





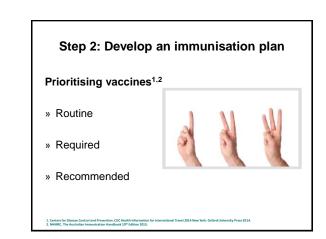
Who are high-risk travellers?

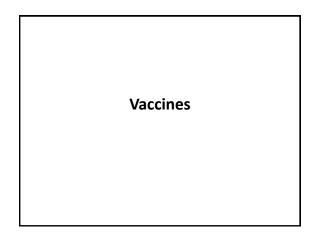
High risk:

- Chronic illnesses
- Immunocompromised
- · Young children
- · Elderly
- Pregnant
- Travellers or expatriates travelling to developing countries for an extended period
- Travellers visiting friends and relatives (VFRs)

Consider consulting a travel medicine expert for highrisk travellers

 Centers for Disease Control and Prevention. CDC Health Information for International Travel 2014 New York: Oxford 2. Yung A et al. Manual of travel medicine, 3st edition. IP Communications, 2011





Update routine vaccines

» With what <u>routine</u> immunisations should this person be up-to-date?

» MEASLES!!!

» What <u>additional</u> immunisations should this patient be up-to-date with, in view of his/her medical history and individual factors?



Required travel vaccines

· Yellow fever vaccination

- is an entry requirement in some countries
- for those who arrive from or transit through yellow fever endemic countries
- Other examples of required vaccinations
 - Vaccination requirements for Hajj & Umrah pilgrims (updated annually)
 - Influenza
 - Meningococcal ACW₁₃₅Y vaccine
 Polio
 - Pollo
 - Polio booster endemic countries can have entry and exit requirements

Recommended travel vaccines

Recommendations should be tailored to the individual traveller –

"This traveller, this trip, this time"

- · Consider:
 - · Potential health risks based on traveller and trip factors
 - · Efficacy and safety of vaccines
 - Likelihood of repeated travel
 - Time until departure
 - Vaccine interactions

1. Centers for Disease Control and Prevention. CDC Health Information for International Travel 2014 New York: Oxford University Press 2014. 2. NHMRC. The Australian Immunisation Handbook 10th Edition 2013. 3. Yung A et al. Manual of travel medicine, 3rd edition. IP Communications, 2011

Standard Vaccines

- No lower age limit
 Polio, HBV
- Lower age limit 6 weeks

 DTP, Hib (T & OMP), MenCCV, 4CMenB, PCV, Rotavirus (both)
- Lower age limit 9 months

 MMR, Varilrix
- Lower age limit 12 months

 Varivax

Standard Vaccines

- Minimum interval between all doses: 4 weeks
 DTP, influenza, polio, PCV, rotavirus
- Minimum interval between D1 and D2: 4 weeks
 All the standard childhood vaccines (except MenCCV)
- Minimum interval between doses: 8 weeks

 MenCCV (all)
 - HBV (D2-3; D3-4)
- Variations from the above
 - Hib (PRP-T) D1/D2/D3: 4wks/4wks/52wks
 - Hib (PRP-OMP) D1/D2: 4wks/52wks
 - 4CMenB interval varies with age &/or insufficient data

Accelerated Schedules

Vaccine	Routine Age (months)	Accelerated
DTP	2, 4, 6, 48	6, 10, 14 weeks & 4 years
Polio	2, 4, 6, 48	0, 1, 2 (+/- 3) months, 4 years
Hib	2, 4, 12	6, 10, 14 weeks but 2, 3, 4, 12 months preferred
MMR	12, 48	9 months, 4 years
Hepatitis B	2, 4, 6 or 12	0, 1, 2 months, 6-12 months
PCV	2, 4, 6	6, 10, 14 weeks

Travel Vaccines

- No lower age limit
 BCG, Rabies
- Lower age limit 2 months*

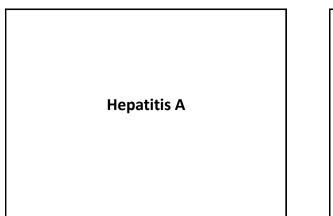
 MenACWY conjugate, Japanese Encephalitis
- Lower age limit 6 months

 Yellow fever
- Lower age limit 12 months

 Hepatitis A
- Lower age limit 24 months

 Cholera, typhoid

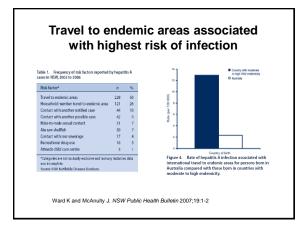
*Varies from product information

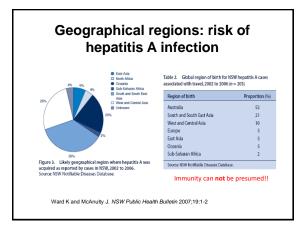


Hepatitis A

- "Hepatitis A is the Most Common Vaccine-Preventable Illness in Travellers"*
- Highly immunogenic
- Virtually 100% seroconversion
- Risk areas are everywhere except:
 - Nth America
 - ANZ
 - Western Europe

*Influenza and TD probably are the most common





Hepatitis A vaccines

All inactivated vaccines (Avaxim, Havrix, VAQTA)

- Safe, highly immunogenic
- Primary course 1 dose + 2nd dose at 6–12 months for long term protection – almost 100% sero-conversion
- Ideally give 1st dose at least 2 weeks prior to anticipated exposure, but 1st dose can be given any time up to (even after) anticipated exposure (?7 days) i.e. "last minute traveller"
- After 2 doses, further doses not needed in healthy individuals

ases and vaccines 2016 . NHMRC. Australia

Hepatitis A & B

- "Hepatitis A & B are the most common Vaccine-Preventable Illnesses in Travellers"
- Most Travellers don't allow **enough time** for Hepatitis B schedule:
 - ~ 70% Travellers who see an MD do so
 - < 1 month before leaving

*Influenza and TD probably are the most common

Twinrix 0,7, 21 days

- 1 week after 3rd dose:
 - > 80 % protected HBV
 - -100 % high levels of protective anti-HAV
- But.....
 - give 4th dose @ 12 months for almost 100%
 SC (& antiHBs +++)

Typhoid

Typhoid – The Illness

Faeco-oral spread

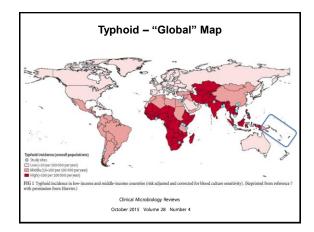
- Usually food, but waterborne outbreaks occur
- Incubation period 7-21 days (range 3-60 days)

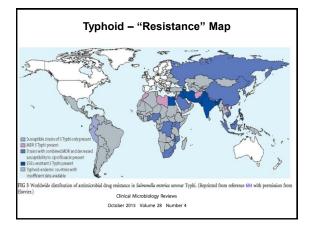
"Enteric Fever" Presentation

- Fever, headache, dry cough, myalgias, rose spots
- Usually constipated diarrhoea not common in travellers
- Preferred diagnostic sample is **blood culture**, stool negative early

Complications

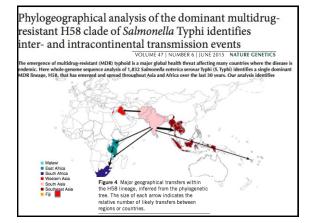
- Increase if untreated after 2 weeks, but may be the initial presentation
- Perforation (SB), massive GI bleeding (SB), seeding to foci bone, artery wall, "typhoid state" = encephalopathy/psychosis

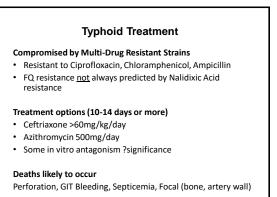




Typhoid Cases Notified in Australia

- Average around 120 150/year
 All (virtually) have travel history
- · Indian sub-continent and South East Asia most cases
 - India
 - Nepal
 - Bangladesh
 Pakistan
 - PakistanIndonesia
 - VFRs (visiting friends and relatives) high risk
 - Less likely to seek pretravel health advice
 - Greater exposure





Typhoid Vaccine: Which Travellers?

- Travellers to endemic regions, where food hygiene may be suboptimal and drinking water may not be adequately treated
 - Travellers to endemic regions to visit friends and relatives (VFR) have considerably greater risk
 - Endemicity data may be unreliable
 - Some areas of countries higher risk than others
 - Popular Oceanic (Fiji!) destinations may have MDR strains

NHMRC. The Australian Immunisation Handbook 10th Edition 2013.

Typhim Vi, Typherix, Vivaxim

- Stimulate antibodies to Vi antigen of S.typhi – Killed, IM/SC (0.5 cc), VICPS
 - Lower age limit (must be aged over 2 years)
 - 1 dose, boost at 3 years
 - 60-80% protection (70% @ 18mths, 50% @36mths)
- Vivaxim for vaccine "virgins" vs TwinRix (for <u>HepA</u>)

 depends on itinerary/risk/time for course
- Injectables v. oral typhoid vaccine
 - Compliance
 - Handling issues
 - No interactions

Oral Typhoid Vaccine (OTV) (Vivotif)

- Oral live attenuated Ty21a vaccine: Vivotif Oral
 - 3 (or 4) doses swallowed (whole) on D1, D3, D5, (D7)
 - Contraindicated if immunesuppressed
 - Can have simultaneously with mefloquine, Malarone
 - Antibiotics avoid 7 days pre-D1 to at least 3 days post-D5(D7)
 Postationa Minimum Alignment (but human and built Conserve)
 - Protection =Vi injectables (but lower age limit = 6 years)
 Duration of protection: 3 doses (1-3 years), 4 doses (5-7 years)
 - Booster (full course): 1 year or 3 years or 5-7 years (USA/Canada)
- Salmonella paratyphoid B protection?
 - Chile & Israel OTV studies not replicated Indonesia
 - Immunological investigation 2012 no explanation ?true effect*
 - No data for other S.paratyphi (A, C)

*Wahid R et al CVI 2012

Cholera

Cholera

Acute diarrhoeal disease

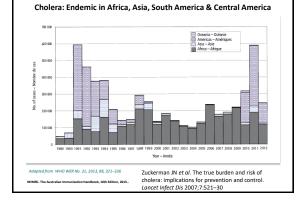
- Enterotoxin-producing Vibrio cholerae
- Serogroups O1 & O139
- In food or water contaminated with Vibrio cholerae

Not everyone is sick

• ~ 75% asymptomatic / ~ 25% symptomatic

But if symptomatic

- ~ 20% > profuse watery diarrhoea > severe dehydration
- · Severe cholera can be rapidly fatal if left untreated
- Focus of treatment is rehydration



Cholera: Who to Vaccinate?

- If considerable risk of exposure to/of acquiring cholera
 - e.g. Humanitarian disaster workers
- · If at risk of severe or complicated diarrhoeal disease
 - e.g. Poorly controlled diabetes, inflammatory bowel disease
- If at risk of acquiring diarrhoeal disease
 e.g. Patients with achlorhydria

NHMRC. The Australian Immunisation Handbook 10th Edition 2013.

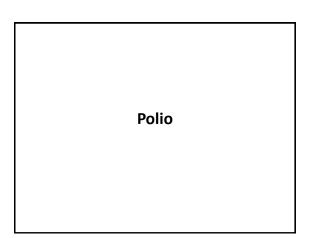
Dukoral – For Cholera

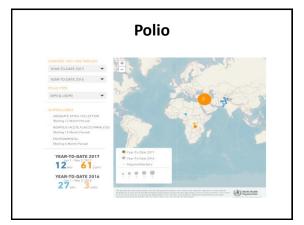
- Oral inactivated vaccine
 For immunisation against cholera caused by serogroup O1 Vibrio cholerae (not against O139)
- For adults & children ≥2 years of age
 Demonstrated 85% protective efficacy against cholera at 6 months after primary course
- Administer doses at an interval of 1-6 weeks:
 Children 2 6 years: 3 doses
 Adults > 6 years: 2 doses
 - If > 6 weeks elapse between doses primary course must start again
- Generally well tolerated
 Occasional GI symptoms

Dukoral - Boosters

- Protection from ~1 week after primary immunisation completed

 <u>Ensure</u> 2nd dose is taken at least 1 week before departure
- Protection lasts 2 years <u>but</u> booster doses my be required every 3-6 months for persons at continuous high risk of contracting cholera
 - Booster (for <u>Cholera</u> protection): 1 dose every 3 months if at continuous significant risk
- Booster Doses for <u>ETEC TD</u> protection:
 - After primary course of 2 doses, a follow-up booster (1 dose) given at <u>any</u> <u>time within 5 years</u> from completion of the primary course (or within 5 years after any booster doses eg for cholera protection) <u>should be</u> <u>sufficient for renewed protection against ETEC.</u>
 - If >5 years has passed since the primary course or last booster dose, the full primary course (2 doses at least 1 week apart) should be given





Polio

- Only 2 countries reporting wild poliovirus (WPV) infection in 2017

 Afghanistan, Pakistan
- Recent WPV cases, other than the above – Nigeria (2016)
- Recent cases of vaccine derived poliovirus (VDPV) infection
 DR Congo, Lao PDR, Syria
- Circulating WPV and VDPV pose risk to unimmunised travellers
 - Detect by environmental sampling may be no detected cases
 Primary immunisation v. booster
 - Polio mass vaccination campaigns implemented
 - Evidence of recent poliovirus vaccination as a requirement for travellers entering and leaving endemic countries
 - Travel to countries bordering the above?

Japanese Encephalitis

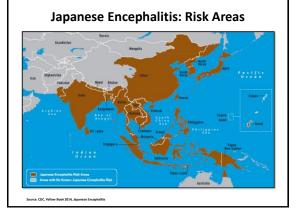
JE – The Disease

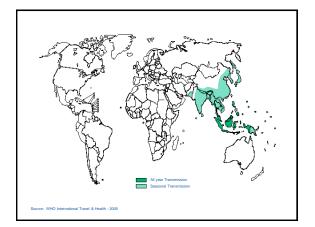
- Flavivirus

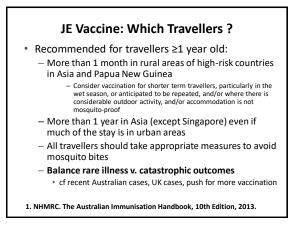
 Mosquito-borne
- Perspective
 Clinical illness very rare in travellers
- Prevention
 - Vaccine efficacy good
 - Immunity
 - Mosquito avoidance

JE – The Disease

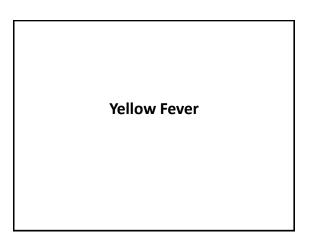
- Many cases asymptomatic
 - Very rare in travellers
 - Very rare in expats (but OH&S)
- Annual global incidence 15,000 / year
 - Case fatality: 10 30%
 - Long term disability: 30%
 - Neurological sequelae: 40 80%
 - Complete recovery: 10 15%







		accine using ye Abs @ 14 days	llow fever virus backbo)	ne
Table 4.8.1: Recom	mended doses o	f JE vaccines		
Age of vaccine recipient	Vaccine	Number of doses	Booster	Jespect
≥2 to <9 months	JEspect	2 doses* (28 days apart)	Not required Refer to Note 2	 Vero cell cultured, inactivated vaccine
≥9 months to <18 years	Imojev	1 dose	1-2 years after primary dose	 2 doses IM: day 0,
	JEspect Refer to Note 1	2 doses* (28 days apart)	Not required Refer to Note 2	28 (or 0,7)
≥18 years	Imojev	1 dose	Not required	
	JEspect	2 doses (28 days apart')	1-2 years after primary dose	
4.8.6 Dosage and ads Note 2: Currently there is JEspect in children w * Each dose of JEspect	ninistration above). very limited evidence ho received JEspect in infants and childr	e available to inform re as primary immunisatio m aged ≥2 months to <	commendations regarding the need as n. 3 years is 0.25 mL.	ot available or is contraindicated (refer to ad appropriate time interval for a booster of readults who are at imminent risk of



Yellow Fever

- Globally 200,000 cases/year – Outbreaks in 2016-2017 (Angola, **Brasil**, others)
- Mosquito transmitted
 - Flavivirus
 - Monkey reservoir
 - Jungle urban
- Africa, South & Central America, T&T
- Not Asia

Yellow Fever Vaccine

- Self-Protection "active" areas
- International regulations
 - WHO & CDC advise for return from "active" areas
 - Many countries insist on the "old endemic zone"
 - "Receptive" countries cautious
 - WHO now advises 1 dose = lifelong protection
 - No boosters required





Contraindications

- Allergy
 - egg, polymixin, neomycin
- < 6, 9 or 12 (?4) months of age
- Cancer, immunosuppressed,
- Pregnant
- HIV (< 200 CD4)
- Thymic Disorders (ever)
- Risk of getting YF vs. Risk of AE from YFV

Yellow Fever - Vaccine Deaths

- Non-allergic causes related to complications of the (usual) YFV strain viraemia
 - Viscerotropism (YF-AVD or MSOF)
 - Neurotropism (YF-AND)
- YFV Serious AE risk increases with age
 - 7.5 /100,000 aged 70 years or more
 - 4 /100,000 aged 60-69 years age
 - Immunocompetent risk is age related
 - Not seen with booster doses

Yellow Fever Vaccine Risk

• YFV - Risk of YF v. Risk of YFV

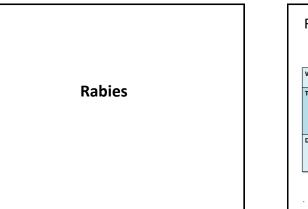
- Get up to date data on active transmission (lags)
- Change itinerary or even go somewhere else

• Exemption/Waiver Certificate

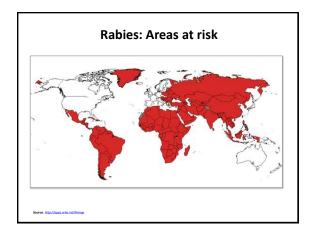
- Age > 60 years & Low Risk Travel
- But if risk significant may still need YFV

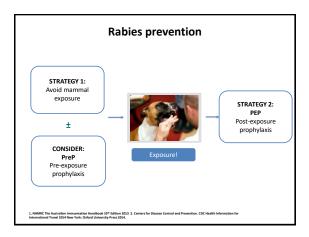
• Anaphylaxis rate: 1/130,000

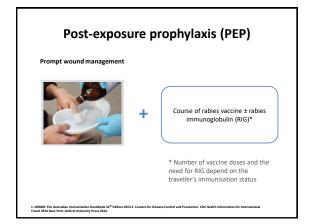
		-		· · · ·	
Agent/ Condition	Mode of Action	Infections®	TB Screen	Live* Vaccines	Killed# Vaccines
TNF Inhibitors	anti-TNF	Bacterial, Granulomatous, Intracellular, Viral	Yes	No	Yes
IL-1 Inhibitors	anti-IL1	Bacterial, Granulomatous, Intracellular, Viral	Yes	No	Yes
Rituximab	CD20, B cells	Bacterial, HBV, Other viral	?No	No	Yes
Abatacept	CD86,80,28T cells	Bacterial, Granulomatous, Intracellular, Viral	Yes	No	Yes
Newer	Various	??	22	No	Yes

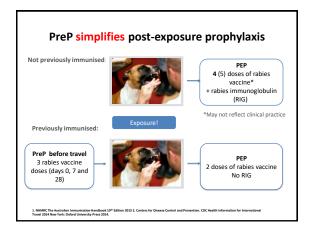


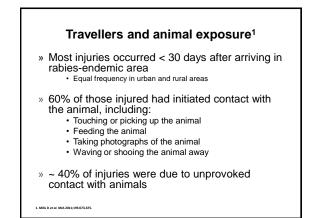
Rabies: Overview		
What is it?	 A viral infection caused by rabies virus belonging to genus Lyssavirus¹ 	
Transmission	 Usually through bite of infected animals Occasionally through scratches, licking of open wounds Rarely through tissue transplants² 	
Disease course	Incubation period is highly variable ¹ Effective and prompt post-exposure treatment can prevent disease Usually fatal once symptoms develop ¹	
1. NHMRC. The Australian Imp	nunitation Handbook 30° Edition 2011. I. Vara NM et al. JAMA 2011.130(4):934-607.	

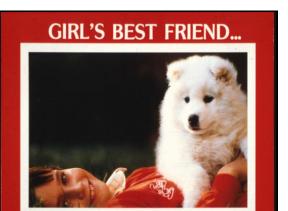














Rabies: Which travellers should receive PreP?

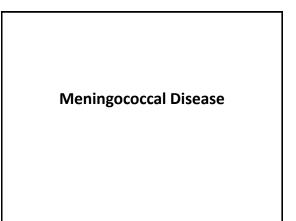
- » Travellers spending time in rabies-enzootic areas. Risk assessment should take into consideration:
 > Potential interaction with terrestrial mammals and bats
 - > Access to emergency medical attention
- » Consider PreP for **younger children** as they are at higher risk of exposure, especially head/neck
- » Persons working with terrestrial mammals and bats in rabies-enzootic areas

1. NHMRC. The Australian Immunisation Handbook 10th Edition 2013.

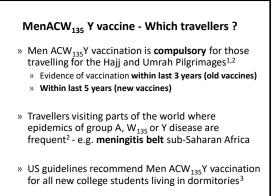
Rabies: Advice for Travellers

- » Avoid mammal bites and scratches1
 - Do not allow young children to play with or feed animals
 Do not pat or feed monkeys, even in popular areas such as
 - temples/markets
 - Avoid contact with stray dogs/cats
 - Bali is no longer rabies free cases occur near tourist areas
- $\,\,{}^{\,\,}$ M now what to do in the event of exposure 1,2
 - Immediately and thoroughly wash wound with copious amount of soap and water, then apply antiseptic e.g. povodine-iodine
 - Seek medical attention as soon as possible, preferably within 48 hours
- » If PEP was commenced overseas, request PEP certificate1

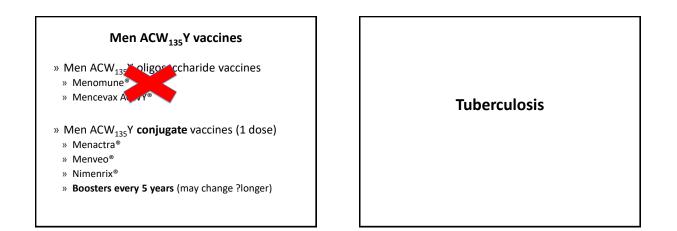
1. NHMRC. The Australian Immunisation Handbook 10th Edition 2013. 2. Centers for Disease Control and Prevention. CDC Health Information for International Travel 2014 New York: Oxford University Press 2014.



What is it?	 Bacterial infection caused by N. meningitidis¹ Serogroups A,B,C, W₁₃₅ and Y cause majority of meningococca disease globally¹
Transmission	Via close contact with respiratory secretions or saliva ² Risk factors include: ¹ Living in crowded conditions Recent illness Multiple kissing partners
Disease course	 May cause meningitis, septicaemia or a combination of both. May be rapidly fatal^{1,3} Of those who survive, ~10–20% are left with permanent sequelae e.g. brain damage, hearing loss^{1,3}

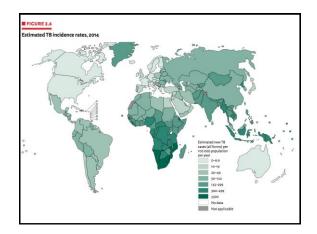


 Saudi Arabian Ministry of Health. http://www.bajinformation.com/main/p1001.htm Accessed May 7 2013 2. NHMRC The Australian Immunisati Handbook 10th Edition 2013 3. Cohn A et al. MMWR Recomm Rep. 2013 Mar 22; 62(RH-2):1-28



Tuberculosis

- Risk for travellers underestimated
 - Higher for VFRs
 - Exposure does not have to be "months"
 - MDR-TB a concern
 - Mycobacterium bovis from food
- Age < 5 years more risk for severe disease & death
 Miliary TB
 - Severe TB meningitis



Tuberculosis

BCG vaccine

- Live vaccine (ID: 0.1cc; half dose age < 12 months)
- P/E: 50% overall; 80% v miliary TB, TBM, TB death
- Children aged less than 5 years
- Stay for > "a few weeks" or > 3 months in HICs
- High incidence countries (>100/100,000 annual incidence)
- Pre-BCG screening not usually necessary
 - Unless previous history of possible exposure
 - Less data on PPD/Mantoux v Quantiferon Gold/IGRA assays in children v adults (but seem to correlate)

Remember When..?

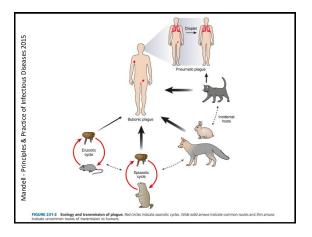
I've got a cough & feel terrible!

Richard aged 45 years

- Has just returned from Madagascar yesterday
- Regular traveller to Madagascar
- Has a lot of local friends
- Stayed with friends and "went everywhere" over 4 weeks
- Was on doxycycline malaria prophylaxis but stopped @ 2 weeks
- Rarely gets sick
- Doesn't believe in influenza vaccination "doesn't work"
- Only developed cough and fever
 Halfway home on the flight from South Africa





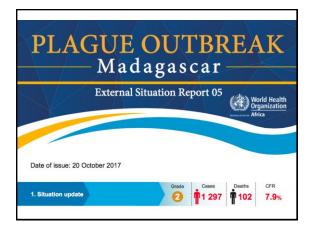


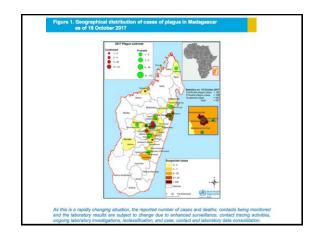
Plague – Prevention

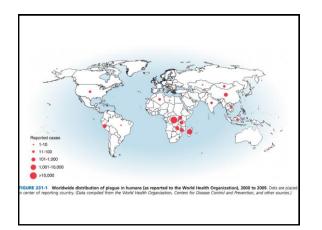
- Avoid
 - Handling or exposure to animals, dead animals
 Rats, fleas
 - Rural areas, slum areas (rats, fleas)
- Use
 - Cough protection (mask), hand hygiene, repellents
- Vaccine (old, killed) not available
- Post-exposure prophylaxis – Doxycycline (100mg bd), ciprofloxacin (500mg bd)
- Malaria chemoprophylaxis (doxycycline)

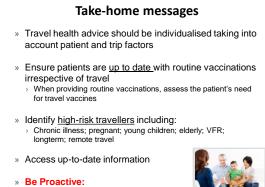
Plague – Key Points

- Bubonic
 - Lymph node enlarged, tender, may discharge
- Septicaemic
 - Sepsis, undifferentiated, may be bubo, eschar
- Pneumonic
 - Cough, very unwell, "flu", death
- Incubation Period
 - 2 7 days (24 hours for pneumonic plague)
- Pneumonic is highly contagious









» Ask patients about their travel plans

