Travel Medicine in the Age of COVID

September 8, 2023 Shingo Chihara, MD





Disclosure Statement:

No relevant financial relationships with ineligible companies to disclose.



Objectives

- 1. Familiarize with risk for travelers in various circumstances and how to discuss with the traveler
- 2. Develop recommendations for traveler's pre-trip planning and preparations
- Identify resources for country-specific requirements with regard to COVID testing, how to access COVID testing and treatment in the US and abroad, how to obtain appropriate medical treatment for serious illness while away from home
- 4. Consider special needs of child travelers



Travel Risk Assessment

- Itinerary
 - Countries and specific regions, including order of countries if > 1 country
 - Outbreaks at destination
 - Rural or urban destinations
- Timing
 - Season of travel
 - Time to departure
 - Trip duration



Travel Risk Assessment

Reason for travel

- Adoption
- Adventure
- Business
- Education or research
- Medical tourism (seeking health care)
- Pilgrimage
- Tourism



- Visiting friends and relatives
- Volunteer, missionary or aid work

Travel Risk Assessment

Travel style

- Accommodations (ex. Camping/tent, dormitory, guest house, hostel/budget hotel, local home or host family, tourist/luxury hotel)
- "Adventurous" eating
- Level of hygiene at destination
- Modes of transportation
- Traveler risk tolerance
- Travel with children



Immunizations



Vaccinations

Routine vaccination

- Tdap
- MMR
- Influenza
- Hepatitis A and B
- Polio
- Meningococcal
- HPV



- VZV
- Pneumococcal

Estimated incidence per month of vaccine preventable diseases in lower-income countries among nonimmune Western travelers

Influenza: 1% (1/100)

Animal bite with rabies risk, latent tuberculosis infection: 0.1-1% (1/1000-1/100)

Typhoid in South Asia, Measles, Pertussis, Hepatitis A: 0.01-0.1% (1/1000-1/10,000)

Tick borne encephalitis in rural Baltics, hepatitis B, typhoid in Africa and south America, Active tuberculosis 0.001-0.01% (1/10,000-1/100,000)

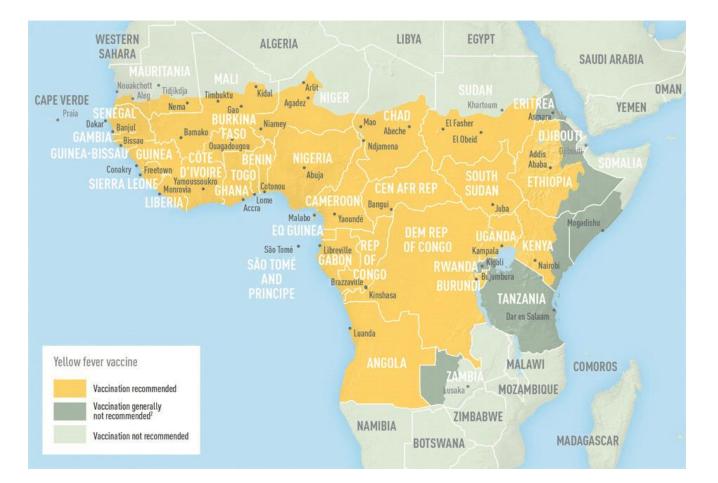
Typhoid in Caribbean and Central America 0.0001-0.001% (1/100,000-1/million) Japanese encephalitis (1/million)

Meningococcal disease, poliomyelitis, cholera, yellow fever (1 in 1-10 million)

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Yellow fever







https://www.cdc.gov/yellowfever/maps/index.html

Yellow Fever Vaccine

- Entry requirements for proof of vaccination under the International Health Regulations
- Recommended for all travelers \geq 9 months old going to areas with YF risk
- 1 dose of YF vaccine provides long-lasting protection
 - Booster
 - Pregnant when receiving initial dose
 - Hematopoietic stem cell transplant after receiving vaccine
 - Infected with HIV



Yellow Fever Card

INTERNATIONAL CERTIFICATE OF VACCINATION OR PROPHYLAXIS Certificat international de vaccination ou de prophylaxie This is to certify that Nous certifions que Image: Angrey Doe Certificat international de vaccination ou de prophylaxie Image: Certificat international de vaccination ou de prophylaxie Insist to certify that Certificat international de vaccination ou de prophylaxie Image: Certificat international de vaccination ou de prophylaxie Image: Certificat international de vaccination ou de prophylaxie Insist to certify that Certification grave Image: Certificat international de vaccination ou de prophylaxie al date indiquée Image: Certificat international de vaccination ou de prophylaxie al date indiquée Image: Certificat international de vaccination ou de prophylaxie al date indiquée Image: Certificat international de vaccination ou de prophylaxie al date indiquée Image: Certificat international de vaccination ou de vaccination								
Vaccine or prophylaxis Vaccin ou agent prophylactique	Date	Signature and professional status of supervising clinician Signature et titre du professionet de santé responsable	Manufacturer and batch no. of vaccine or prophylaxis Fabricant du vaccin ou de l'agent prophylactique et numéro du lot	Certificate valid from: until: Certificat valable à partir du : jusqu'au :			al stamp of the administering center Gachet officiel du centre habilité	
(+) Yellow Fever	5) 15 June 2018	6 John M. Smith, MD	Batch (or lot) #	 June 25 June 2018; life of person vaccinated 			6	



Yellow fever vaccine-associated neurologic disease

- Acute disseminated encephalomyelitis, Guillain-Barre syndrome, meningoencephalitis or cranial nerve palsies
- Onset 2-56 days following vaccination
- Rarely fatal
- 0.8 per 100,000 doses administered (2.2 per 100,000 doses administered in age ≥60 years



Yellow fever vaccine-associated viscerotropic disease

- Severe illness similar to yellow fever disease
- Multiorgan failure with case fatality rate of 48%
- Median onset 4 days after vaccination (1-18 days)
- 0.3 cases per 100,000 doses of vaccine
- 1.2 cases per 100,000 doses for age ≥ 60



Yellow fever vaccine contraindications

- Age \leq 6 months
- Allergy to vaccine component (chicken protein, eggs, egg products, gelatin)
- HIV infection (symptomatic) or CD4 <200/mL (or <15% of total lymphocytes)
- Primary immunodeficiencies
- Immunosuppressive and immunomodulatory therapies
- Malignant neoplasms
- Thymus disorder associated with abnormal immune cell function
- Transplantation



Yellow fever vaccine precautions

- Age 6-8 months
- Age \geq 60 years
- Breastfeeding
- HIV infection (asymptomatic) and CD4 T lymphocyte counts 200-499/mL (or 15-24% of total lymphocytes)
- Pregnancy



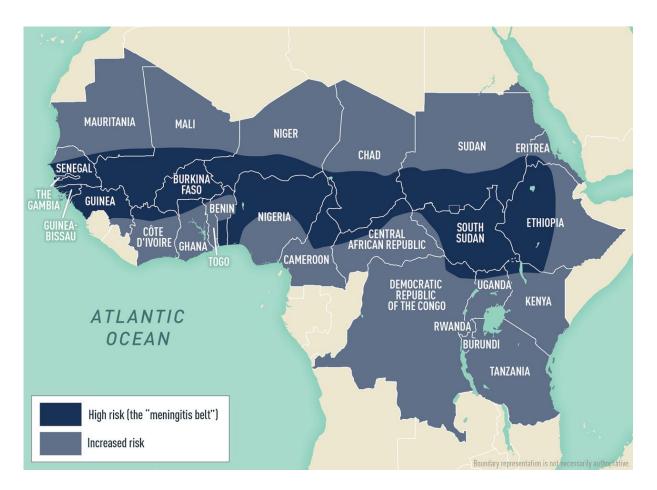
Medical Contraindication to Vaccination

MEDICAL CONTRAINDICATION TO VACCINATION	
Contre-indication médicale à la vaccination	
This is to certify that immunization against Je soussigné(e) certifie que la vaccination contre	
	for
(Name of disease – Nom de la maladie)	pour
	is medically
(Name of traveler – Nom du voyageur)	est médicalement
contraindicated because of the following conditions: contre-indiquée pour les raisons suivantes:	
	(Signature and address of physician) (Signature et adresse du médecin)



https://wwwnc.cdc.gov/travel/yellowbook/2024/infections-diseases/yellow-fever

Meningococcal vaccine



- Most risk during dry season: December to June
- Quadrivalent conjugate vaccine
- Saudi Arabia require polysaccharide vaccine for travelers making the Umrah or Hajj pilgrimage 10 days and ≤ 3 years before arrival



https://wwwnc.cdc.gov/travel/yellowbook/2024/infections-diseases/meningococcal-disease

Typhoid vaccine

- Oral, live attenuated vaccine for age 6 and above, every 5 years if needed
 - 1 capsule every other day for 4 doses
 - Complete 1 week before travel
 - Refrigerate capsules
 - Complete 1 week before travel
 - No antibiotics for 72 hours after completion
- IM polysaccharide vaccine for age 2 and older evey 2 years if needed
- 50-80% effective



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Japanese encephalitis

Mosquito borne illness

<1 case per million travelers

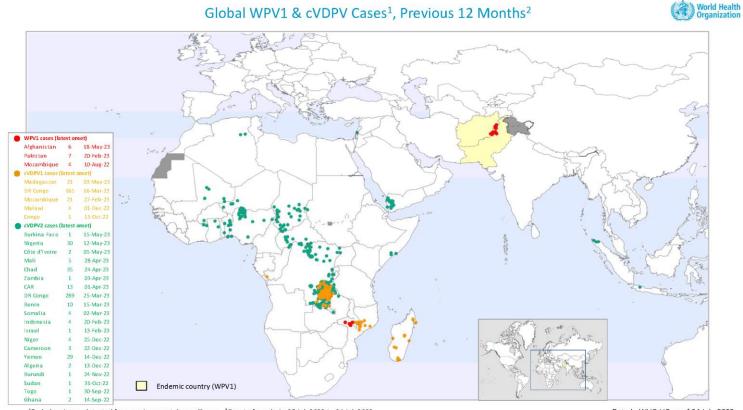
2 shot series: Day 0, 7-28 days for 18-64 years Booster considered after 1 year of vaccine





https://www.cdc.gov/japaneseencephalitis/maps/index.html

Polio



¹Excludes viruses detected from environmental surveillance; ²Onset of paralysis: 05 Jul. 2022 to 04 Jul. 2023

Data in WHO HQ as of 04 Jul. 2023



https://polioeradication.org/polio-today/polio-now/



Most common vaccine-preventable infections acquired during travel

2 shot series: 0, >6 months

Boosters not recommended





Fatal, acute progressive encephalomyelitis

All mammals are susceptible: Mainly dogs and bats

Vaccine schedule

Preexposure prophylaxis: 0, 7 days (no longer 21-28 days for travelers)

If bitten or scratched, clean the wound with copious amounts of soap and water, povidone

iodine or other products with virucidal activity.

Preexposure vaccine: Vaccines at Day 0 and 3

No preexposure vaccine:

Rabies immunoglobulins and vaccines at (0, 3, 7, 14) 28 if immunocompromised



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Tick Borne Encephalitis

Western and northern Europe, extending to northern and eastern Asia

Spring and summer months

Indications

- Adventurous eaters
- Expatriates living in endemic area
- Travelers participating in outdoor
 - activities in forested areas





Tick Borne Encephalitis Vaccine

Approved in August 2021

3 shot series for adults 16 and older

Day 0, 14 days to 3 months, 5-12 months

4th dose could be given at least 3 years after completion of primary series



Malaria prevention



Malaria Prophylaxis

- A: Awareness of Risk
- B: Bite Prevention
- C: Chemoprophylaxis
- D: Diagnosis-early detection



Awareness of Risk

Traveler Categories at Greatest Risk for Exposure and Infection

- Children
- Long-term travelers and expatriates
- Pregnant travelers
- Travel to Sub-Saharan Africa
- No use of chemoprophylaxis
- Travelers visiting friends and relatives in areas with malaria



Locally Acquired Malaria Cases Identified in the United States

<u>Print</u>





Distributed via the CDC Health Alert Network June 26, 2023, 5:00 PM ET CDCHAN-00494

Summary

The Centers for Disease Control and Prevention (CDC) is issuing this Health Alert Network (HAN) Health Advisory to share information and notify clinicians, public health authorities, and the public about—

- 1. Identification of locally acquired malaria cases (*P. vivax*) in two U.S. states (Florida [4] and Texas [1]) within the last 2 months,
- 2. Concern for a potential rise in imported malaria cases associated with increased international travel in summer 2023, and
- 3. Need to plan for rapid access to IV artesunate, which is the first-line treatment for severe malaria in the United States.



https://emergency.cdc.gov/han/2023/han00494.asp

Ecuador: CDC Yellow Book

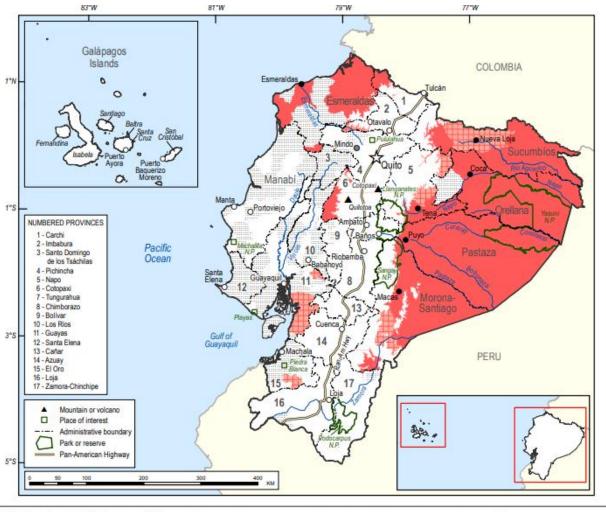
- Areas <1500 m (5000 feet) elevation in the provinces of Carchi, Cotopaxi, Esmeraldas, Morona-Santiago, Orellana, Pastaza and Sucumbios
- Rare cases <1500m (5000 feet) in all other provinces
- No malaria transmission in the cities of Guayaquil or Quito (the capital)
- No malaria transmission on the Galapagos Islands

https://wwwnc.cdc.gov/travel/yellowbook/2024/preparing/yellow-fever-vaccine-malaria-prevention-by-country/ecuador#seldyfm1118



Ecuador

Travax





- Chemoprophylaxis is recommended for all travelers.*
- Chemoprophylaxis is recommended for certain travelers; see Issues to Consider box.
- Insect precautions only are recommended* (negligible transmission is reported).
- No preventive measures are necessary (no evidence of malaria transmission exists).
- * Exceptions may apply. For more information, see Technical Explanation of Malaria Mapping.

- City where preventive recommendations are the same as the surrounding region.
 City where insect precautions only are recommended* (negligible transmission is reported).
- O City where no preventive measures are necessary (no evidence of malaria transmission exists).
- A National capital (no preventive measures are necessary).



https://www.shoreland.com/downloads/pdf/ecuador-malaria-map_sample.pdf

Bite Avoidance Measures (Bite Prevention)

Sustained-release or controlled release formulation of DEET (20-50%) or Picardin

(>20%)

- Reapply often and when outdoors
- Apply sunscreen prior to skin repellents
- Wear long sleeve shirts, long pants and socks
- Use permethrin treated clothing
- Check for ticks
- Sleep under long-lasting insecticide treated bed nets



Chemoprophylaxis

Atovaquone-proguanil

Doxycycline

Chloroquine/hydroxychloroquine

Mefloquine

Tafenoquine

Primaquine



Atovaquone-proguanil

Worldwide use

1 tab daily 1-2 days prior to travel, during travel then 7 days after travel

Highly effective and well tolerated

Take with food to minimize GI side effect

Bitter taste

Contraindicated when CrCl <30





Worldwide use

Take 100mg daily: 1-2 days prior to travel, during travel and 28 days post travel

Covers rickettsial infection and leptospirosis

Least expensive

Good for long term travelers

Side effect profile: nausea, photosensitivity, esophagitis

Not during pregnancy or breast feeding



Chloroquine and hydroxychloroquine

Only for Central America and Caribbean due to resistance

Dose weekly: 1-2 weeks pretravel, during travel and 4 weeks post travel

500mg weekly for chloroquine; 400mg weekly for hydroxychloroquine

Safe in pregnancy and with breast feeding

Avoid in liver disease, psoriasis and retinal disease



Mefloquine

Some resistant areas in Southeast Asia

Weekly dose: 1-2 weeks pretravel, during travel and 4 weeks post travel

Not for history of depression, psychosis, seizures, prolonged QT

May be used during pregnancy and breast feeding



Tafenoquine

FDA approved in July 2018

Needs G6PD testing

200mg daily x 3 days before trip, 200mg weekly then 1 dose after return

Only for >18 years old

Not during pregnancy



Traveler's diarrhea



Travelers' diarrhea

- 30-70% of travelers during 2 week period depending on destination and season of travel
- "boil it, cook it, peel it or forget it"
- Poor hygiene practices in local restaurants and underlying hygiene and sanitation infrastructure deficiencies are largest contributor
- Bacterial : 80-90%; virus: 5-15%; protozoal pathogens seen in long-term travelers

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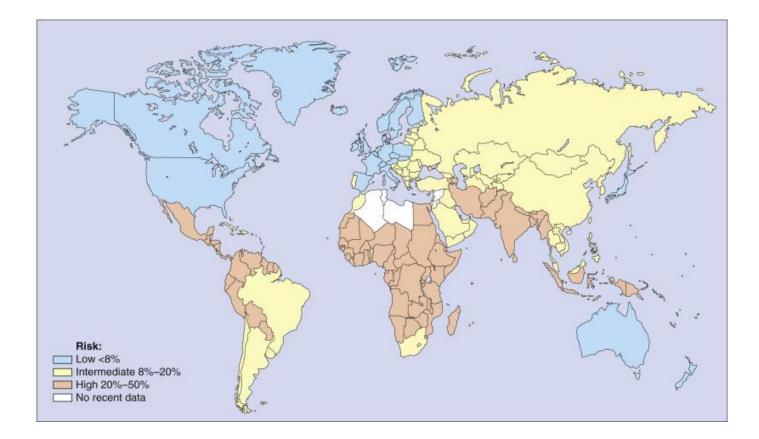


Food and Water Precautions

- Avoid salads, uncooked vegetables, raw unpeeled fruits and unpasteurized fruit juices
- Food of animal origin including meat and eggs should be cooked thoroughly
- Least risk: fully cooked foods that are served hot and foods that travelers carefully prepare themselves
- Wash hands frequently
- Avoid tap water as well as ice made from tap water



Incidence rate of travelers' diarrhea





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Treatment

- Mild diarrhea: Tolerable, not distressing, does not interfere with planned activity: Antibiotic not recommended; consider bismuth subsalicylate or loperamide
- Moderate diarrhea: Distressing or interferes with planned activities Antibiotics can be used for treatment (Azithromycin, fluoroquinolones or rifaximin); Consider loperamide for use as monotherapy

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Treatment

Severe Diarrhea: Incapacitating or completely prevents planned activities, all dysentery is considered severe

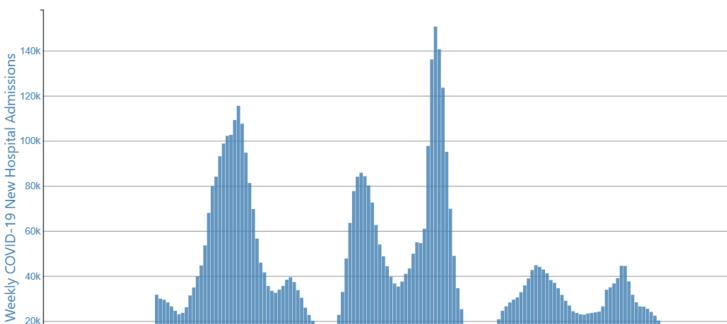
- Antibiotics advised: Azithromycin preferred; fluoroquinolones or rifaximin can be used for non-dysenteric diarrhea
- Antimotility agents not recommended as monotherapy for bloody diarrhea or diarrhea with fever







COVID-19 New Hospital Admissions by week



COVID-19 New Hospital Admissions, by Week, in The United States, Reported to CDC

Centers for Disease Control and Prevention. COVID Data Tracker. Atlanta, GA: U.S. Department of Health and Human Services, CDC; 2023, August 19. https://covid.cdc.gov/covid-data-tracker

Nov 13, '21

Jun 25. '22

Feb 4, '23

Aug 12, '23

Apr 3, '21



Jan 11, '20

Aug 22, '20

COVID Variants

Omicron EG.5 (Eris)

Most dominant subvariant (20.6% as of 3rd week of August)

One mutation in spike protein: potential for evading immunity acquired after infection or vaccination

Moderna announced preliminary clinical trial data confirm that updated vaccine for fall 2023 showed significant boost in neutralizing antibodies against EG.5. and FL1.5.1 variants (8/17/2023)

https://investors.modernatx.com/news/news-details/2023/Moderna-Clinical-Trial-Data-Confirm-Its-Updated-COVID-19-Vaccine-Generates-Robust-Immune-Response-in-Humans-Against-Widely-Circulating-Variants/default.aspx Virginia Mason Franciscan Health^{*}



Make sure vaccination is up to date

Risk factors of traveler for severe disease

Check COVID-19 status in visiting country

Bring test kits

Nirmatrevir and ritonavir?



Traveling with children



Vaccines

Vaccines: Routine and Travel vaccines

Age restrictions on vaccines

Minimum ages in the US

Yellow fever: 9 months

Hepatitis A 6 months

Typhoid: Injectable 2 years

Oral: 6 years

Influenza 6 months



Refusal of recommended travel-related vaccines among U.S. International travelers in Global TravEpiNet

J Travel Med 2017 Jan;24(1); taw075

23,768 travelers eligible for at least one vaccine from July 2012 to June 2014

25% refused one or more vaccines

Providers documented reason for refusing vaccine

- 1. Cost concerns
- 2. Safety concerns
- 3. Not concerned with the illness



Reason for refusing vaccines among travellers in the GTEN study population

Vaccines	Reason traveler refused vaccine		
	Not concerned with illness	Concerned with vaccine safety	Concerned with vaccine cost
Influenza	2851 (81%)	526(15%)	150(4%)
Meningococcal	1744 (78%)	311(14%)	177(8%)
Typhoid	1230 (73%)	171(10%)	289(17%)
Hepatitis A	1169 (73%)	245(15%)	184 (12%)
Tetanus	1140 (76%)	257 (17%)	101 (7%)
Polio	1098 (80%)	181 (13%)	88 (6%)
Rabies	3340 (78%)	421 (10%)	517 (12%)
Yellow fever	612 (67%)	225 (25%)	80 (9%)
Japanese encephalitis	460 (60%)	35 (5%)	266 (35%)



Other tips

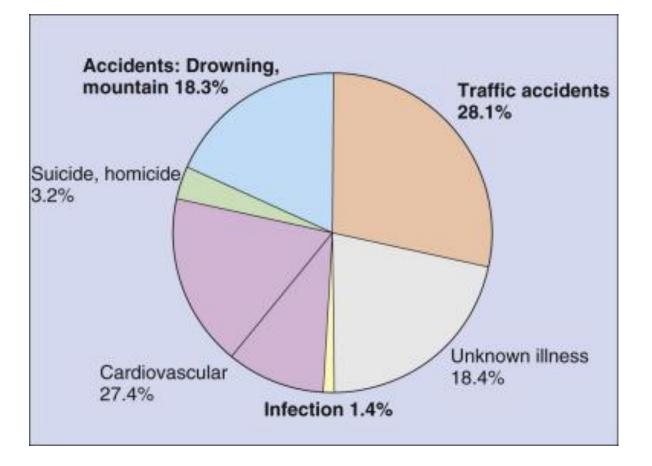
- Medications: provide liquid when possible
- Safety: bring car seats
- Medical kit: COVID test kit
 - Medical card with age, weight, medical history, allergies, blood type
 - Immunization records
 - OTC meds
 - Acetaminophen/ibuprofen, antihistamines
 - 1% hydrocortisone cream, antibacterial, antifungal ointment
 - Cough suppressant
 - Bismuth subsalicylate, loperamide



Non-infectious counseling



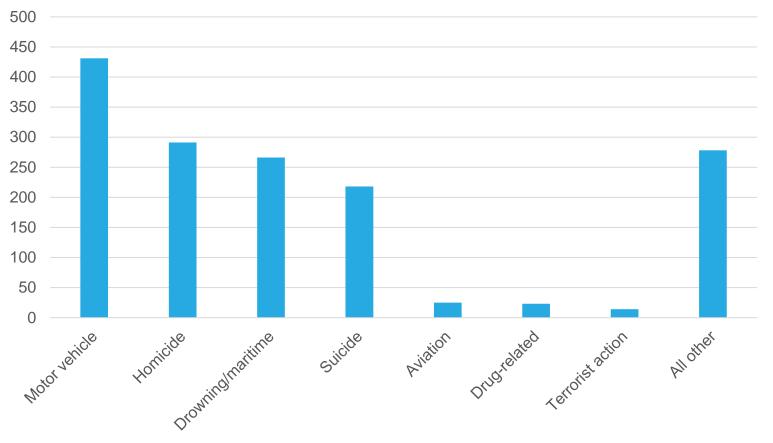
Fatalities among French abroad 2001-2004





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Leading cause of injury death for US citizens in foreign countries, 2016 & 2017



Virginia Mason Franciscan Health[®]

https://wwwnc.cdc.gov/travel/yellowbook/2024/environmental-hazards-risks/injury-and-trauma

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Typhoid in Caribbean and Central America 0.0001-0.001% (1/100,000-1/million)

Japanese encephalitis (1/million)

Rabies, meningococcal disease, poliomyelitis, cholera, yellow fever (1 in 1-10 million)

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Health Care Abroad

Medicare, Medicaid and most health insurance plans do not cover international medical care

- Travel insurance: lost baggage and trip cancellation
- Supplemental travel health and medical evacuation insurance

US Department of State (travel.state.gov/content/travel/en/international-travel/before-you-go/your-health-abroad/insurance-providers-overseas.html.)

International Association for Medical Assistance to Travelers (www.iamat.org)

US Travel Insurance Association (www.ustia.org)

American Association of Retired Persons (www.aarp.org)

https://wwwnc.cdc.gov/travel/yellowbook/2024/health-care-abroad/insurance



Take Home points

- Review vaccines: Routine and travel vaccines
- Malaria prevention with medications/insect precaution
- Food/water precaution and travelers' diarrhea
- Non-infectious disease risk
- Medical kit
- Look into travel / medical evacuation insurance

