Infectious Diseases Issues in Central Australia

IKTO Meeting – October 2013

Dr Saliya Hewagama Infectious Diseases Physician Alice Springs Hospital



Remote Clinics

Lake Nash

Ti Tree







Socioeconomic determinants of health – NT statistics

- 62.6% of Indigenous people were in community housing
- 34% of Indigenous households in the NT were overcrowded
 - 60.8% in those in community housing
- Australia-wide in "discrete Indigenous communities"
 - 53% used bore water as major water source, 11% used a river or reservoir
 - 53% used community generators for electricity

What's Common in Alice Springs?

- Skin & soft tissue infections
- Bronchiectasis
 - Prevalence 147 cases per 10 000 Indigenous children in Central Australia¹
- Strongyloides
- Sexually transmitted infections
 - NT has highest rate of Chlamydia, Gonorrhoea & Syphilis in the country ²
- Bacteraemia
 - Especially Staph aureus and Strep pneumoniae
- Human T cell lymphotropic virus 1 (HTLV-1)

² Chang AB, MJA 2008 ¹ Kriby Institute, 2012

Two nations: racial disparities in bloodstream infections recorded at Alice Springs Hospital, central Australia, 2001–2005

Lloyd J Einsiedel and Richard J Woodman

MJA 2010; 192: 567–571

3 Population-based BSI incidence rates among Indigenous and non-Indigenous patients admitted to Alice Springs Hospital, by age, 2001–2005



1 Patient characteristics, outcomes and comorbid conditions for Indigenous and non-Indigenous adults admitted to Alice Springs Hospital with BSIs, 2001-2005

	Indigenous patients	Non-indigenous patients	Р
Characteristics	(n=614)	(n = 69)	
Mean age, in years (SD)	44.0 (15.2)	57.3 (20.4)	< 0.001
Age > 65 years, n (%)	64 (10.4%)	29 (42.0%)	< 0.001
Male, n (%)	271 (44.1%)	43 (62.3%)	< 0.001
Residence outside region, n (%)*	110 (17.9%)	24 (34.8%)	< 0.001
Outcomes			
30-day mortality, n (%)	76 (12.4%)	6 (8.7%)	0.38 [†]
Overall mortality, n (%)	197 (32.1%)	9 (13.0%)	0.003 ⁺
Mean age at death, in years (SD)	48.5 (16.2)	75.1 (18.7)	< 0.001
Recurrent BSI, n (%) [‡]	91 (16.5%)	3 (5.0%)	0.019
Comorbid conditions, n (%)§	(n = 392)	(n = 44)	P¶
Diabetes mellitus	164 (41.8%)	6 (13.6%)	0.16
Alcohol dependence	152 (38.8%)	6 (13.6%)	0.21
Chronic renal failure	59 (15.1%)	2 (4.5%)	0.68
Haemodialysis	49 (12.5%)	0	< 0.001
Chronic liver disease	14 (3.6%)	0	< 0.001
Congestive cardiac failure	8 (2.0%)	1 (2.3%)	0.99
Malignancy	5 (1.3%)	4 (9.1%)	0.58
HTLV-1**	116/270 (43.0%)	nt	
Strongyloidiasis ^{††}	73/206 (35.4%)	nt	

BSI = bloodstream infection. HTLV-1 = human T-cell lymphotropic virus type 1. nt = not tested.

* Residence outside Alice Springs rural area. † Log-rank test. ‡ Among patients who survived > 30 days after initial BSI (Indigenous [n = 550]; non-Indigenous [n = 60]). § Comorbidities recorded for patients admitted over the period 2003-2005. ¶ Binomial test of proportions. ** Positive Western blot test for HTLV-1. †† Positive serology for Strongyloides stercoralis.

Non-communicable diseases, infection and survival in a retrospective cohort of Indigenous and non-Indigenous adults in central Australia

Lloyd Einsiedel, Liselle Fernandes, Sheela Joseph, et al.

BMJ Open 2013 3:

Table 1 Demographics and comorbidities for indigenous and non-indigenous BSI patients 2003–2007

	Indigenous (n=558)	Non-indigenous (n=55)	p Value for difference	
Age (years) (±SD)	44.7±15.2	57.5±21.1	<0.001	
Gender (M/F) (%)	234/324 (42/58)	31/23 (57/43)	0.03	
Mortality				
28 days	62 (11.1)	7 (12.7)	0.72	
All deaths	145 (26.0)	15 (27.3)	0.84	
Age of death (years)	47±15	68±21	<0.001	
Major BSI pathogens ^{††}	1029	110		
Enterobacteriaceae	370 (36.0)	38 (34.5)	0.56	
Escherichia coli	246 (66.5)	28 (73.7)	0.37	
Klebsiella pneumoniae	57 (15.4)	2 (5.3)	0.09	
Staphylococcus aureus	191 (18.6)	20 (18.2)	0.83	
MRSA	53 (27.8)	1 (5.0)	<0.001	
Streptococcus pneumoniae	136 (13.2)	8 (5.9)	<0.001	
Streptococcus pyogenes	68 (6.6)	8 (7.3)	0.42	
Haemophilus influenza	22 (2.14)	0	0.14	
Enteric pathogens‡‡	29 (2.81)	1 (0.91)	0.27	

Invasive Pneumococcal Disease

Map: Notification rates for invasive pneumococcal disease, Australia, 2008, by Statistical Division of residence



CDI

Staphylococcus aureus bacteraemia



Figure 1 Annual incidence rates of *Staphylococcus aureus* bacteraemia, 2003–2006, among indigenous and non-indigenous residents of Central Australia. Annual incidence rates of *S. aureus* bacteraemia for the indigenous and non-indigenous resident population of the Alice Springs Region using the Australian Bureau of Statistics annual estimated resident population as the denominator. (—) indigenous; (....) non-indigenous.

¹ Hewagama S, IMJ 2012

Human T Cell Lymphotropic Virus 1 (HTLV-1)

- Cell associated retrovirus
 - Vertical transmission
 - Breast feeding
 - Sexual
 - Blood products
- Endemic to certain areas:
 - Japan
 - Sub-Saharan Africa
 - Caribbean
 - South America
 - Central Australia
 - 31% of dialysis patients in Alice Springs in 2011

HTLV-1

	Epidemiological evidence			Biological evidence		
	Case reports or series	Case control studies	Cohort studies	HTLV-1 in lesions	Animal model	
Inflammatory syndromes						
HAM/TSP	Yes	Yes	Yes	Yes	Yes	
Uveitis	Yes	Yes		Yes	Yes	
Arthropathy	Yes	Yes		Yes	Yes	
Sjögren's syndrome	Yes			Yes	Yes	
Polymyositis	Yes			Yes	Yes	
Thyroiditis	Yes			Yes		
Pneumopathy	Yes					
T lymphocyte alveolitis	Yes	••				
Malignant diseases						
ATL	Yes	Yes	Yes	Yes	Yes	
Cutaneous T-cell lymphoma	Yes			Yes		
Infectious complications						
Strongyloides stercoralis	Yes	Yes	Yes			
Crusted scabies	Yes					
Infective dermatitis	Yes			••	••	
Tuberculosis	Yes	Yes				
Leprosy	Yes	Yes				

HAM/TSP=HTLV-1-associated myelopathy/tropical spastic paraparesis. ATL=adult T-cell leukaemia/lymphoma. ..=unknown. References 1, 5, 55, 103, 105–126. See webappendix for supplemental list of references, and an indication of which studies relate to each association and basis for association.

Table 2: Diseases reported in association with HTLV-1 and basis for this association

Bronchiectasis Is Associated With Human T-Lymphotropic Virus 1 Infection in an Indigenous Australian Population

Lloyd Einsiedel,^{1,2} Liselle Fernandes,² Tim Spelman,² Daniel Steinfort,¹ and Eduardo Gotuzzo³

- 52/89 (58.4%) of Indigenous adults with bronchiectasis were HTLV-1 positive
- HTLV-1 associated with:
 - ↑ number of bronchiectatic lobes (OR 1.51; 95% CI 1.03-2.20)
 - ground glass opacities on HRCT (OR 8.54; 95% CI 1.04-70.03)
 - **↑** disease specific mortality (OR 5.78; 95% Cl 1.17-26.75)
- Presence of skin lesions was associated with mortality (OR 6.77; 95% CI 1.46-31.34)
- Mortality 34.2% median age 42.5

Clinical Infectious Diseases 2012;54(1):43–50

INFECTIONS IN RENAL TRANSPLANT PATIENTS

Renal Transplant Outcomes in Central Australia: 2000-2011 Retrospective Review

- 42 transplants in 38 recipients
 - i.e. 4 re-transplants
 - 26 Indigenous, 16 non-Indigenous
- All non-Indigenous patients from/in Alice Springs
- 6 of Indigenous patients in Alice Springs, 10 in Tennant Creek
- 21 recipients alive with functioning graft & 12 alive with a failed graft
- 8 deaths in patients with a functioning graft (7 Indigenous)

Cause of Death



SEPSIS

CARDIAC ARREST POST OPUNCLEAR

CANCER

The past 10 years

- Functioning graft
 - Enterococcal bacteraemia
 - CMV Colitis
 - BK viraemia
 - Cutaenous CMV
- Failed graft
 - Rhizoctonia bataticola skin infection, Cryptosporidial diarrhoea, disseminated microsporidiosis, Nocardia bacteraemia
 - Cryptosporidial diarrhoea
- Deaths
 - ESBL UTI, CMV viraemia, BK viraemia
 - Apophysomyces elegans skin infection (Zygomycete)
 - E.coli urosepsis
 - Fungal pneumonia

Mr W.E

- 59yo Indigenous male from Tennant Creek
- ESRF presumed secondary to diabetic nephropathy
 - PHx: T2DM, dyslipidaemia, past *M.bovis* lung infection, cleared Hepatitis B, (IHD)
- Cadaveric renal Tx in 2010
 - Acute vascular rejection day 7 Rx: Atgam
- Cutaneous fungal infection Rhizoctonia bataticola
 - Rx: liposomal Amphotericin B then voriconazole
- Cryptosporidial diarrhoea
 - Rx: 2/52 nitazoxanide
- Scabies

2011 – Disseminated Microsporidiosis

- presented with a LRTI
- Deteriorated despite broad spectrum antibiotics
- Intubated in ICU
- Bronchoscopy:
 - microsporidial spores
- Renal biopsy:
 - extensive granulomatous inflammation with spores of microsporidia
- Started on albendazole Rx
- Transferred to RAH
 - Microsporidia seen again on BAL (*E.intestinalis*)
 - CMV colitis on rectal biospy
 - C.difficile diarrhoea



Progress

- October 2011 Nocardia bacteraemia
 - Presented with fevers & new lung infiltrate on CXR
 - Complicated by seizures (CT brain NAD)
 - Rx: meropenem & bactrim subsequently changed to imipenem and ciprofloxacin, then ciprofloxacin maintenance Rx for 1 year
- Decision to withdraw immunosuppression and return to haemodialysis

Microsporidiosis

- Intestinal disease
 - Usually Enterocytozoon bieneusi
- Disseminated disease
 - Usually *Encephalitozoon* species
- Often associated with HIV infected individuals
- A number of case reports of disseminated disease in SOT patients; the majority in renal transplantation
 - Most E.cunuculi infection
 - 1 other case report of *E.intestinalis* disseminated infection (also a renal transplant patient)¹
- E.intestinalis infection usually associated with gastrointestinal disease and is associated with poor socioeconomic living conditions²

¹ Latib MA, Transpl Int 2001 ² Enriquez FJ, CID 1998

Cryptosporidiosis

- Intestinal protozoa
- Faecal-oral spread, requires a low infectious inoculum
 - often transmission from swimming pools
- Oocyst survives for long periods in the environment & is relatively resistant to chlorine
- Often infects children <5 years old or immunocompromised patients
- Immunocompetent pts:
 - Self-limiting diarrhoea but 40% have recurrent symptoms
 - In developing countries can cause more persistent diarrhoea (predictor of malnutrition)
- HIV patients
 - CD4 >150 usually self-limited diarrhoea but more frequent relapse
 - CD4 <150 persistent diarrhoea & disseminated disease
 - Respiratory tract & biliary tree

Cryptosporidiosis in the NT

- In 2012
 - 243 notifications
 - 75% higher than the 5 year mean
 - 156% higher than 2011
 - 74% were between 0-4 years old
 - Overall:116 cases per 100,000 Indigenous vs 90 cases per 100,000 non-Indigenous
 - Rate ratio 1.3 (p=0.04)
 - Non-significant in the 0-4 year olds only

Figure 4. Cryptosporidiosis rates, by age and Indigenous status, Northern Territory, 2012



ANY QUESTIONS?