







National and Regional Antibiotic Guidelines

Bart Currie

Infectious Diseases Department, Royal Darwin Hospital
Global and Tropical Health Division, Menzies
Northern Territory Medical Program, Flinders & CDU

Standard Treatment Guidelines

- Delegation of most management to non-medical staff
- Availability of therapy for the whole population
- Early institution of appropriate therapy
- Implementation of a rational essential drugs policy
- Minimisation of inappropriate therapy
- Correct dosing



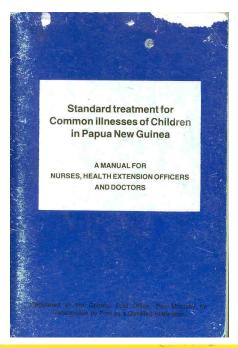
Standard Treatment Guidelines

Antibiotic Guidelines

Delay emergence of antimicrobial resistance

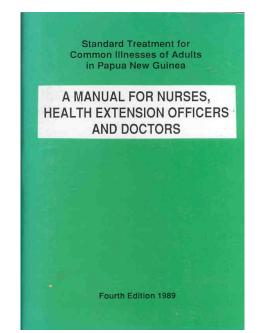
Year	PNG	NT	TG: Abiotic
1966	MO's Pocket Book	Diarrhoea "slide-rule"	
1974	1st Ed Paediatric STM		
1977	1st Ed Adult STM		
1978			1 st Ed
1980s		ARI & Syphilis Protocols	
1992		1st Ed CARPA STM	7 th Ed
			Remote Section
1998			10 th Ed Remote Mainstreamed
2003		4 th Ed CARPA	12 th Ed
2009/10		5 th Ed CARPA	14 th Ed

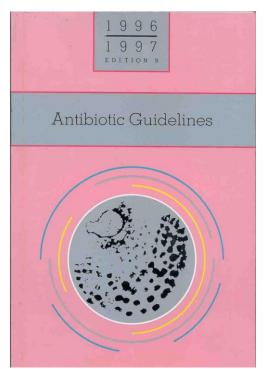
Year	PNG	NT	TG: Abiotic
1966	MO's Pocket Book	Diarrhoea "slide-rule"	
1974	1st Ed Paediatric STM		
1977	1st Ed Adult STM		
1978	1996		1 st Ed
1980s	1997 EDITION 9	CENTRAL AND NORTHERN AUSTRALIA AND OTHER REMOTE AREAS The following guidelines have been developed primarily for use by doctors, nurses and Aboriginal health workers in rural Aboriginal	
1992	Antibiotic Guidelines	communities. High morbidity and mortality from bacterial infections justify early appropriate empirical antimicrobial therapy. Emphasis is placed on supervised regimens where possible, utilising intramuscular and single dose therapies and less frequent dosing. Storage requirements of antimicrobials must also be considered. Bicillin All-Purpose is a heat-stable powder for intramuscular injection which contains a mixture of benzyl, procaine and benzathine penicillin. If it is not available benzathine penicillin alone (Bicillin L-A) can be used, see Table 6, p. 201.	7 th Ed Remote Section
1998		Studies on organism prevalence and antimicrobial resistance patterns in the Northern Territory, northern Western Australia and northern Queensland are reflected in the guidelines. Streptococcus pyogenes remains important, with subsequent rheumatic heart disease and glomerulonephritis occurring. Erythromycin and roxithromycin resistance in Staphylococcus aureus is common, as is Escherichia coli resistance to (amoxy)ampicillin, sulphonamides and trimethoprim. Beta-lactamase production in Haemophilus influenzae has emerged. Penicillinase-producing Neisseria gonorrhoeae (PPNG) remains rare in communities, but ongoing	10 th Ed Remote Mainstreamed
2003		surveillance is critical. Reference is made to the main guidelines where regional circumstances do not apply. Doses are generally not specified in the text of this section. Refer to Table 6, p. 199, for weight-related drug doses unless detailed in the text.	12 th Ed
2009/10		Timing of doses in relation to food appear in the main guidelines, see Table 7, p. 203.	14 th Ed

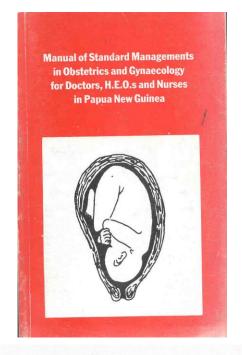












CENTRAL AND NORTHERN AUSTRALIA AND OTHER REMOTE AREAS

The following guidelines have been developed primarily for use by doctors, nurses and Aboriginal health workers in rural Aboriginal communities. High morbidity and mortality from bacterial infections justify early appropriate empirical antimicrobial therapy.

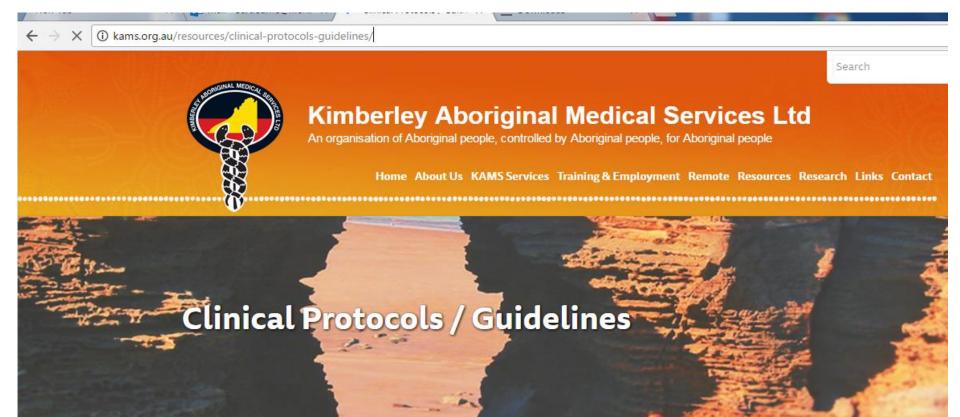
Emphasis is placed on supervised regimens where possible, utilising intramuscular and single dose therapies and less frequent dosing. Storage requirements of antimicrobials must also be considered. Bicillin All-Purpose is a heat-stable powder for intramuscular injection which contains a mixture of benzyl, procaine and benzathine penicillin. If it is not available benzathine penicillin alone (Bicillin L-A) can be used, see Table 6, p. 201.

Studies on organism prevalence and antimicrobial resistance patterns in the Northern Territory, northern Western Australia and northern Queensland are reflected in the guidelines. Streptococcus pyogenes remains important, with subsequent rheumatic heart disease and glomerulonephritis occurring. Erythromycin and roxithromycin resistance in Staphylococcus aureus is common, as is Escherichia coli resistance to (amoxy)ampicillin, sulphonamides and trimethoprim. Beta-lactamase production in Haemophilus influenzae has emerged. Penicillinase-producing Neisseria gonorrhoeae (PPNG) remains rare in communities, but ongoing surveillance is critical.

Reference is made to the main guidelines where regional circumstances do not apply.

Doses are generally not specified in the text of this section. Refer to Table 6, p. 199, for weight-related drug doses unless detailed in the text.

Timing of doses in relation to food appear in the main guidelines, see Table 7, p. 203.



The Kimberley Protocols were developed from a desire to standardise the screening for and management of health conditions more prevalent in the Kimberley and recognising that the management of these conditions differs from standard management due to the extreme remoteness of the area. Their purpose is to enable care to be initiated by a range of health care professionals working within their own scope of practice when direct access to

Minimalist Antibiotics in Alice in the early 1990s

Penicillin

Gentamicin

Chloramphenicol

Metronidazole



Therapeutic Guidelines: Antibiotic

- 1st Edition 1978
 - 31 Pages and 31 Antimicrobial agents
- 11th Edition 2000
 - 318 Pages and 100 Antimicrobial agents
- 15th Edition 2014
 - 677 Pages and ?? Antimicrobial agents

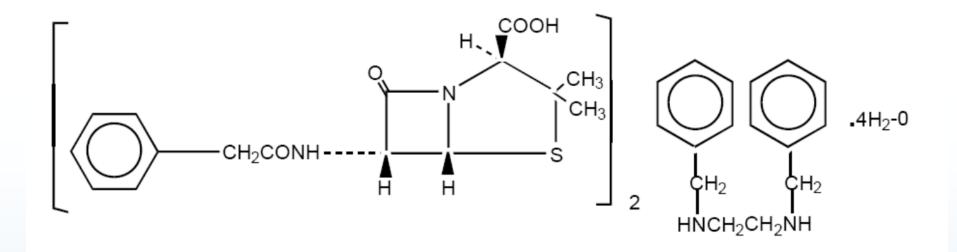


Supervised Therapy

- Single dose IM benzathine penicillin for throat/skin
- 4-weekly IM benzathine penicillin for ARF prophylaxis
- DOTS for TB
 - √ 95% completion of 6 months Tx in NT
- Daily IM procaine penicillin for pneumonia
- Single dose nitrofurantoin for cystitis
- Topical scabies therapy
- STI protocols
 - ZAP pack: azithro/amoxy/probenecid syndromic Tx
 - IM benzathine penicillin for syphilis
- IM iron for anaemia



Benzathine Penicillin G (BPG)



- Composed of two molecules of benzyl penicillin attached to a compound called benzathine which allows it to be released slowly from the muscle
- Allows low concentrations for 2-4 weeks



BPG – history

- BPG was developed by J. Lester Szabo in 1951, and the first BPG patent appears to have been granted in the United States of America in 1953
- □ The original clinical application was for the syphilitic infections, which it is still used for today
- A patent was granted for a stabilised powdered formulation in 1967



BPG – history prior to 1995

The availability of an appropriate formulation of BPG in Australia has been a challenge with multiple manufactures and shortages



- 4 mL Bicillin LA with large 18 gauge "horse" needle
- Cillicaine 1.5g syringe
- Bicillin AP powder



BPG – history prior to 1995

The availability of an appropriate formulation of BPG in Australia has been a challenge with multiple manufactures and shortages



MUCH CONFUSION!
About dosing



BPG – history 1995 to 2006

After much lobbying a 900mg formulation of Bicillin became available in 1995



- Administered via Tubex device
- Major stock outs in 2001 and 2004
- Withdrawn from Australia in July 2006



BPG – history 2006 to 2008

- □ Therapeutic Goods Administration (TGA) provided S19A approval for the use of "Pan Benzathine Penicillin"
- Powdered product with large volume
 - 4.6mL vs 2mL
 - Issues with clogging of needles (40%)
 - Use of lignocaine as diluent to reduce pain
 - Less viscous
 - Not cold





BPG – history 2008 to present

- Bicillin LA returned to Australian market in 2008
- Pre-filled syringe with 21 gauge needle
- Only available as a 900mg product
- No markings for dosing in children
- Needs refrigeration





BPG – history 2008 to present

- □Stock shortages continue; December 2012 to January 2013 and February to May 2014
- ■No ability to deliver smaller doses accurately without decanting from original syringe
- ■No data on mixing formulation with lignocaine
 - □ The addition of 0.5–1 mL of 1% lignocaine is used with powdered products but is not recommended with preloaded syringes currently available in Australia

Azithromycin

- Early 1990's marketed in Australia in adults for respiratory and skin infections
- Not initially marketed for chlamydia
- Initially no pediatric formulation
- But MAGICAL for
 - Chlamydial urethritis
 - Trachoma
 - Donovanosis

Azithromycin

- 1994 STI protocols in WA, NT genital chlamydia
- 1995 First use for trachoma in Australia NT
 - capsule contents mixed with Xantham gum
 - NT granted TGA SAS Category A approval for importation of pediatric suspension – trachoma deemed "life-endangering"
- 1996/7 9th Edition Antibiotic Guidelines indications
 - genital chlamydia STIs
 - trachoma
 - donovanosis

Azithromycin

- Emergence of resistance with antibiotic selection pressure
 - not in chlamydia, but concern in skin & throat bacteria
 - >S. aureus, S. pyogenes and S. pneumoniae

long tissue half life – low levels > 1 week – make azithromycin theoretically more likely to select resistance than other macrolides

 macrolide resistance emerged in WA in 1970's, common in Europe, Japan (> 50% in *S. pyogenes* in Japan, 1970's)

The role of gentamicin (in the NT)

- Ceftriaxone plus single dose gentamicin prior to evacuation of septic patient to RDH/ASH
- Ceftriaxone plus single dose gentamicin for adults with risk factors and moderate/severe community-acquired pneumonia
- Ampicillin plus 1-2 doses gentamicin for pyelonephritis





The body of a PNG national who died of TB in Australia is repatriated across the border last Thursday, 20 October, from Saibai. Photo: STEFAN ARMBRUSTER, SBS.



 What was the commonest antibiotic used on the adult medical wards at Port Moresby General Hospital in 2013?





- What is the commonest antibiotic used on the adult medical wards at Port Moresby General Hospital in 2013?
 - chloramphenicol

Steven Yennie, M Med Thesis 2013

- What is the commonest antibiotic used on the adult medical wards at Port Moresby General Hospital in 2013?
 - chloramphenicol

Steven Yennie, M Med Thesis 2013

Mortality in patients admitted to the adult wards at PMGH was:

4% in 1964, 10% in 1989 and 7% in 1997.

What was mortality in 2013 and what are the commonest causes?

- What is the commonest antibiotic used on the adult medical wards at Port Moresby General Hospital in 2013?
 - chloramphenicol

Steven Yennie, M Med Thesis 2013

Mortality in patients admitted to the adult wards at PMGH was:

4% in 1964, 10% in 1989 and 7% in 1997.

What was mortality in 2013 and what are the commonest causes?

- 32% of adult patients have died
- TB 25% of deaths and HIV/AIDS 24%

Vincent Pyakalyia, M Med Thesis 2013