# TROPICAL FEVER EVALUATION





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# PRESENTATION OBJECTIVES



- Identify the unique challenges to evaluating tropical fever
- Employ an effective rationale in differential diagnosis
- Develop reliance upon only limited diagnostic aids

### NOT ADDRESSED TODAY: FEVER AMONG TRAVELERS





Scenario 1

WHY IS **EVALUATING FEVER IN A** TROPICAL SETTING UNIQUELY **CHALLENGING?** 



### TROPICAL FEVER CHALLENGES

- Unfamiliar causes
- Unusual causes
- Limited laboratory and imaging
- Cultural unfamiliarity
- Out of practice clinical skills
- Time pressure



### WHAT DO YOU KNOW ABOUT PATHOGENESIS OF FEVER?





### FEVER MOST COMMONLY IS...

- A physiological response to infection, mediated principally via cytokines.
- Cytokines alter thermoregulation in the hypothalamus through a process mediated by prostaglandins.



### WHAT NON-INFECTIOUS ETIOLOGIES CAUSE FEVER?



# NON-INFECTIOUS CAUSE OF FEVER

- Malignancies, especially lymphomas, leukemias, and renal cell CA
- Autoimmune diseases, including rheumatoid arthritis, temporal arteritis, polyarteritis nodosa
- Drug reactions



### FOR FEBRILE, TROPICAL-LOCALE PATIENTS, MULTIPLE CAUSES MUST BE CONSIDERED



### CLINICAL APPROACH TO FEBRILE TROPICAL-LOCALE PATIENTS



### A Search For Clues

# APPROACH TO FEBRILE TROPICAL-LOCALE PATIENTS: EPIDEMIOLOGY

- Knowledge of local epidemiology is extremely useful in assessing risk
- Local epidemiology may change suddenly with occurrence of epidemics, such as Marburg virus



### WHERE IS INFLUENZA COMMON?



### New Influenza A (H1N1), Number of laboratory confirmed cases as reported to WHO

Status as of 26 June 2009 06:00 GMT



**INFLUENZA** 

The boundaries and names shown and the designations used on this map do not imply the expression of any opinion whatsoever on the part of the World Health Organization concerning the legal status of any country, territory, city or area or of its authorities, or concerning the delimitation of its frontiers or boundaries. Dotted lines on maps represent approximate border lines for which there may not vet be full agreement. Data Source: World Health Organization Map Production: Public Health Information and Geographic Information Systems (GIS) World Health Organization



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### WHERE IS TYPHOID COMMON?



# Typhoid fever 20 Risk area

### **TYPHOID DISTRIBUTION**



# WHERE IS YELLOW FEVER COMMON?



### **YELLOW FEVER DISTRIBUTION**

### WHERE IS MALARIA COMMON?





### MALARIA DISTRIBUTION

### WHERE IS MEASLES COMMON?





### **MEASLES VACCINE COVERAGE**

### WHERE IS TB COMMON?





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### **TUBERCULOSIS INCIDENCE**

### WHERE IS HIV COMMON?



### ditante Occum Partific Ocean Indian Ocean Prevalence of HIV in Adults 10% 5% - 10% %-4.9% <1% No Data

### **HIV PREVALENCE**

### WHERE IS DENGUE COMMON?





### **DENGUE DISTRIBUTION**



### Scenario 1 Diagnosis: Dengue Fever

### WHAT DO YOU KNOW ABOUT DENGUE FEVER?



### **DENGUE SIGNS & SYMPTOMS**

- Abrupt onset of fever and chills
- Severe frontal headache, painful eye movement, and musculoskeletal and lumbar pain present in 70% of patients.
- Anorexia, nausea and vomiting occur in half of patients.
- Initial examination is usually nonspecific and may reveal scleral injection, generalized lymphadenopathy and bradycardia relative to fever.

### QUESTION

Which ONE statement about dengue fever is NOT true?

- A The dengue vaccine is highly protective.
- B The vector for dengue is the *Aedes aegypti* mosquito.
- C Dengue most commonly occurs in Southeast Asia and Latin America.
- D Treatment is essentially supportive with particular attention to hemodynamic status.
- E Infection may progress to complications that include hemorrhage and shock.
#### ANSWER

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# APPROACH TO FEBRILE TROPICAL-LOCALE PATIENTS



WHAT **CLUES CAN** BE **GLEANED FROM THE HISTORY**?



# HISTORY

- Rigors & night sweats suggest a febrile illness even if the patient is afebrile
- Weight loss suggests significant disease
- Pattern of fever is rarely helpful
- A self-recorded temperature chart may be useful
- Upper Respiratory Infection symptoms
  suggest viral cause

# HISTORY

- Freshwater exposure suggests schistosomiasis
- In children, feeding, weight change, and activity are useful measures
- Localizing symptoms should be sought, such as headache, photophobia, cough, sputum, pleurisy, painful sites, diarrhea, urinary symptoms

WHAT **CLUES CAN** BE **GLEANED FROM THE** PHYSICAL EXAM?



## **PHYSICAL EXAM**

- Pulse rate > 120 or systolic BP <100 suggest serious illness
- Spontaneous hemorrhage suggests viral hemorrhagic fever
- Psychosis (typhoid)
- Altered consciousness or neck stiffness (meningitis)

#### WHAT IS THIS FINDING?



# APPROACH TO FEBRILE TROPICAL-LOCALE PATIENTS: EXAM

- Cervical and axillary adenopathy (TB, HIV, CMV, EBV, lymphoma, toxo, syphilis
- Occipital adenopathy (rubella, American trypansomiasis)
- Inspect eyes for anemia, jaundice, conjunctival injection (measles, leptospirosis)
- Inspect fundi for endocarditis
- TMs in children

#### **PHYSICAL EXAM**

- Chest exam for pneumonia, pleural or pericardial effusion (TB, empyema)
- Heart murmurs (endocarditis, rheumatic disease)
- Abd tenderness (appendicitis, peritonitis, PID, amebic liver abscess)

### **PHYSICAL EXAM**

- Hepatomegally (malaria, TB, hepatitis, schisto, hepatoma, liver abscess)
- Splenomegally (malaria, typhoid, leishmaniasis, HIV, mono, lymphoma, leukemia, portal HTN, disseminated TB, brucellosis)
- Ascities (tap)
- Joint effusions (tap)

# Put Some Skin in the Game!

#### WHAT DOES THIS SUGGEST?



## WHAT IS THIS FINDING?



## **SKIN EXAM**

- Inspect mouth for candidiasis (HIV), Koplick spots (measles), and pharyngitis
- Rash (viral exanthems, meningococcal patechiae)
- Eschar (tick-borne rickettsia infection)
- Anesthetic patches with pigmentary changes (leprosy)
- Skin signs for cellulitis

WHAT **CLUES CAN** BE **GLEANED FROM LAB** & IMAGING?



## LAB & IMAGING

- Malaria smears and/or rapid test in malaria areas
- Renal functions (sepsis, severe malaria)
- Liver functions (viral hepatitis)
- ESR or CRP (general assessment, SLE)
- Acute & convalescent serum for retrospective diagnosis
- Ultrasound for liver abscess, splenomegally

# LAB & IMAGING

- CXR (TB, pneumonia)
- Blood cultures (sepsis, typhoid)
- UA (bacteria infection, schistosomiasis)
- Stool O & P
- Sputum AFB
- Lumbar puncture
- HIV and hepatitis ab tests
- Syphilis serology
- Genital swab for micro

# APPROPRIATE USES OF TECHNOLOGY





For what tropical febrile diseases are rapid diagnostic tests available?

# APPROPRIATE USES OF TECHNOLOGY



Rapid diagnostic tests for leptospirosis, syphilis, malaria, visceral leishmaniasis, typhoid, trypanosoma, and more





WHAT PRINCIPLES CAN BE USED TO DETERMINE WHETHER A TECHNOLOGY IS APPROPRIATE OR NOT?



#### **IS IT RELIABLE?**



## **IS IT SUSTAINABLE?**



#### **IS IT ACCEPTABLE?**



- Based on timeline, malaria smears and WBC count and differential
- Omitted are those conditions with obvious localizing signs, such as pyogenic arthritis
- Includes only more common causes



- Negative malaria smears
- Neutrophilia: pyogenic infections, leptospiral infections, amebic liver abscess, relapsing fevers (*Borrelia*)
- No neutrophilia: viral infections, rickettsial infections, typhoid

- Negative malaria smears
- *Neutrophilia*: sepsis, amebic liver abscess, cholangitis, relapsing fever, erythema nodosum leprosum

- Negative malaria smears
- Eosinophilia: invasive hepatic schistosomiasis (S. mansoni), invasive Fasciola hepatica infection, acute exacerbations of lymphatic filariasis (Wuchereria bancrofti), visceral larva migrans (Toxocara canis)

- Negative malaria smears
- Neutropenia: malaria, disseminated TB, visceral leishmaniasis, brucellosis, influenza, dengue, yellow fever

- Negative malaria smears
- Normal WBC: HIV, localized TB, brucellosis, secondary syphilis, trypansomiasis, toxoplasmosis, endocarditis

# WHAT IS THE NATURAL HISTORY OF TROPICAL FEVERS?



## NATURAL HISTORY OF TROPICAL FEVERS



At least 50% of adults and older children with genuine fever will have no cause identified and fever will resolve spontaneously in a few days.

#### **BUT REMAIN VIGILANT!**



# If the cause is not immediately apparent, it may become so in days.

# WHEN SHOULD EMPIRICAL TREATMENT BE INITIATED?


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- In adults it is preferable to delay anti-infective treatment until diagnosis is made
- It is poor practice to treat malaria empirically unless no diagnostic test is available

# WHEN SHOULD EMPIRICAL TREATMENT BE INITIATED?

 In ill appearing children with leukocytosis empirical treatment may be prudent

• Suspected sepsis or meningitis should be treated immediately

• Severe pharyngitis should be treated with a low threshold given the increased frequency of post-streptococcal complications.

# EMPIRICAL TREATMENT: BROAD GENERALIZATION

- In many low-resource, tropical regions malaria is the most common cause of febrile illness.
- If malaria treatment fails, consideration is next given to typhoid fever.
- If typhoid treatment fails, consideration is then given to tuberculosis



Scenario 2

#### LAB RESULTS

Malaria Test: Negative

WBC 15,500 Segs 67, Lymphs 15, Eos 18

# WHAT DO YOU KNOW ABOUT HEPATIC SCHISTOSOMIASIS?





#### SCHISTO BACKGROUND

- Schistosomes are trematodes

   (flatworms) that contaminate freshwater
   by infected persons. *S. mansoni* and *S.
   japonicum* ultimately reside in the
   intestine and most disease
   manifestations are hepatic.
- Schistosomiasis is most common in communities with poor sanitation. The prevalence of schistosomiasis is some 200 million people worldwide. In terms of morbidity, this disease is second only to malaria as the most dangerous parasitic disease.

#### **SCHISTO SIGNS & SYMPTOMS**

- Most infected persons in endemic areas are asymptomatic. Infection may manifest as skin rash and itching, fever, cough, muscle aches, headache, diarrhea, and hepatosplenomegaly.
- Chronic schistosomiasis from *S. mansoni* and *S. japonicum* may cause weight loss, anorexia, melena, jaundice, hepatosplenomegaly, intestinal ulceration and polyposis, portal hypertension, and liver failure.
- Diagnosis of schistosomiasis is usually made by identification of ova in feces, tissue or urine.



#### **SCHISTOSOME EGGS (OVA)**

## QUESTION

All are true concerning schistosomiasis of the GI tract are true EXCEPT which ONE?

A. Persons infected with *S. mansoni commonly* present with clinical symptoms that include rapid onset lower gastrointestinal bleeding.

B. People with *S. haematobium* infection usually have a history of fresh water exposure.

C Humans are infected by free-swimming *Schistosoma* cercariae (larvae) that have developed in snails.

D. *S. mansoni* and *S. japonicum* reside in the intestine and primarily cause hepatic disease.

#### ANSWER

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TROPICAL **FEVER EVALUATION: ONE OF** CLINICAL **MEDICINE'S** GREATEST **CHALLENGES!** 



#### EQUIP YOURSELF TO BETTER SERVE FORGOTTEN PEOPLE



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