

## Transfusion-Transmitted Malaria in the United States: Focus on FFP

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#### **Transfusion-Transmitted Malaria**

- Background
- Epidemiology in the United States, as relevant to FFP
- Lookback at outcome of recipients of FFP from known malaria-infected donors
- Medical literature





# Malaria

Protozoan parasitic disease

 Caused by 1 of 4 species of *Plasmodium*:
 *P. falciparum*, *P. vivax*, *P. malariae*, *P. ovale*

• Transmitted by bite of infected female anopheline mosquito



### Malaria in the United States

- Eradicated in late 1940s
- Approximately 1,000 1,400 cases reported annually

   Almost all imported
   US travelers or foreigners
- Each year a few cases (<10) are transmitted within US
   Transfusion or organ transplantation
   Congenitally acquired
   Local mosquito-borne transmission





#### **Imported Malaria**

 Approximately half of cases immigrants, refugees, residents of malarious countries who come to US

 Half in US travelers - civilians and military

 Major risk factor is failure to use effective chemoprophylaxis
 85% of cases use no or ineffective prophylaxis





#### Malaria Transmission Cycle

Sporozoites injected into human host during blood meal Exo-erythrocytic (hepatic) Cycle: Sporozoites infect liver cells and develop into schizonts, which release merozoites into the blood (7-16 days)

Dormant liver stages (hypnozoites) of *P. vivax* and *P. ovale* 

Parasite undergoes sexual reproduction in the mosquito (9-35 days) Some merozoites differentiate into male or female gametocyctes Erythrocytic Cycle: Merozoites infect red blood cells to form schizonts (36-72 hours)



#### **Malaria Species**



No persistent liver stage (non-relapsing)

- P. malariae can persist at low levels in blood for years to decades
- P. falciparum rarely persists beyond 1-2 years
   Responsible for most deaths due to malaria
- <u>Persistent liver stage (relapsing)</u> rarely last beyond 3 years
- P. vivax
- P. ovale



#### Transfusion-Transmitted Malaria

- Rare in the US
   Estimated incidence: 1 case/ 4 million units collected
- High case fatality rate

• 3 complicated *Plasmodium falciparum* infections reported during 1996-1998





# Transfusion-Transmitted Malaria in the United States, 1963-1999

- 93 reported cases
- Implicated blood component (n = 70)
  Whole blood 63%
  Packed RBCs 31%
  - Platelets 6%
- 67 implicated donors

• No change in incidence in last 15 years



#### What about "unsolved cases"?

- Any suggestion FFP could have been the infective blood product?
- Case-patient got FFP in 10 of 92 cases where data were available
  - In 6 cases, an implicated donor was found
    - In 5 of those 6, whole blood or PRBCs
    - In 1 of those cases, implicated product not mentioned in records (patient got PRBCs, platelets and plasma)



#### What about "unsolved cases"?

#### • What about the other 4 cases?

- 1 case PRBCs/FFP both RBC donors serologically neg but had traveled (unknown where); couldn't track FFP donors
- 1 case PRBCs/ WB/FFP all donors neg but 1 donor had traveled to Ghana – unknown which component he donated
- 2 cases incomplete investigation couldn't even track the whole blood donors
- Bottom line: no cases where evidence for transmission from other components definitively ruled out

#### Lookback: Recipients of FFP from Malaria-Infected Donors, 1990-2000

14 cases (one investigation still in progress)

Of the 13 remaining cases:
No FFP transfused = 4
FFP transfused – no evidence subsequent malaria = 3
Investigation pending = 6



#### Evidence in the Medical Literature Related to Risk from FFP?

Bruce-Chwatt\*
 Reports to WHO from 1911-1972

 2001 reported cases without mention of infection associated with FFP transfusion

\* Bruce-Chwatt LJ. Transfusion malaria. Bull World Hlth Organ 1974;50:337-46.



#### Evidence in the Medical Literature Related to Risk from FFP?

- Lozner & Newhouser, 1943\* transfused plasma from patients with malaria into 35 recipients
  - No transmission from frozen (n=20) or dried (n=3) plasma

 Liquid plasma – transmission from plasma stored 1 day (2 of 2), ? in that stored for 1 week (1 of 5), and none stored for 2 weeks (n=5)

\*Lozner EI, Newhouser LR. Studies on the transmissibility of malaria by plasma transfusions. Am J Med Sci 1943;206:141-46.

