

*VIII Curso Alma  
Cartagena, Colombia  
Agosto, 2009*



# **CAMBIOS CLIMÁTICOS E INFECCIONES**

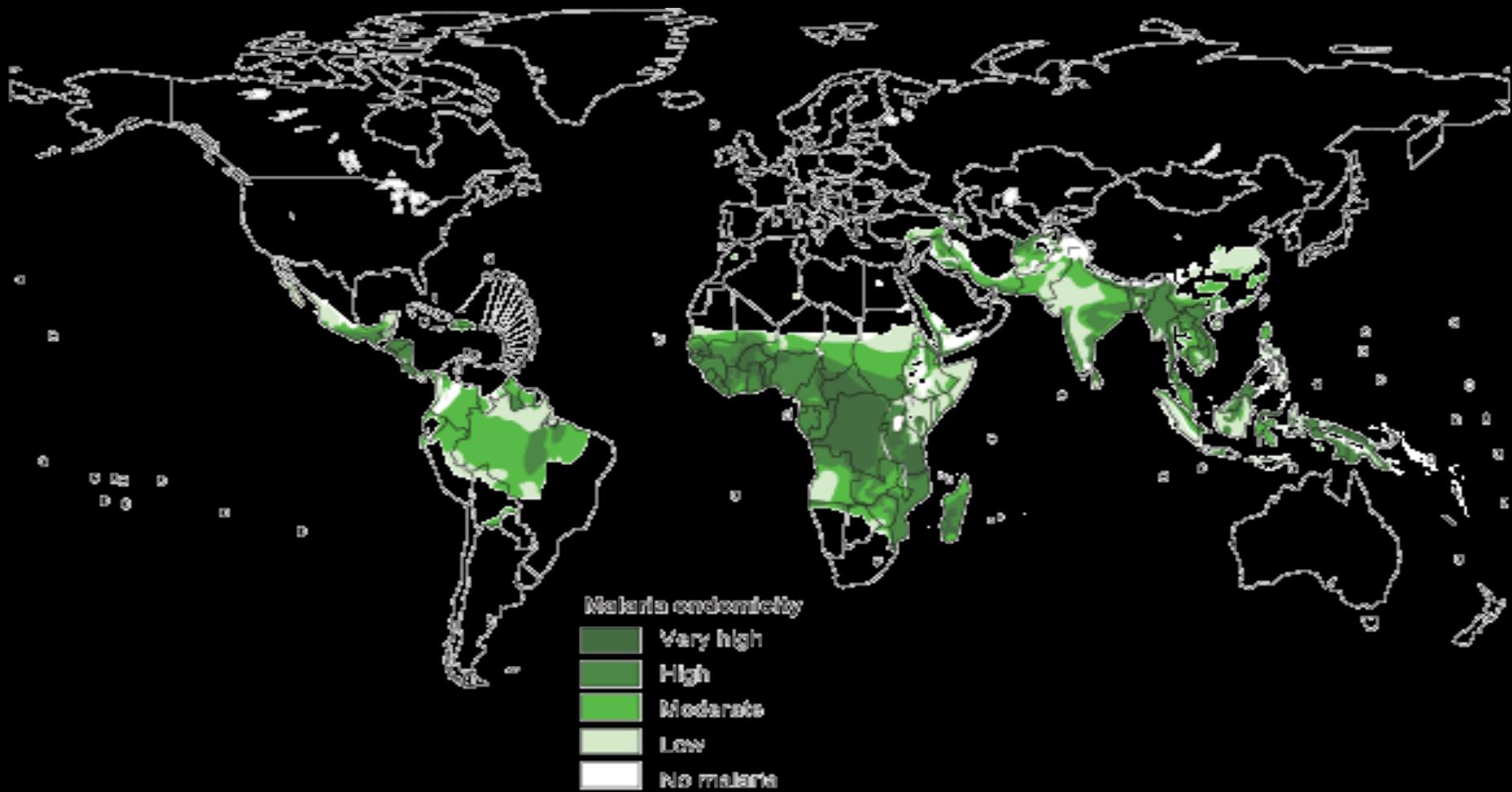
## **PALUDISMO**

Ulises Perez - Mexico

Virgílio Garcia Moreira - Brasil

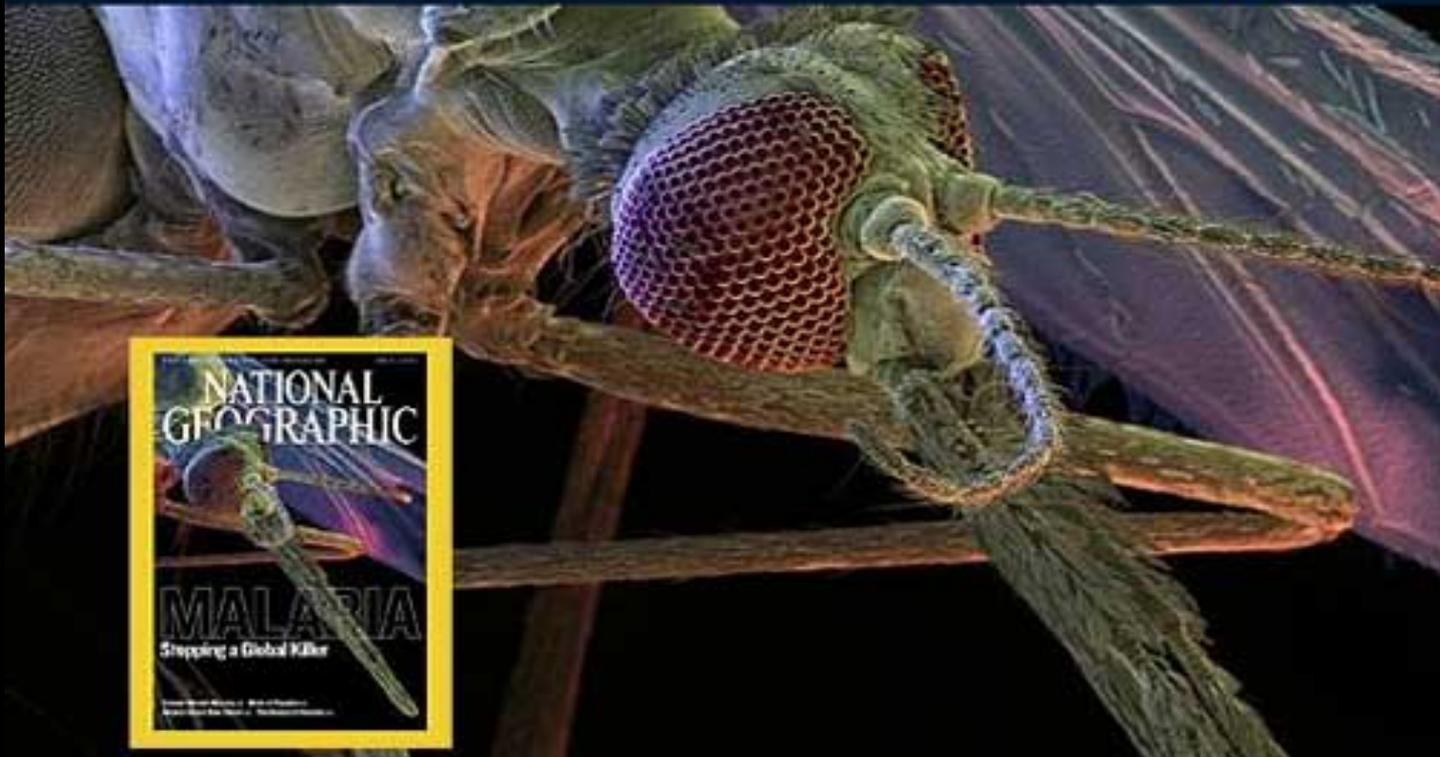


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# Malaria: Bedlam in the Blood

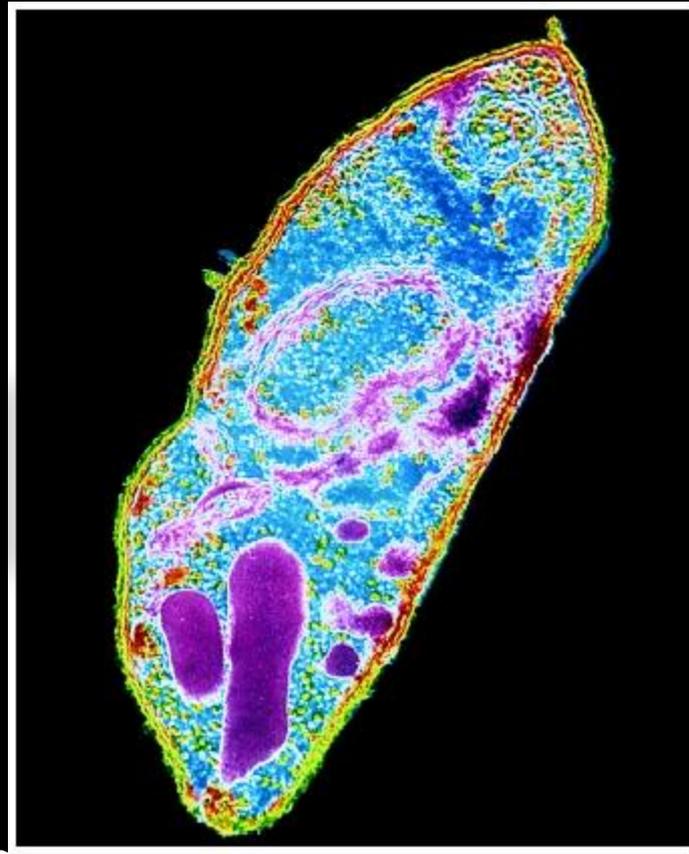
1 2 3 4



# Malária



**Mais comuns do planeta**



**300 Milhões**

**2 a 3 milhões de óbitos**



M A L A R I A

**KILLS  
3000  
CHILDREN  
EVERYDAY**

IN AFRICA. PLEASE HELP.

[HTTP://MALARIANOMORE.ORG](http://malariaanomore.org)

# Malária

## Definição

- Lise Sincrônica de Hemácias parasitadas

# Malária



# Age as a Risk Factor for Severe Manifestations and Fatal Outcome of Falciparum Malaria in European Patients: Observations from TropNetEurop and SIMPID Surveillance Data

N. Mühlberger, T. Jelinek, R. H. Behrens, I. Gjørup, J. P. Coulaud, J. Clerinx, S. Puente, G. Burchard, J. Gascon, M. P. Grobusch, T. Weitzel, T. Zoller, H. Kollaritsch, J. Beran, J. Iversen, C. Hatz, M. L. Schmid, A. Björkman, K. Fleischer, Z. Bisoffi, W. Guggemos, J. Knobloch, A. Matteelli, M. H. Schulze, H. Laferl, A. Kapaun, P. McWhinney, R. Lopez-Velez, G. Fätkenheuer, P. Kern, B. W. Zieger, A. Kotlowski, G. Fry, J. Cuadros, and B. Myrvang, for the TropNetEurop and Surveillance importierter Infektionen in Deutschland (SIMPID) Surveillance Networks\*

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R. Lopez-Velez, G. Fätkenheuer, P. Kern, B. W. Zieger, A. Kotlowski, G. Fry, J. Cuadros, and B. Myrvang,  
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Clinical Infectious Disease. 2003;36:990-5

**Table 2. Age-specific frequency of outcomes of severe falciparum malaria.**

Age group, years	No. of patients	Percentage of patients with			
		Fatal cases	Cerebral complications	Overall complications	Hospital admission <sup>a</sup>
10–19	50	0.0	0.0	4.0	70.0
20–29	290	0.7	1.0	6.2	79.0
30–39	369	0.5	1.9	7.3	79.7
40–49	225	2.2	6.2	13.8	81.8
50–59	169	2.4	4.7	14.2	85.2
60–69	56	3.6	5.4	8.9	73.2
70–79	20	10.0	10.0	30.0	95.0
80–89	2	0.0	0.0	0.0	100.0
Total	1181	1.4	3.1	9.6	80.3

100.0  
 80–89 5 0.0 0.0 0.0 100.0  
 10–19 50 0.0 0.0 4.0 70.0  
 Clinical Infectious Disease. 2003;36:990-5 990



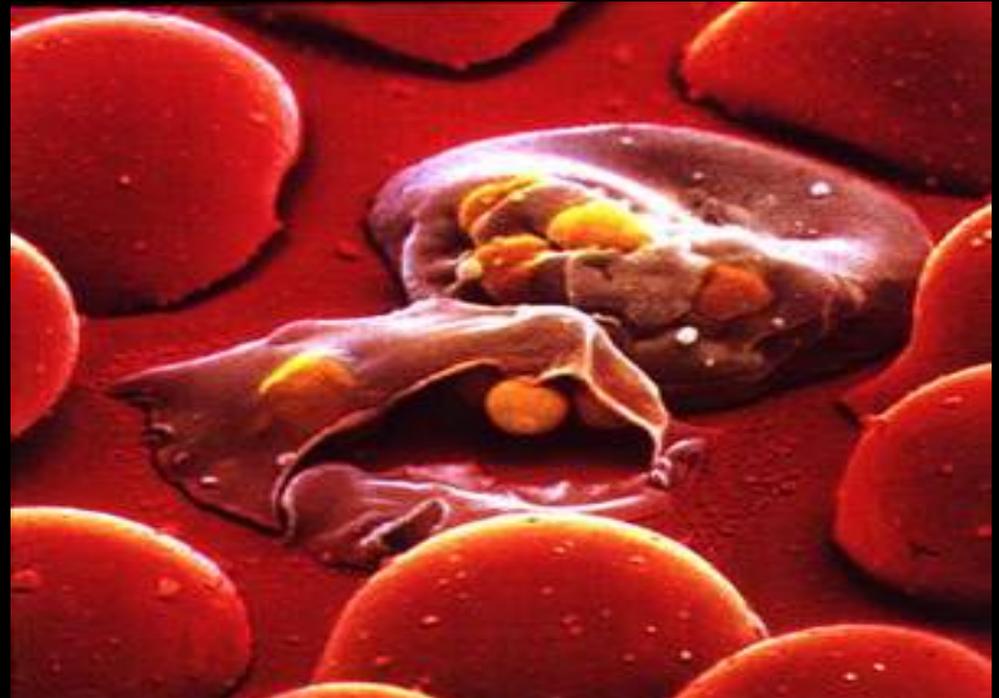
⇒ Estômago e glândulas salivares

Anopheles sp.



# Malária

## Gravidade da Doença



- Espécie Infectante
- Magnitude da Parasitemia
- Efeitos Metabólicos do Parasita no organismo

# The Relationship between Age and the Manifestations of and Mortality Associated with Severe Malaria

**Arjen M. Dondorp,<sup>1,3</sup> Sue J. Lee,<sup>3</sup> M. A. Faiz,<sup>4</sup> Saroj Mishra,<sup>6</sup> Ric Price,<sup>3,7</sup> Emiliana Tjitra,<sup>8</sup> Marlar Than,<sup>9</sup> Ye Htut,<sup>10</sup> Sanjib Mohanty,<sup>6</sup> Emran Bin Yunus,<sup>5</sup> Ridwanur Rahman,<sup>5</sup> Francois Nosten,<sup>1,2,3</sup> Nicholas M. Anstey,<sup>7</sup> Nicholas P. J. Day,<sup>1,3</sup> and Nicholas J. White<sup>1,3</sup>**

<sup>1</sup>Faculty of Tropical Medicine, Mahidol University, Bangkok, and <sup>2</sup>Shoklo Malaria Research Unit, Mae Sot, Thailand; <sup>3</sup>Centre for Tropical Medicine, Nuffield Department of Clinical Medicine, John Radcliffe Hospital, University of Oxford, Oxford, United Kingdom; <sup>4</sup>Department of Medicine, Dhaka Medical College, Dhaka, and <sup>5</sup>Chittagong Medical College, Chittagong, Bangladesh; <sup>6</sup>Department of Medicine, Ispat Hospital, Rourkela (Orissa), India; <sup>7</sup>International Health Division, Menzies School of Health Research and Charles Darwin University, Darwin, Northern Territory, Australia; <sup>8</sup>National Institute of Health Research and Development, Jakarta, Indonesia; and <sup>9</sup>Clinical Research Unit (Malaria), Defence Services General Hospital, Mingaladon, and <sup>10</sup>Department of Medical Research, Yangon, Myanmar

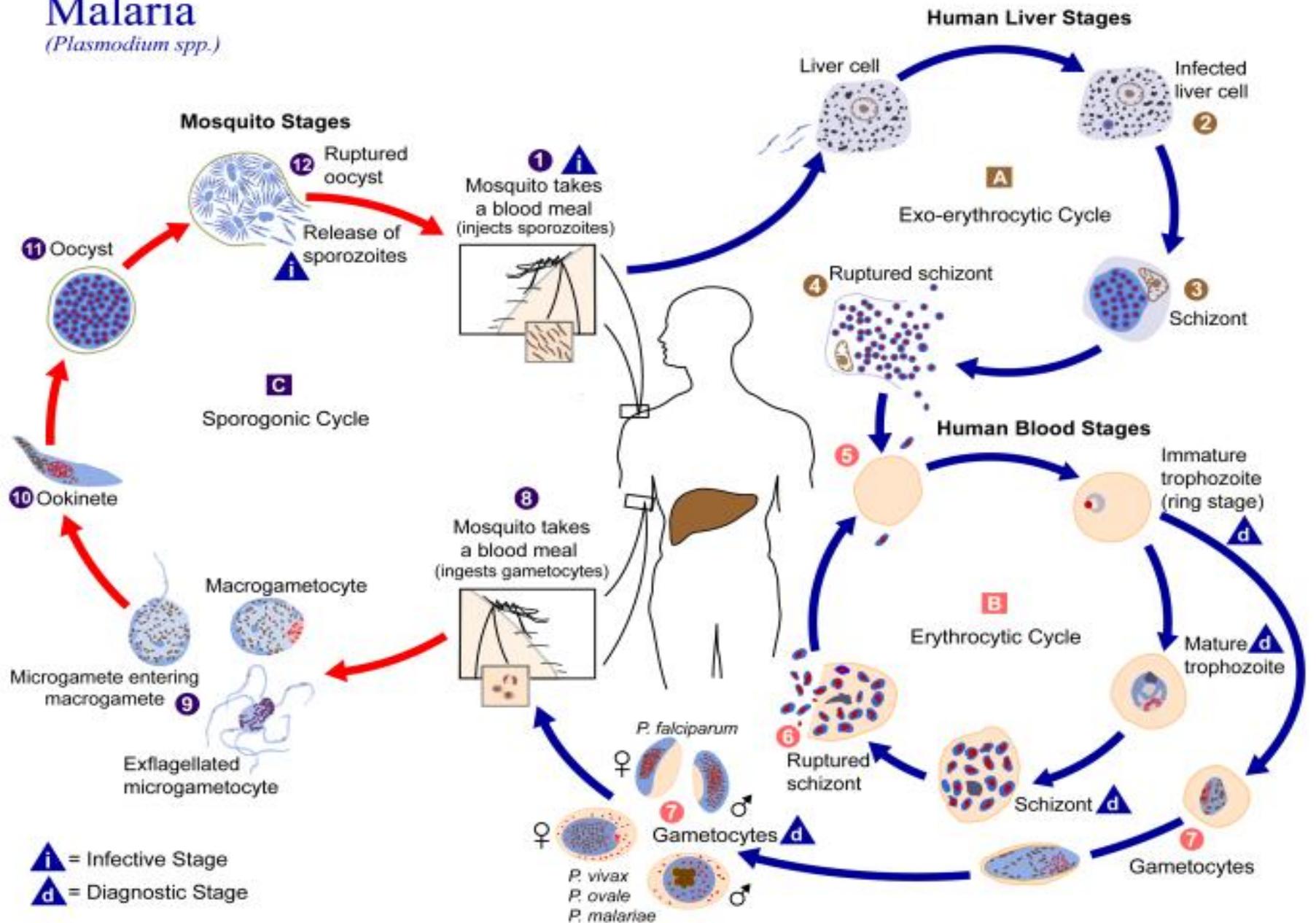
Clinical Infectious Disease. 2008. 47:151-7

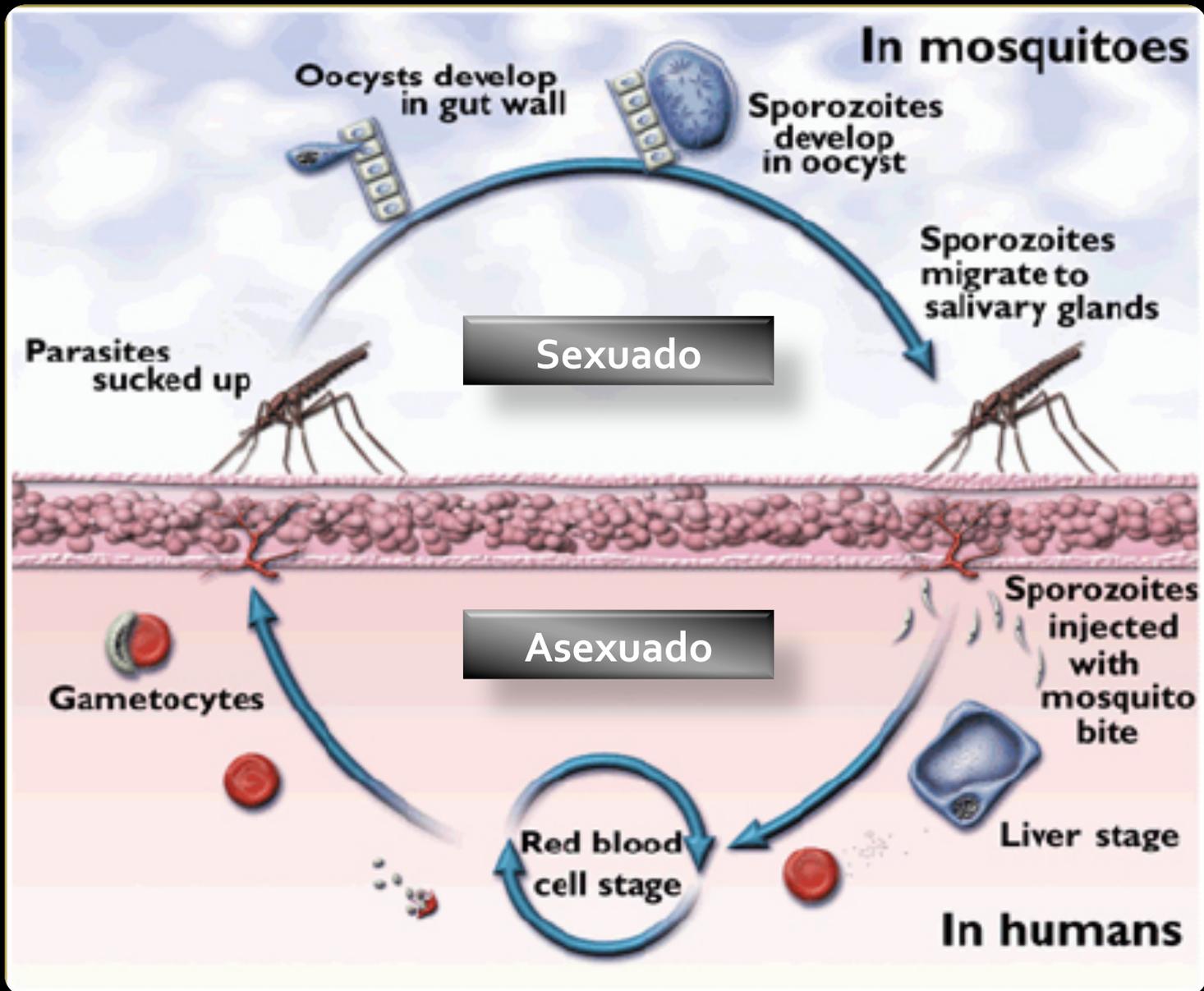
**Table 2. Univariate analysis of risk of death among patients with and without severity syndromes (SS) at hospital admission.**

Age group, years	Coma				Convulsions				Severe anemia			
	Patients who died, %		OR (95% CI)	P	Patients who died, %		OR (95% CI)	P	Patients who died, %		OR (95% CI)	P
	With SS	Without SS			With SS	Without SS			With SS	Without SS		
0-10	11.7	0	NA	.01	7.1	5.6	1.31 (0.28-6.19)	.73	2.3	4.8	0.47 (0.05-4.70)	.51
11-20	32.6	10.5	3.29 (1.73-6.25)	<.001	23.7	21.6	1.13 (0.50-2.53)	.77	21.6	21.8	0.99 (0.47-2.09)	.97
21-50	35.5	15.8	3.11 (2.03-4.76)	<.001	32.1	26.1	1.34 (0.74-2.44)	.33	25.6	25.6	1.00 (0.58-1.74)	.99
>50	48.3	16.7	4.25 (1.5-11.48)	.002	66.7	33.3	4.0 (0.90-17.82)	.05	28.6	37.4	0.67 (0.12-3.71)	.65
All	33.6	12.6	3.44 <sup>a</sup> (2.46-4.79)	<.001	24.8	24.0	1.41 <sup>a</sup> (0.92-2.18)	.11	18.9	24.2	0.94 <sup>a</sup> (0.62-1.44)	.79

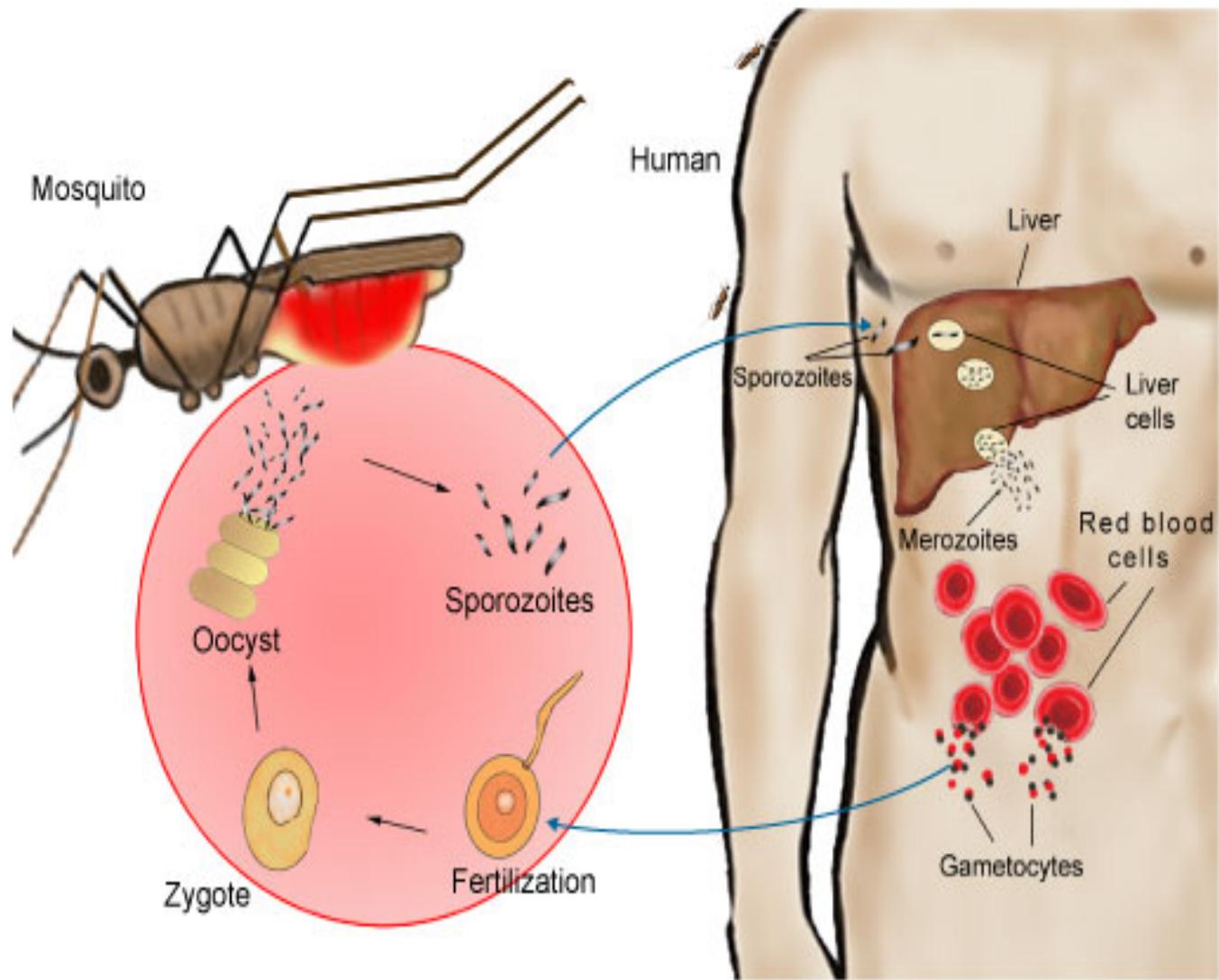
# Malaria

(*Plasmodium spp.*)

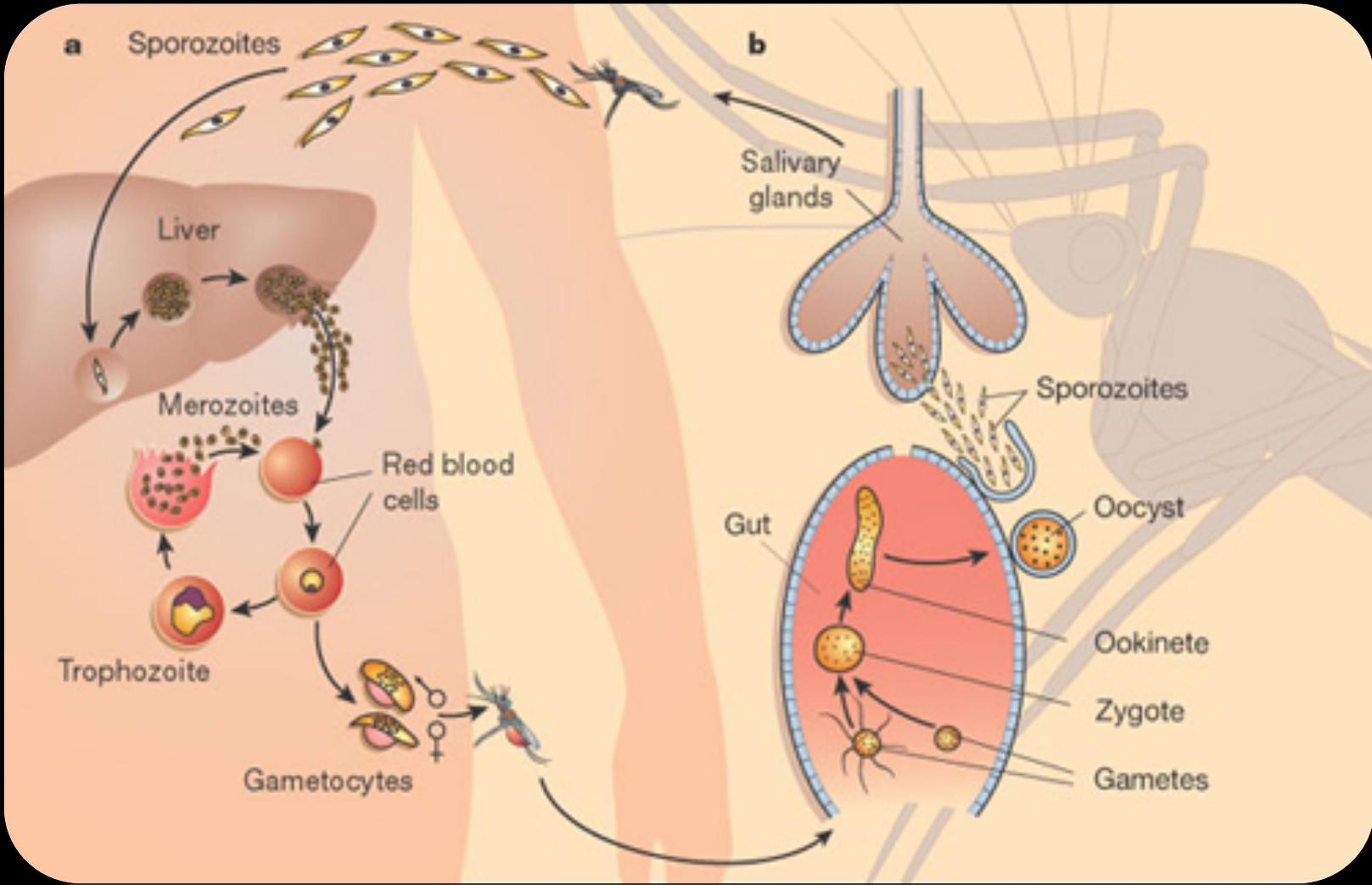




In humans



Life cycle of *Plasmodium*, protist that causes Malaria



## Life Cycle of Plasmodium

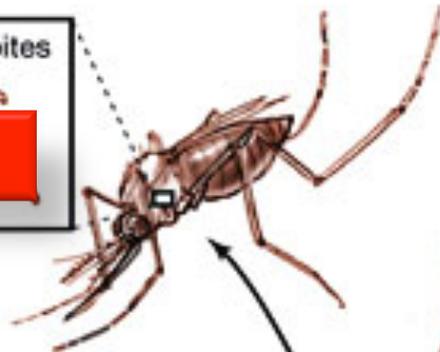
- ① Female *Anopheles* mosquito infected with the malaria parasite (*Plasmodium*) bites a human and transmits the infective form of the parasite (sporozoites) into the human's blood.

Mosquito

- ② Sporozoites enter liver cells and multiply, forming merozoites that burst out of liver cells.



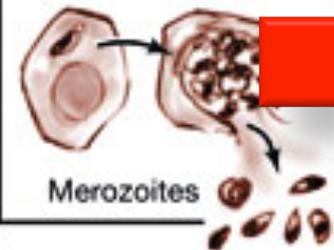
Esporozoíto



Liver

Infected Liver Cells

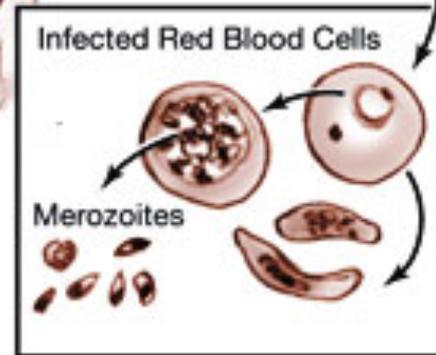
Merozoíto



- ⑤ Sporozoites (infective form), which develop from gametocytes, travel to the salivary glands of the mosquito.

Mosquito

- ④ Female *Anopheles* mosquito bites a human infected with *Plasmodium*, taking up gametocytes with its blood meal.



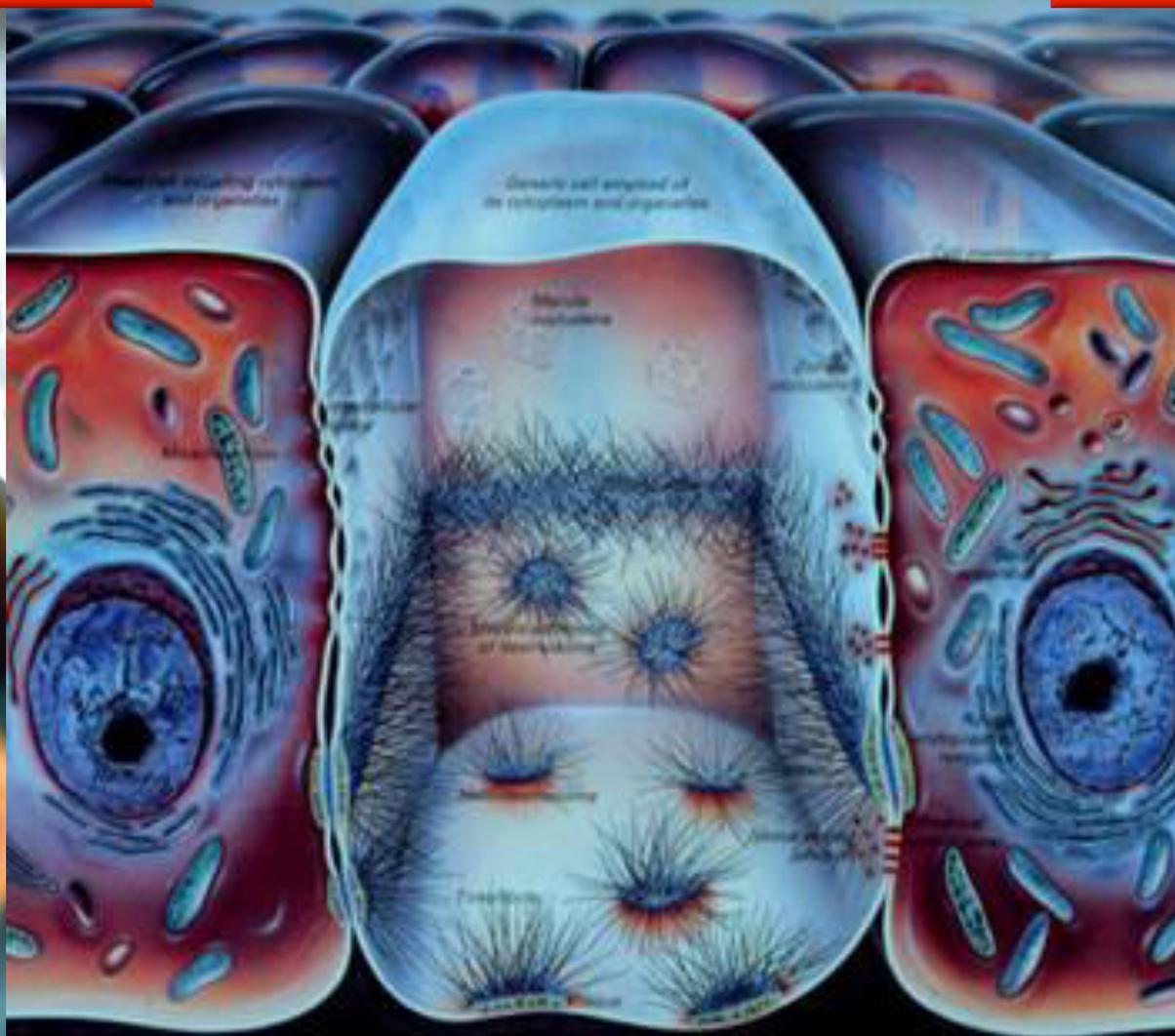
- ③ Merozoites enter red blood cells and multiply, forming new merozoites or developing into gametocytes (reproductive form).

Gametócitos

C. Lynn

Interferon  $\mu$

TNF



# Niveles séricos elevados de interleuquina-10 en pacientes con malaria aguda

Acta méd. colomb;24(1):15-8, ene.-feb. 1999. graf.

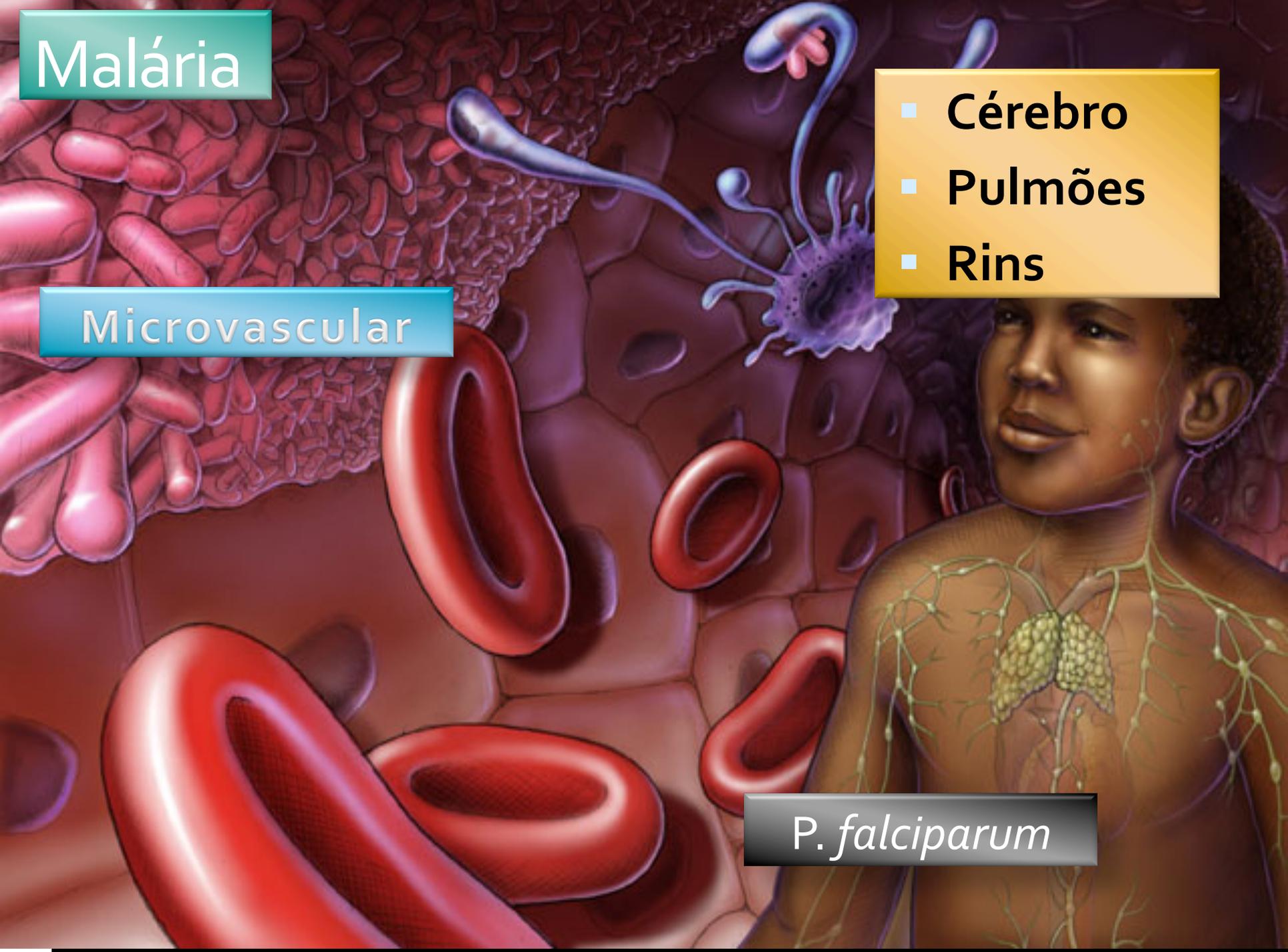
La malaria continúa siendo un problema grave de salud en nuestro país y en el mundo. La respuesta inmune que se genera frente al Plasmodium es bastante compleja y aún no es completamente clara. El objetivo de nuestro trabajo fue investigar los niveles séricos de factor de necrosis tumoral alfa (TNF-Ó) y de interleuquina 10 (IL-10) en personas con malaria y sin ella, y correlacionarlos con el número de parásitos y la temperatura corporal. La población de estudio provenía de El Bagre, municipio de Antioquia, y comprendía 49 niños sin malaria y 51 con malaria infectados por Plasmodium vivax (67 por ciento), por plasmodium falciparum (29 por ciento) o por ambos (4 por ciento). El número promedio de parásitos circulantes de P. vivax fue 5.495/mm<sup>3</sup>. Para la cuantificación de citoquinas se emplearon estuches comerciales basados en una prueba de ELISA en «sandwich». Los resultados mostraron un aumento significativo de IL-10 en los maláricos (266.18±47.9 pg/ml) comparado a los no maláricos (8.52±1.17pg/ml) ( $p < 0.0001$ ). Hubo una correlación directa entre los niveles de IL-10 con el número de parásitos ( $p < 0.0001$ ) y el aumento en la temperatura corporal ( $p < 0.0001$ ). En la mayoría de la población estudiada con malaria (88.2 por ciento) y sin ella (87.8 por ciento) no se detectó TNF-Ó y en el resto, la concentración fue mínima. La no detección de TNF-Ó puede tener dos explicaciones fundamentales: la baja parasitemia, el aumento de IL-10, o ambos. Es difícil explicar la relación de la IL-10 con la parasitemia y la fiebre dado que no se le ha atribuido acción pirogénica a esta citoquina. (AU)

# Malária

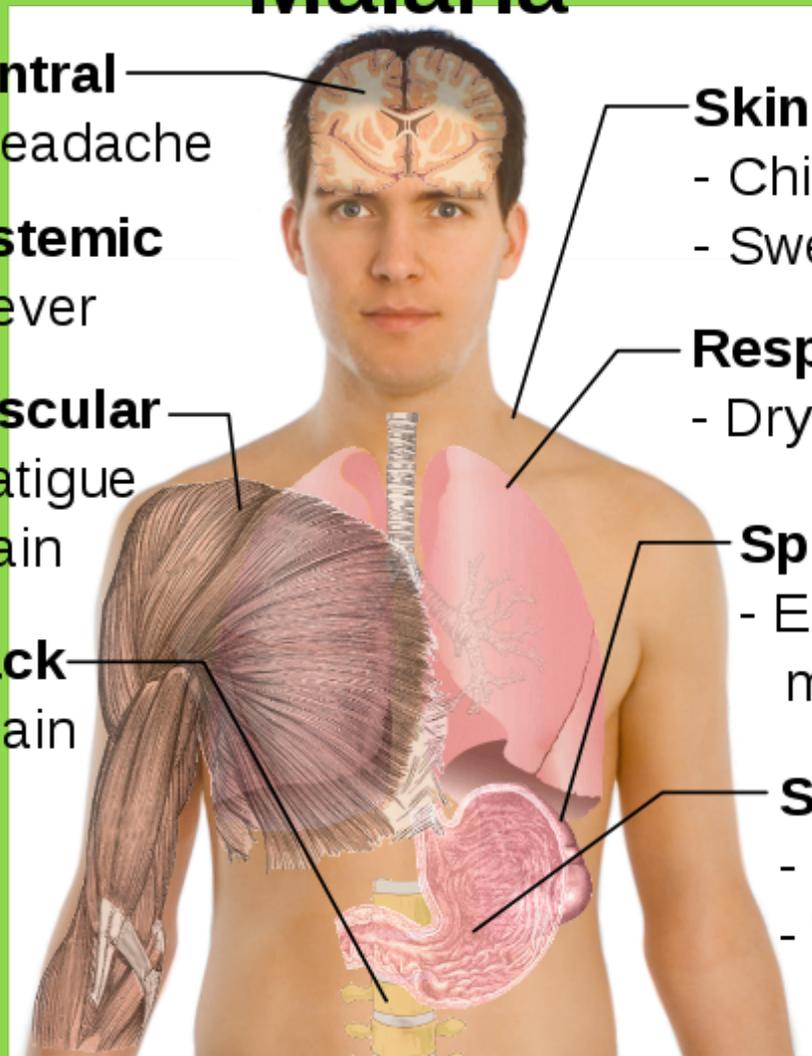
Microvascular

- Cérebro
- Pulmões
- Rins

*P. falciparum*



# Symptoms of Malaria



## Central

- Headache

## Systemic

- Fever

## Muscular

- Fatigue
- Pain

## Back

- Pain

## Skin

- Chills
- Sweating

## Respiratory

- Dry cough

## Spleen

- Enlarge-ment

## Stomach

- Nausea
- Vomiting

Anemia

Coma

Edema  
Agudo  
Pulmonar

Renal

Diarréa

Hepático

Hipoglicemia

# MALÁRIA



## ■ DIAGNÓSTICO

\* CLÍNICO

\* Laboratorial

⇒ Pesquisa do Plasmódio

→ Esfregaço sanguíneo e em gota espessa (Giemsa)

⇒ Sorológico:

→ RIFI, ELISA, HAI, FIXAÇÃO DO  
COMPLEMENTO, PCR.

# MALÁRIA

## ■ TRATAMENTO

- \* Quinina ⇨ Age sobre os trofozoítos, esquizontes e merozoítos
- \* Amino-4-quinolonas (Cloroquina, Aralen, Amodiaquina ou camoquim) ⇨ Age sobre todas as formas sanguíneas exceto gametócitos de *P. falciparum*
- \* Amino-8-quinolinas (Plasmoquina, Primaquina) ⇨ Age sobre os esquizontes hepáticos e sobre os gametócitos.
- \* Pirimidina (pirimetamina) ⇨ Age sobre formas hepáticas e sanguíneas
- \* Sulfonamida (Fanasulf) ⇨ *P. falciparum* cloroquino resistente

# MALÁRIA

- PROFILAXIA

- \* INDIVIDUAL

- ⇒ Evitar a aproximação às áreas de risco após o entardecer e logo ao amanhecer do dia

- ⇒ Uso de repelentes, dormir com mosquiteiros, telar janelas e portas

- ⇒ Uso de quimioproláticos → Iniciar 01 semana antes  
(Mefloquina) → Terminar 30 dias após

# MALÁRIA

## \*COLETIVA

**CONTROLE** ⇒ Reduzir a incidência da doença em determinadas áreas.

Medidas de combate ao vetor

Medidas de combate às larvas

Medidas de saneamento básico

Medidas para melhorar as condições de vida

Tratamento dos doentes

# Malária – Propostas de Tratamento

