 [37] | 2.5 [11] | 4.4 [12] | 3.6 [397] |
| <i>Candida albicans</i> | 1.5 [42] | 4.5 [97] | 4.4 [98] | 3.1 [9] | | 5.3 [9] | | | | | 2.3 [255] |
| Total Isolates | 2863 | 2141 | 2237 | 286 | 692 | 170 | 542 | 1152 | 448 | 274 | 10916 |

[] No. isolated

HPP - Hospital Pulau Pinang HMEL - Hospital Melaka HIPH - Hospital Ipoh
 HKL - Hospital Kuala Lumpur HTAA- Hospital Tengku Ampuan Afzan HTJ - Hospital Tuanku Jaafar
 HSAJB - Hospital Sultanah Aminah HTAR - Hospital Tuanku Rahimah HSB - Hospital Sultanah Bahiyah
 HSNZ - Hospital Sultanah Nur Zahirah HTF - Hospital Tuanku Fauziah HSEL - Hospital Selayan

Appendix 10 (ii)

COMMON ISOLATES FROM INTENSIVE CARE UNIT (ICU)
2007

| Organism | HSAJB | HKL | HSNZ | HPP | HKGR | HSEL | HMEL | HTAA | HSB | HIPH | HTJ | All Hospital |
|-------------------------------|-------------|-------------|------------|-------------|------------|------------|-------------|------------|------------|------------|------------|--------------|
| <i>Staphylococcus aureus</i> | 12 [362] | 18 [133] | 11 [54] | 15 [65] | 8 [20] | 15 [33] | 17 [254] | 13 [27] | 15 [53] | 8 [3] | 11 [52] | 14 [1056] |
| <i>Pseudomonas aeruginosa</i> | 12 [366] | 25 [188] | 14 [72] | 21 [93] | 20 [54] | 12 [26] | 11 [161] | 14 [28] | 11 [37] | 3 [1] | 10 [50] | 19 [1076] |
| <i>Klebsiella pneumoniae</i> | 21 [654] | 14 [103] | 14 [70] | 11 [48] | 6 [16] | 15 [34] | 19 [285] | 11 [22] | 14 [49] | 8 [3] | 9 [43] | 18 [1327] |
| <i>Coag-negative Staph</i> | 6 [173] | 5 [36] | 17 [87] | 10 [45] | 15 [41] | 17 [39] | 6 [91] | 18 [38] | 9 [30] | 11 [4] | 12 [57] | 10 [641] |
| <i>Acinetobacter sp.</i> | 14 [441] | 14 [100] | 15 [78] | 18 [80] | 11 [29] | 14 [31] | 13 [198] | 10 [20] | 19 [67] | 32 [12] | 13 [66] | 15 [1122] |
| <i>Escherichia coli</i> | 4 [139] | 2 [18] | 5 [26] | 4 [17] | 7 [19] | 7 [15] | 6 [93] | 5 [11] | 5 [18] | 11 [4] | 4 [20] | 4 [380] |
| <i>Candida albicans</i> | 2 [47] | 2 [14] | 2 [10] | | 2 [6] | | 3 [40] | | | 3 [1] | 2 [12] | 1.5 [130] |
| <i>Candida sp.</i> | 9 [274] | 3 [21] | 3 [14] | | 3 [7] | | 2 [26] | | | 8 [3] | 2 [12] | 7 [357] |
| <i>Enterobacter sp.</i> | 4 [126] | 4 [33] | 2 [10] | 4.4 [20] | 0.7 [2] | 2 [5] | 2.6 [39] | 3 [6] | 3 [11] | 8 [3] | 2 [9] | 3 [264] |
| Total Isolates | 2582 | 646 | 421 | 368 | 194 | 183 | 1187 | 152 | 265 | 34 | 321 | 6353 |

[] No. isolated

HPP - Hospital Pulau Pinang HMEL - Hospital Melaka HIPH - Hospital Ipoh
 HKL - Hospital Kuala Lumpur HTAA - Hospital Tengku Ampuan Afzan HTJ - Hospital Tuanku Jaafar
 HSAJB - Hospital Sultanah Aminah HTAR - Hospital Tuanku Rahimah HSB - Hospital Sultanah Bahiyah
 HSNZ - Hospital Sultanah Nur Zahirah HTF - Hospital Tuanku Fauziah HSEL - Hospital Selayang

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