

Eosinophilia and Schistosomiasis

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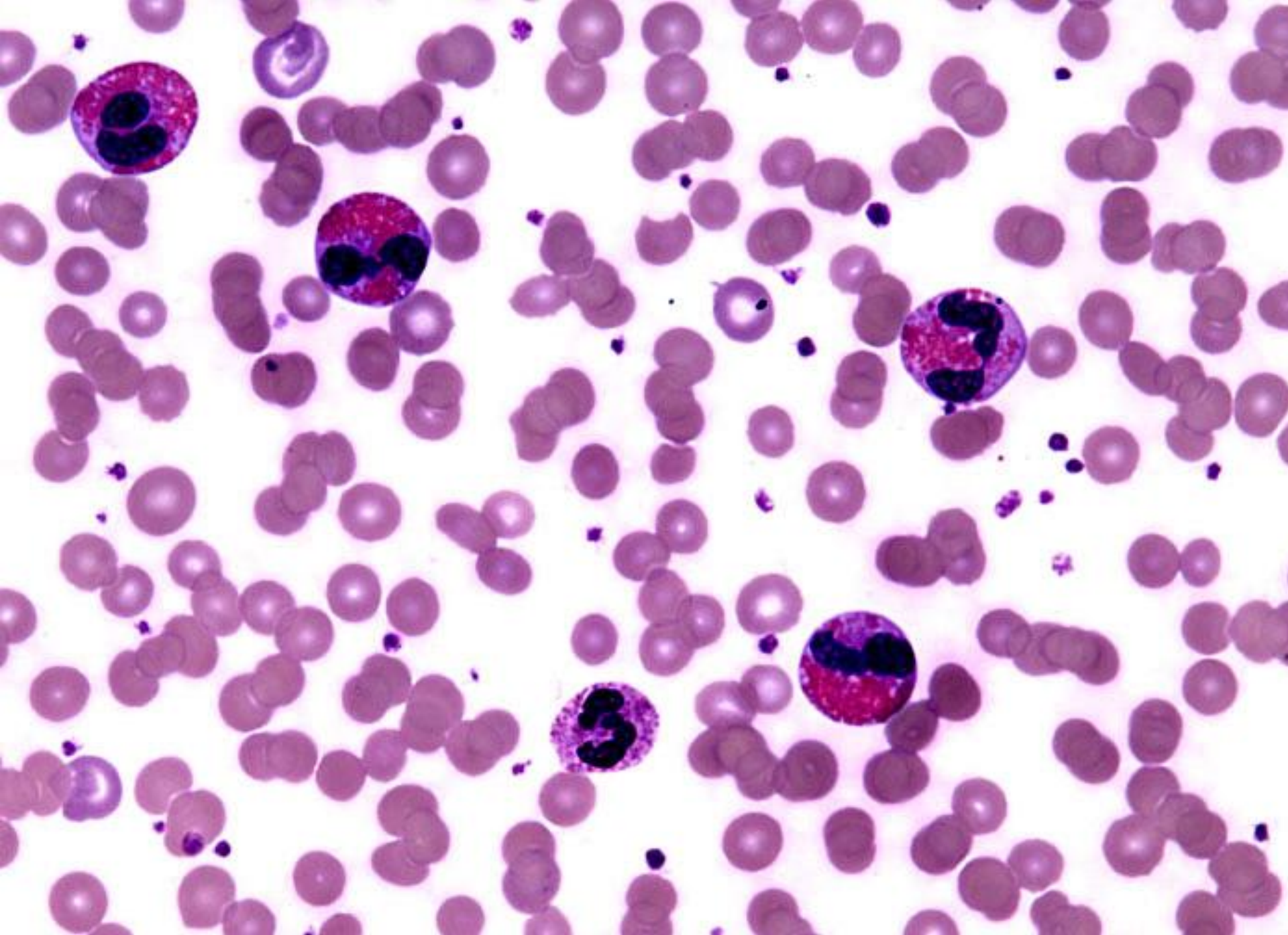
Eozinophilia

= Increased level of eosinophils:

Alergic reaction

Parasitic infection

Not present in protozoal infections!!!



Level of eosinophils

Very high (30-80% WBC)

Trichinella, Toxocara, Fasciola

Medium high (10-30% WBC)

Strongyloides, Ancylostoma, Necator

Low, non-existing (0-10% WBC)

Enterobius, Ascaris, Trichuris

Deposition of helminth in the tissue

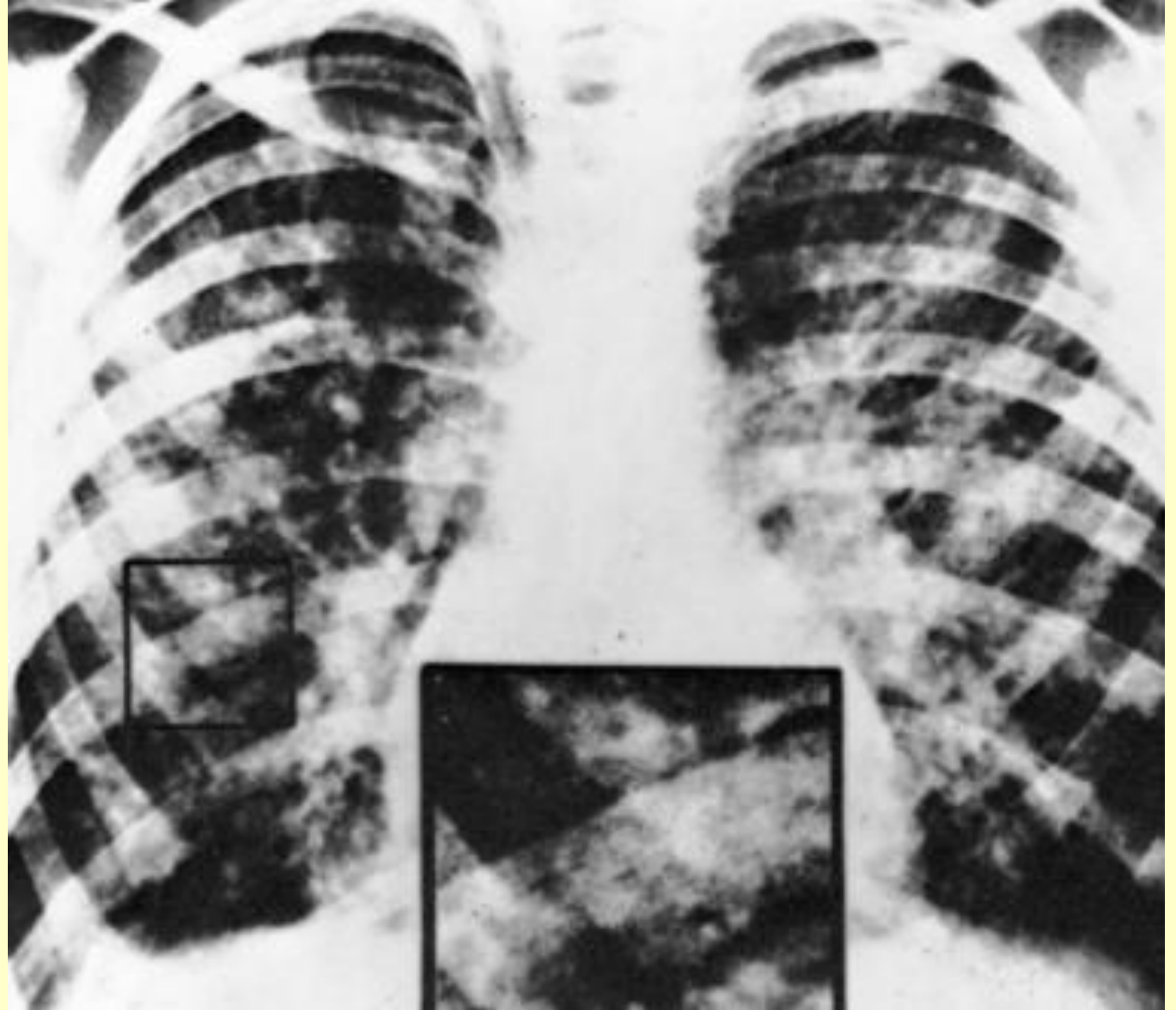
Final site of development –
Final host

Final site of development
- **Intermediate host**

Transitory site of development

Tissues as Transitory site of development

- *Ascaris lumbricoides*
- *Ancylostoma, Necator, Strongyloides*
- As part of development, helminths migrate within the host
- Transitory site of development – **Lungs**
- Host **usually asymptomatic**; heavy infection
- **pneumonia**



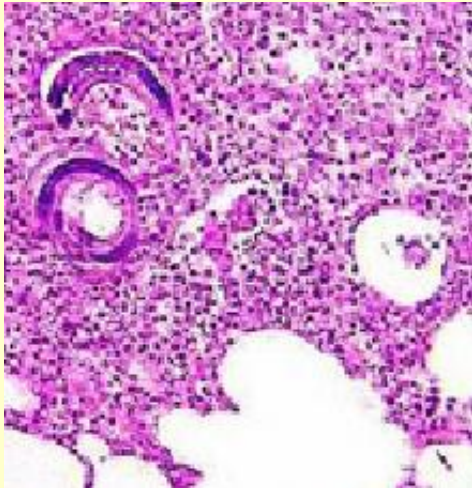
Pneumonia

Cough, dyspnea, nausea, vomiting

Lofflers eosinophilic syndroma

Blood in sputum

Larvae of parasites



Symptoms associated with the migration within the host disappear after residing within intestine

Eosinophilia low or non-existent

Symptoms depend on the final site of development

GIT problems: asymptomatic vs symptomatic

Individual

DIAGNOSTICS!!!!

(ova/parasites in the stool 50 – 80 days pi)

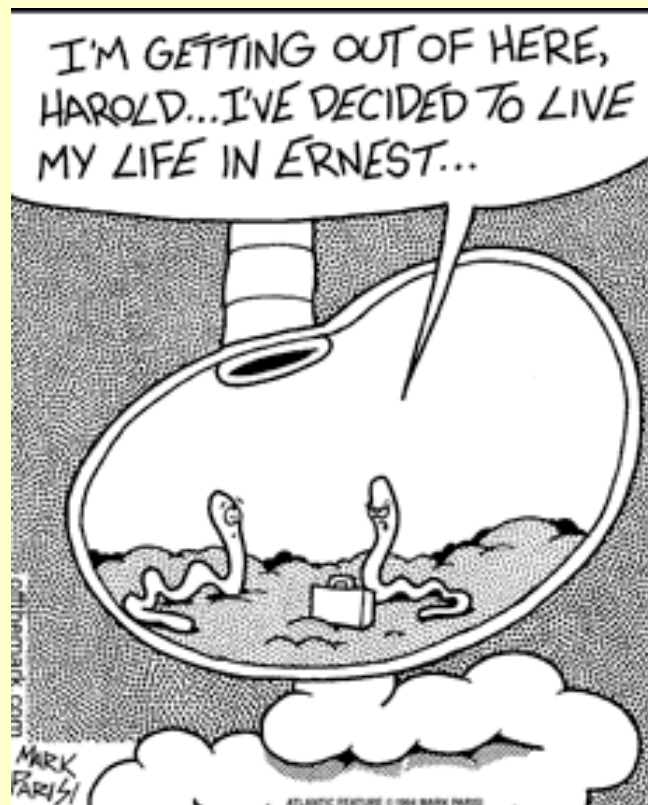
Therapy

Pneumonia: symptomatic; prednisone

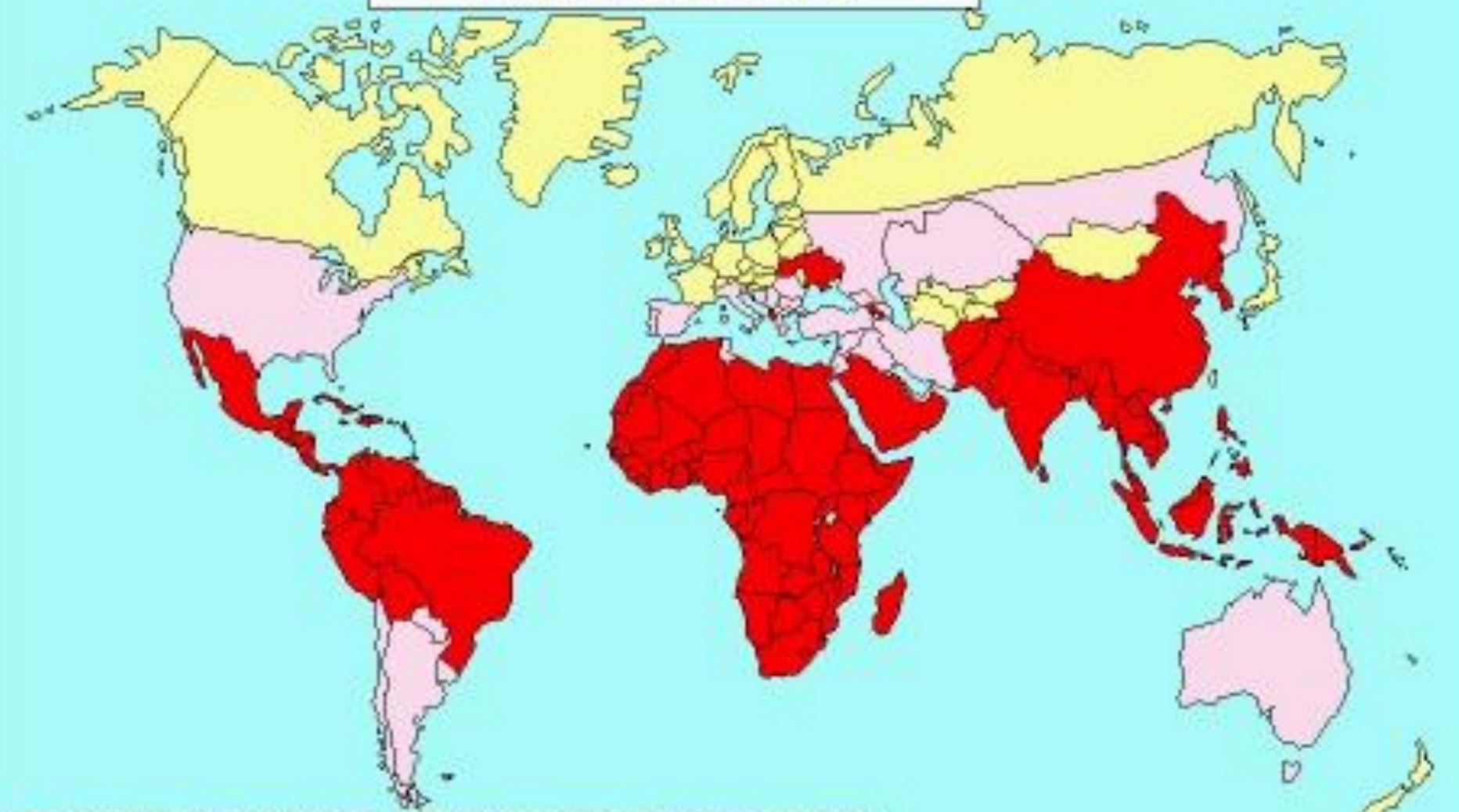
GIT: (nematoda):

- Albendazol, Mebendazol, Levamisol

Helminthic infections



Intestinal Helminths



■ Countries where intestinal helminths are a public health problem
■ Countries where intestinal helminths are transmitted

Source: WHO/CID, 1997

Schistosomiasis



Trematoda; Blood fluke

Snail fever, bilharziosis

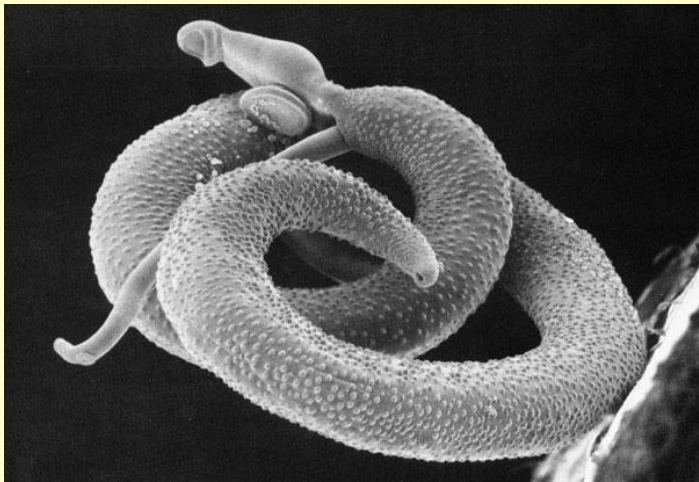
Distribution: tropical and subtropical countries

Most important helminthosis

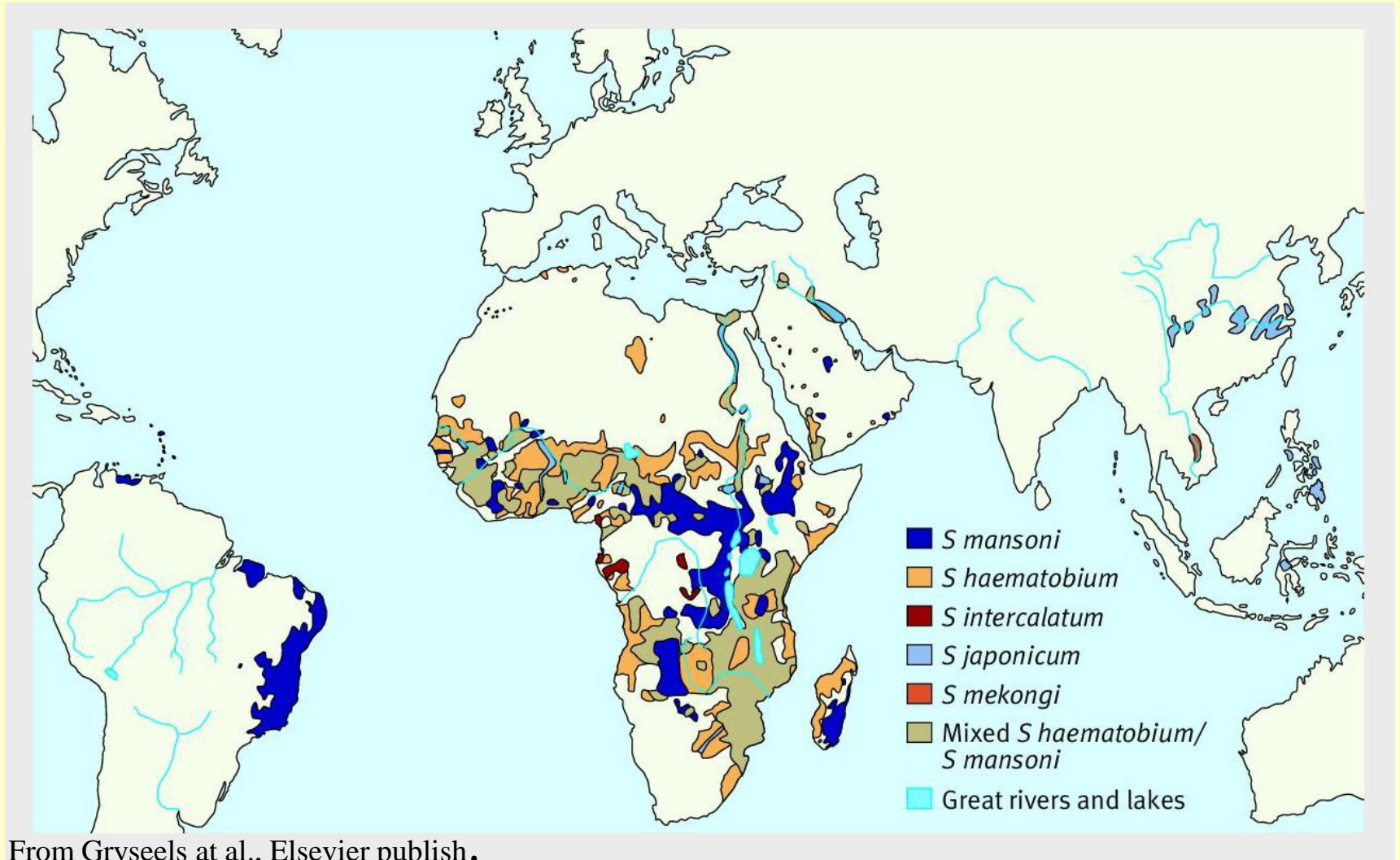
200 mil infected; 85% infections in Africa

600 mil at the risk of infection

Archeology : eggs of schistosomes in mummies from XX. dynasty (1250-1000 years BC)



Global distribution of schistosomiasis



From Gryseels et al., Elsevier publish.

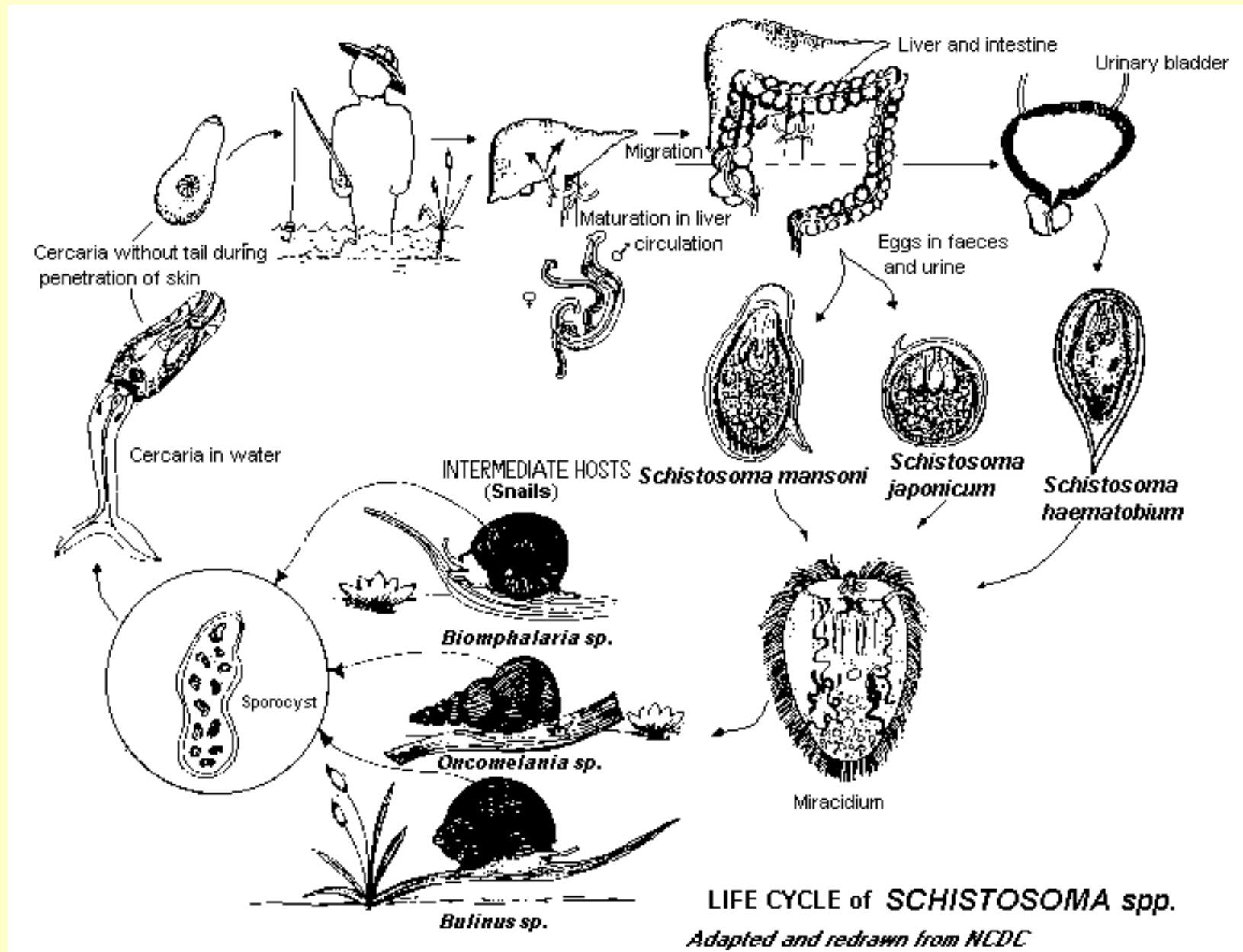
Complicated life cycle

Development in **one intermediate host** –
specific snail

Final host – **human**

Both sexes

Life cycle





For establishment of infection:

Fresh water lake/River

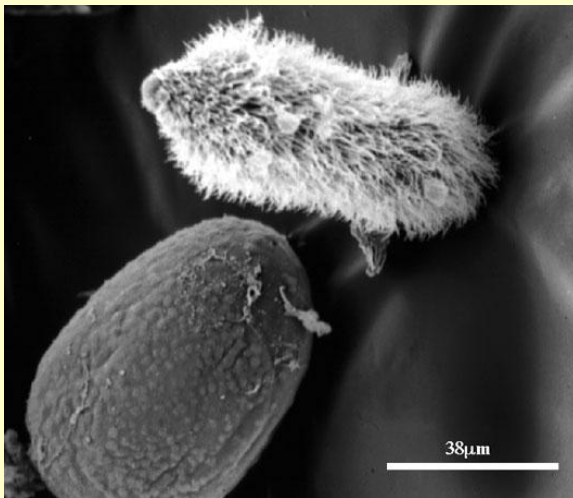
Final host discharging
live eggs

(water contaminated by human faeces)

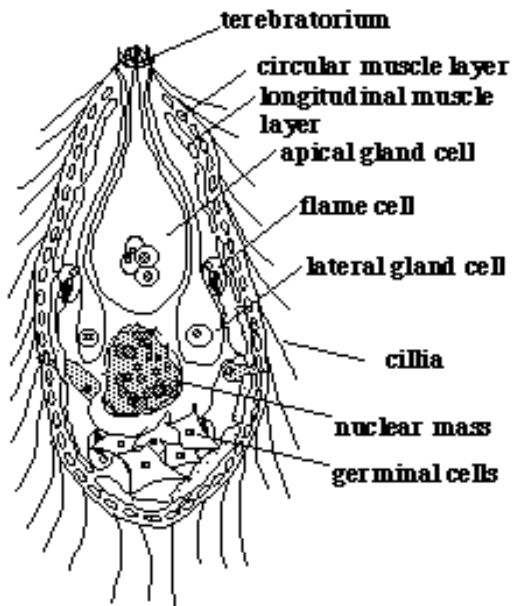
Presence of susceptible
Intermediate host

Miracidium

Larva released from the egg
Infective for specific aquatic snail

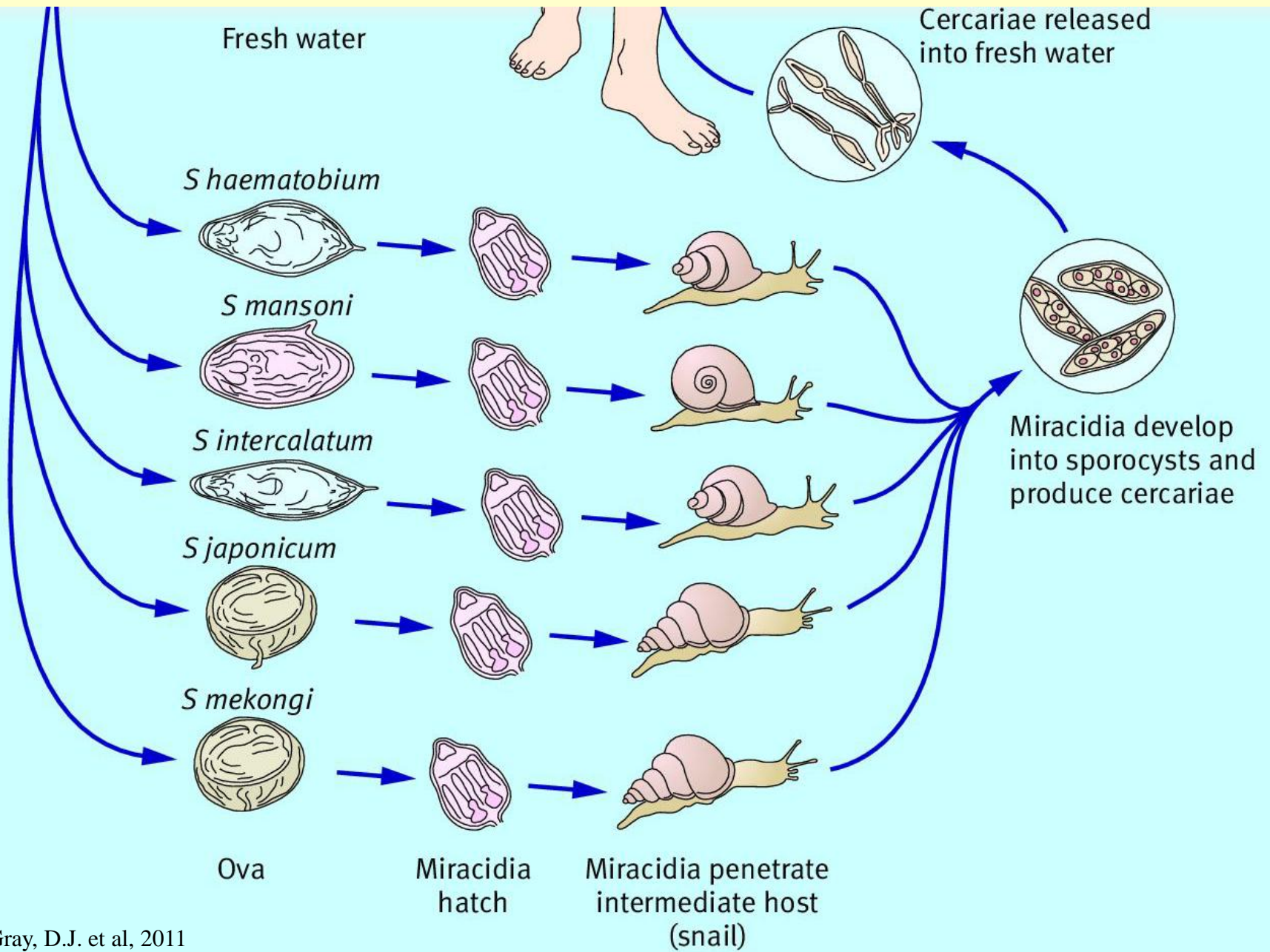


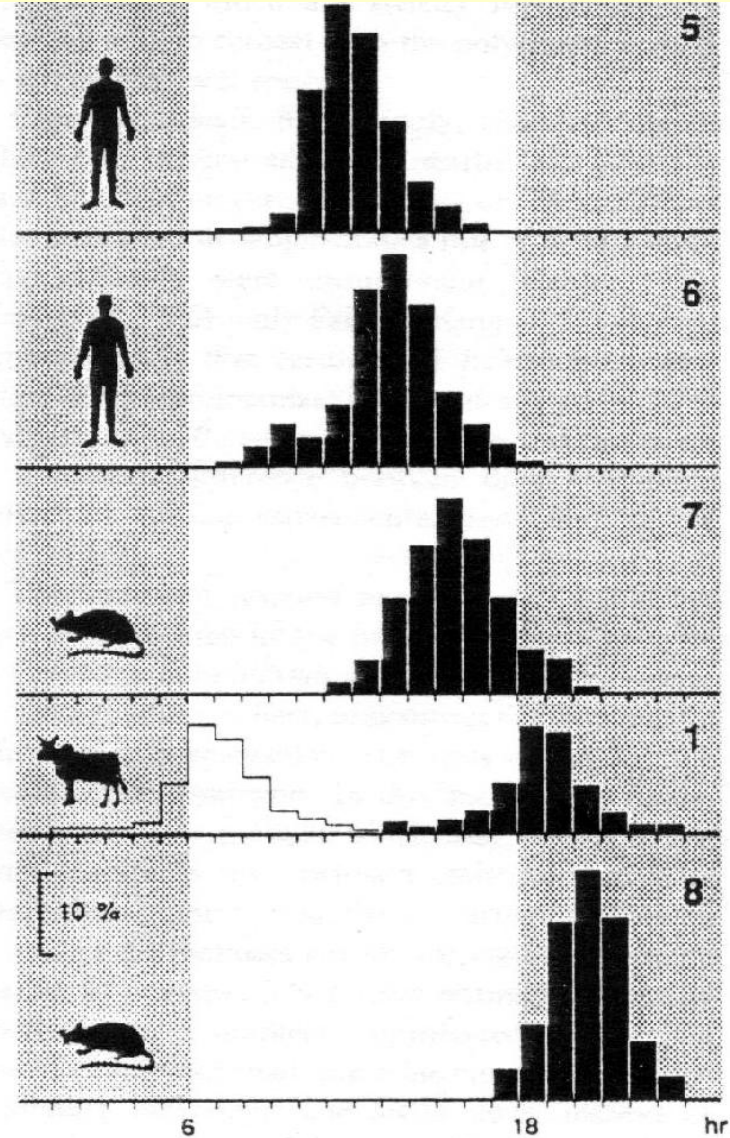
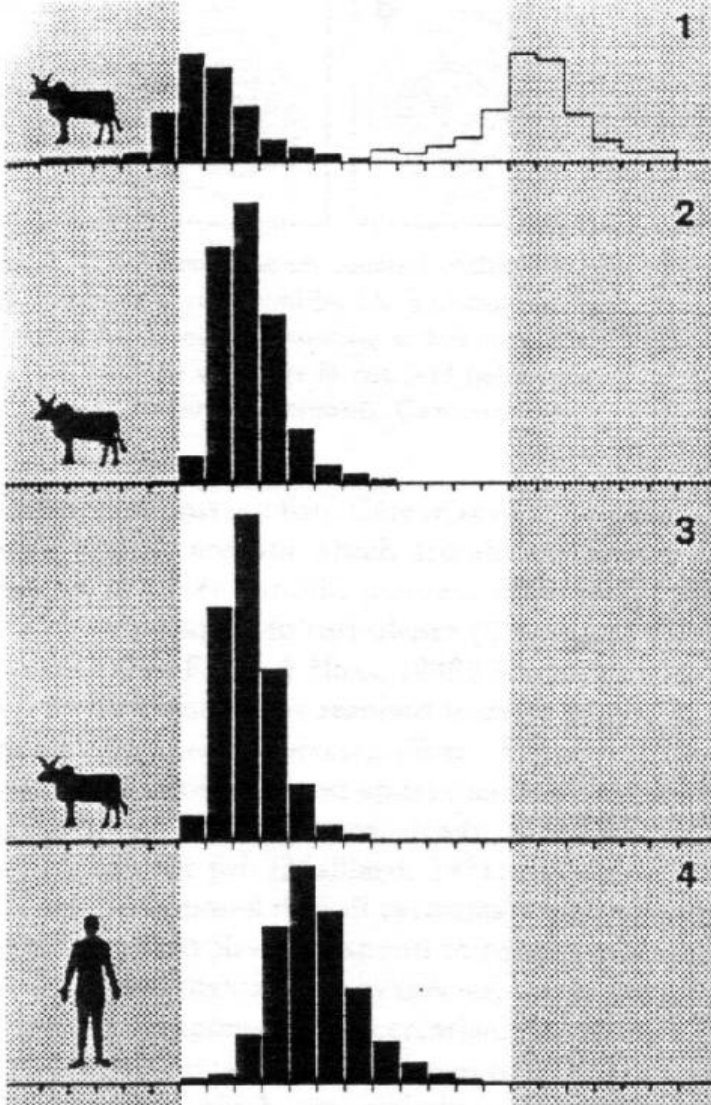
Morphology of Miracidium



Oncomelania spp. – intermediate host of *S. japonicum*







Cercaria

0,5 mm

Energy source: glykogen (24 hrs)

Actively searching for the host:

Arginine (*S. mansoni*)

Lipidic components of skin

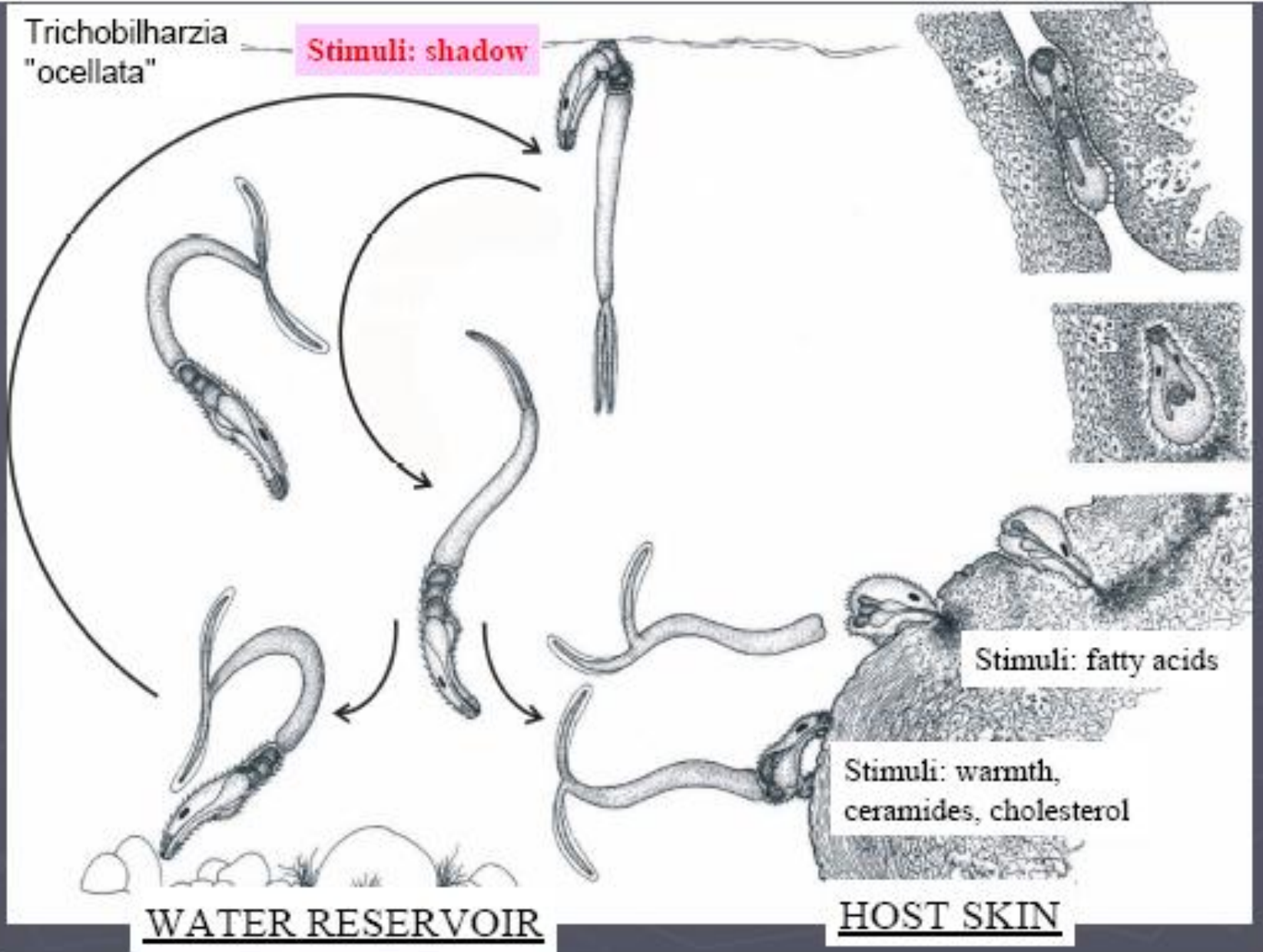
Temperature

Phototaxy

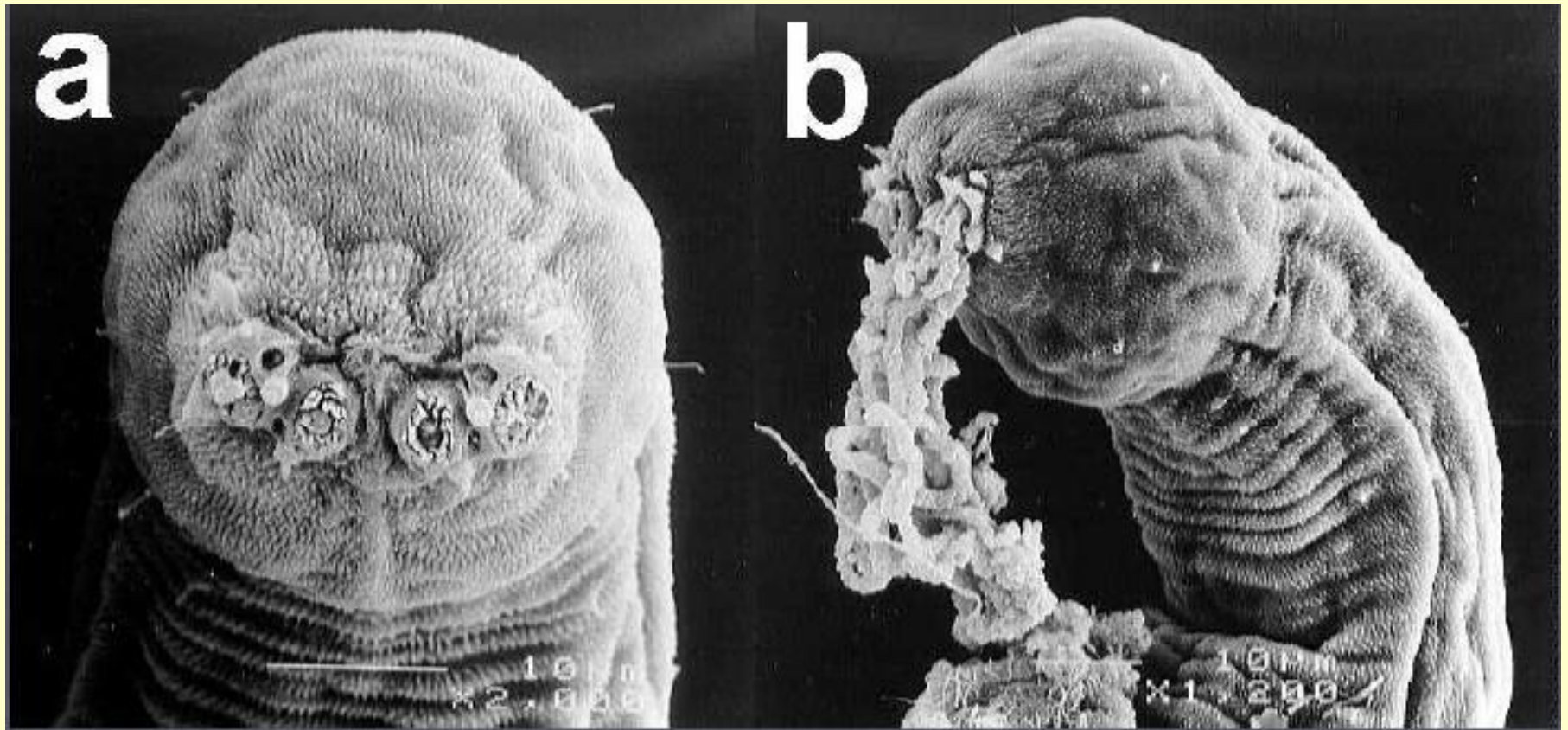


Trichobilharzia
"ocellata"

Stimuli: shadow



Penetration into the skin and through the skin is enabled by **secretion of proteases**: elastase, collagenase, hyaluronidase...



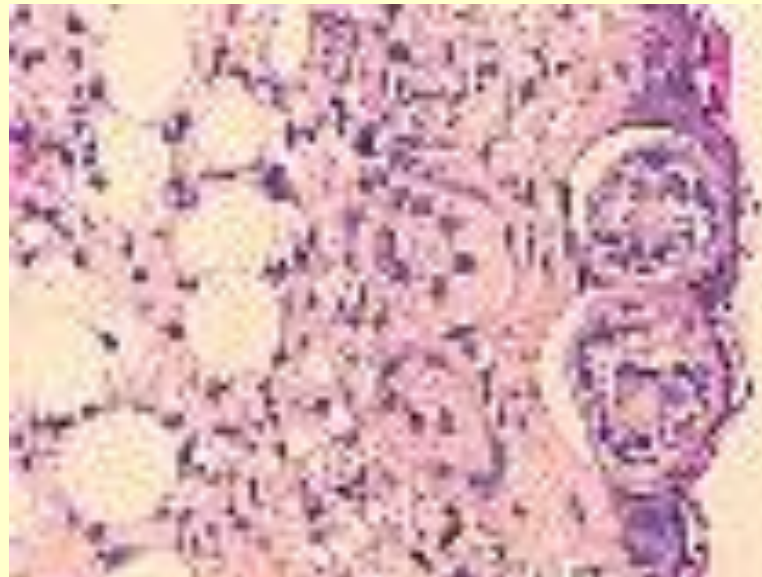
Formation of the **schistosomula** in the skin

Cercariae are differentiating to schistosomulas in the skin

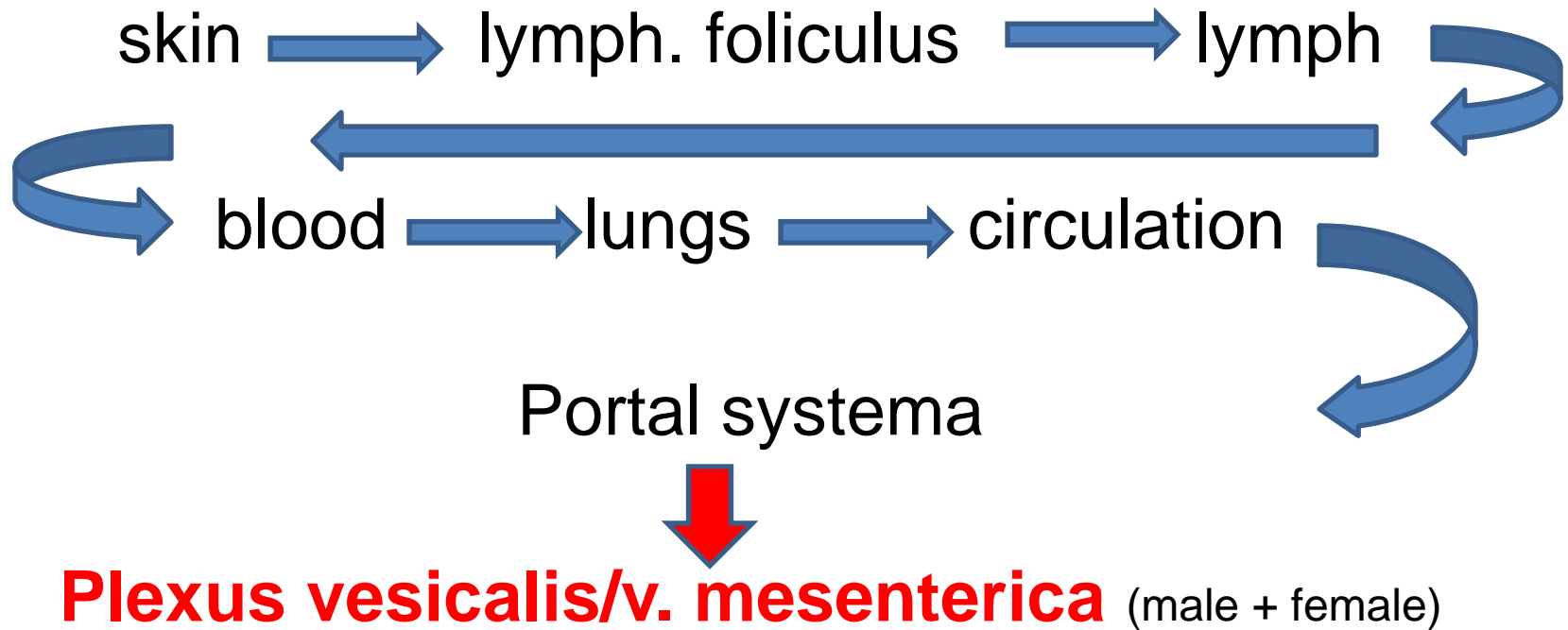
Masking with **host antigens**

Change of **metabolism** – aerobic to anaerobic

Deposition in skin appr. **2 days**



Schistosomula masked by host antigens and unrecognised by immunity migrates into the circulation



In blood – schistosomae absorb
erythrocytes and catabolise haemoglobin
(production of haemozoin)

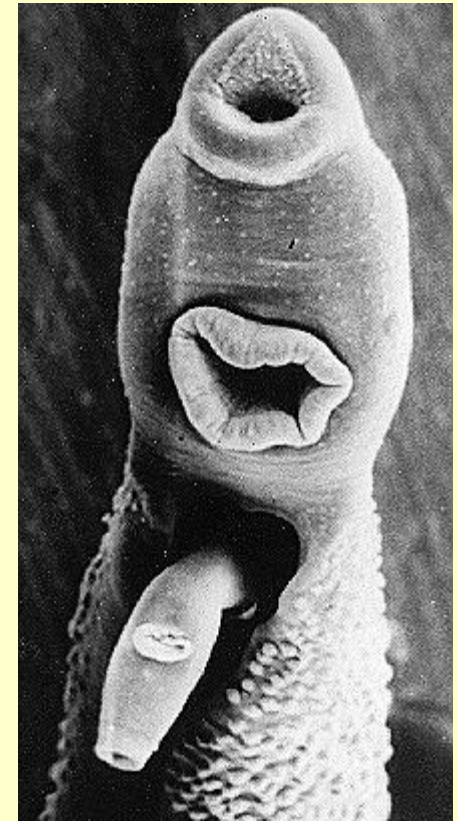
Copulation – deposition of eggs

(dilatation of terminal venules; 300-3000 eggs per day)

1. Antibody response – 50 days pi

Life expectancy: 3 years – *S. haematobium*

6 years – *S. mansoni*



虫

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♂

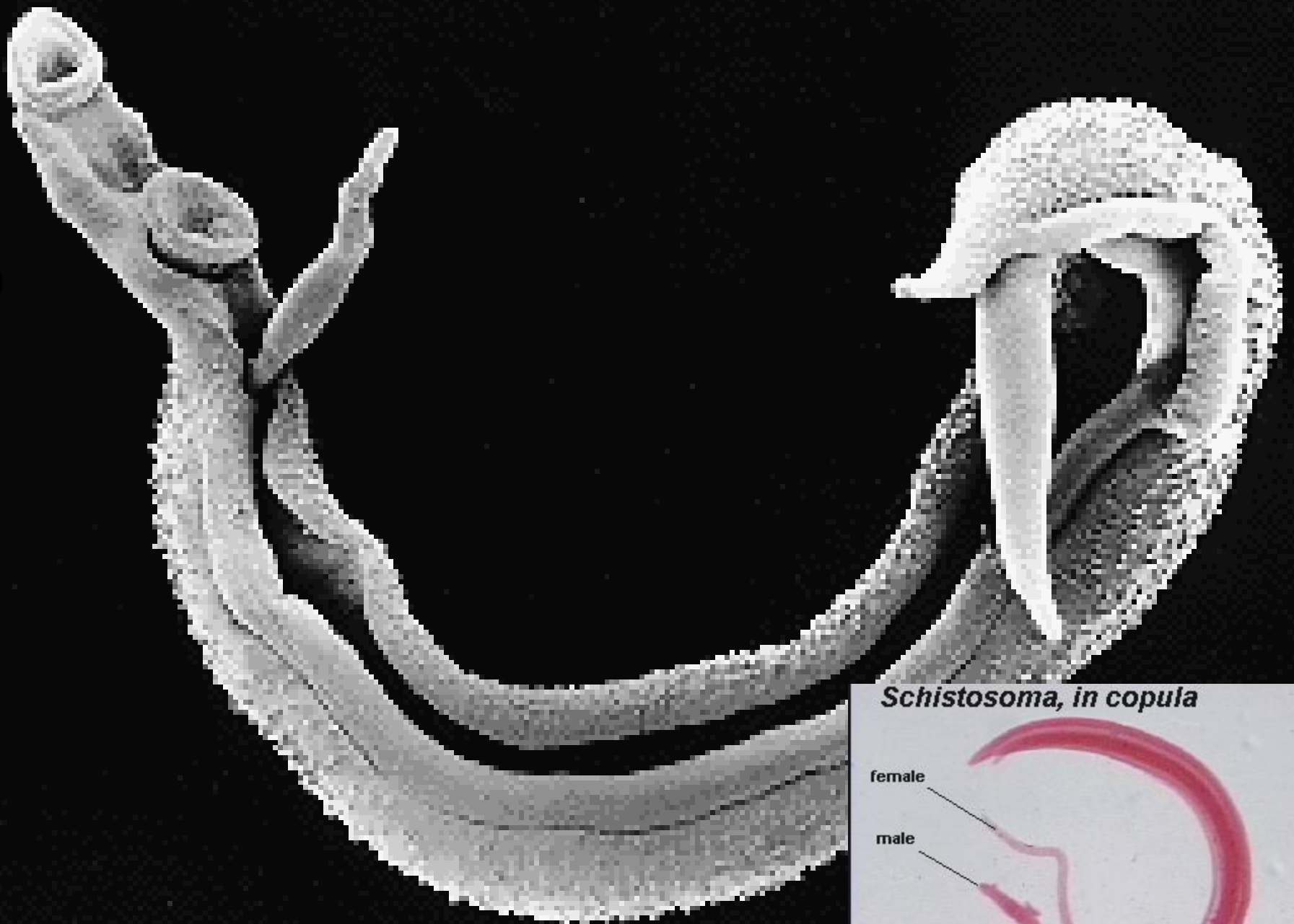
♂



♂



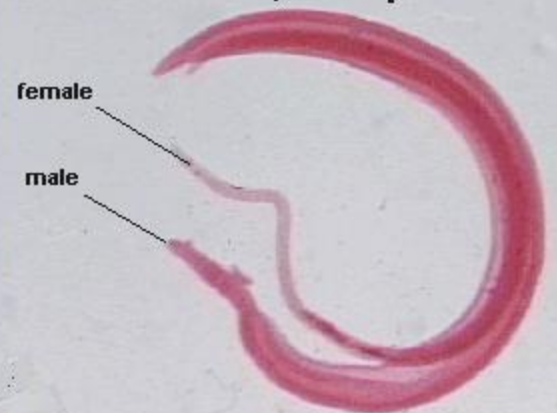
成虫



Schistosoma, in copula

female

male



Infection proceeds through few different phases

Skin infection

Cercaria dermatitis



Pulmonary

subacute pulmonary granulomatic schistosomiasis



Acute schistosomiasis



Chronic schistosomiasis

The course and symptomatology of the disease is dependent on several factors

Species of schistosoma

Number of parasites

Phase of the infection

Immune status of the host

Localisation of parasites

Dermatitis

Makulopapulous dermatitis

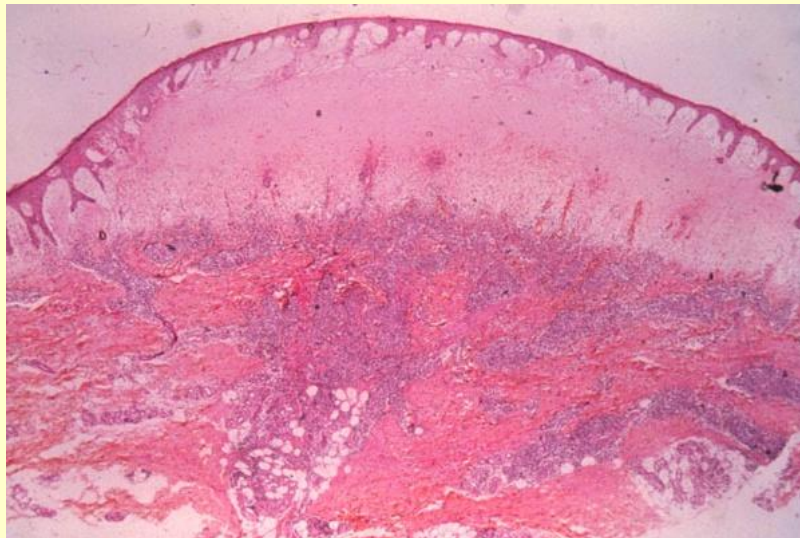
Urticaria within 12-48 hrs after contact with contaminated water

pruritus, oedemas, lymphadenopathy, temperature

Symptoms disappear without therapy within 14 days

Dg.: histology within 3 days pi, serology (50 days pi)

Cercarial dermatitis/swimmers itch



Pulmonary phase

7-14 days pi

Migration of schistosomulas – pulmonary infiltrates (not visible on X-Ray)

Dry cough, hemoptysis, temperature, chest pain, myalgia, diarrhoea, rash...

Bronchiolitis, interstitial pneumonia, thrombosis of pulmonary veins

Acute infection: toxic stage

Schistosoma japonicum

Katayama fever;

4-6 weeks pi (2-16 weeks)

Migration of schistosomulas



Hyperallergic reaction

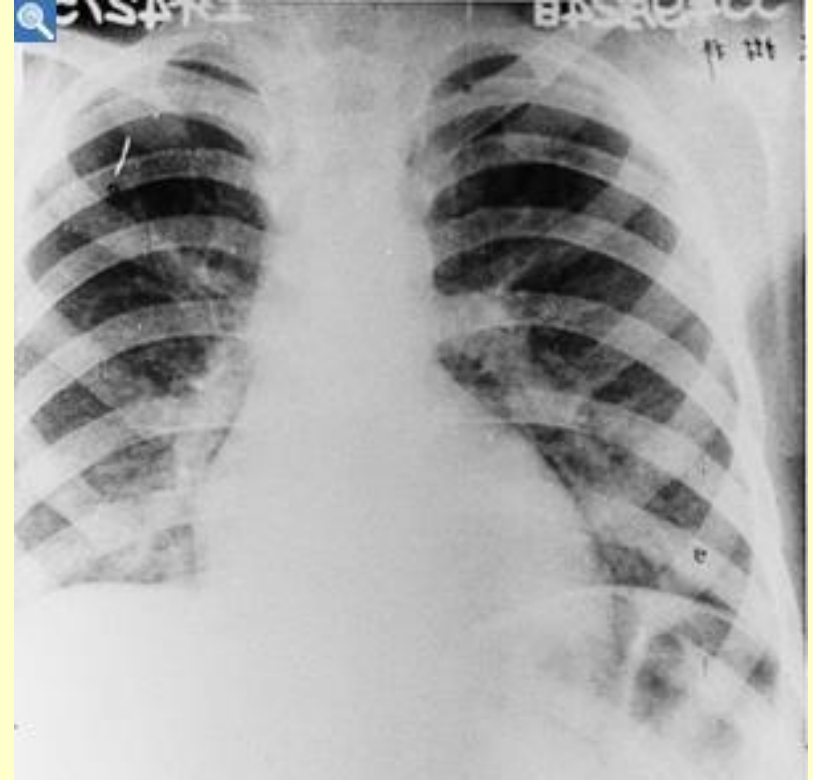
fever, tiredness, myalgia, headache, abdominal pain,
diarrhoea, urticaria, cutaneous oedemas

Hepatomegaly, splenomegaly

Generalized lymphadenopathy

Eosinophilia

Acute phase of the infection



Chronic phase: traumatic stage

- 3-6 months pi
- **Granulomatous reaction** around deposited eggs
- T lymphocytes activation
- symptomatology: asymptomatic vs **severe damage to the affected organs**

Affected organs

Intestine

Hepar

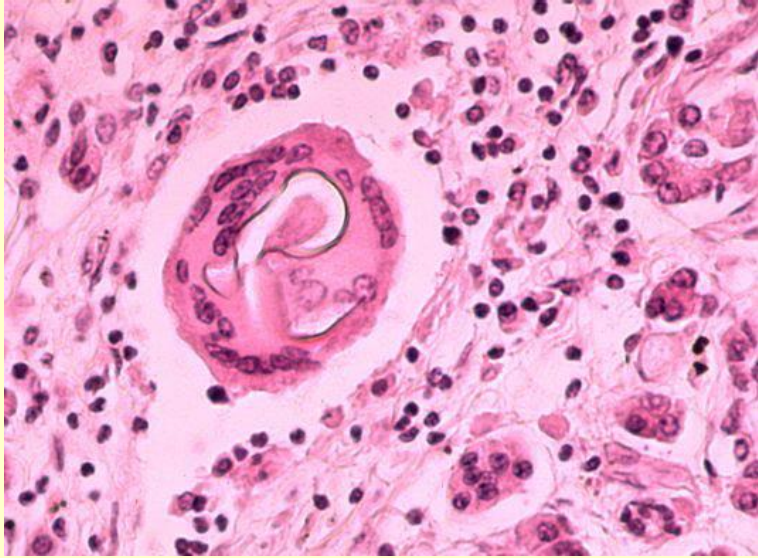
Lien

Ren/Urinary bladder

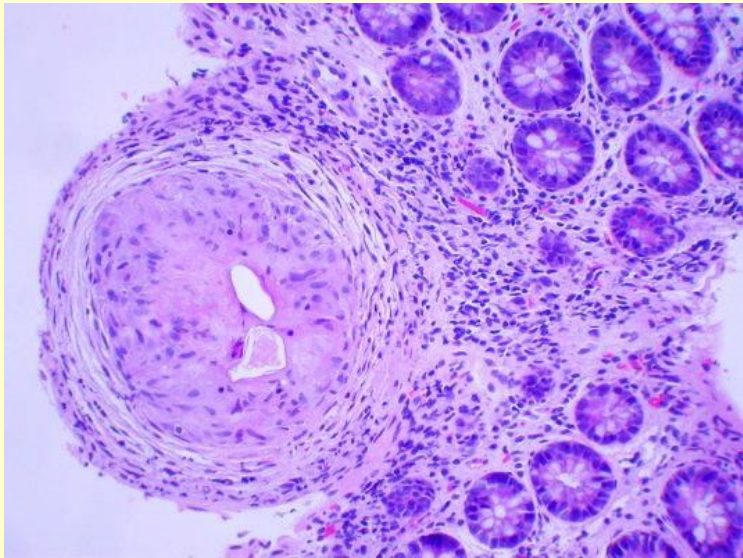
Lungs and heart

CNS

Reproductive organs

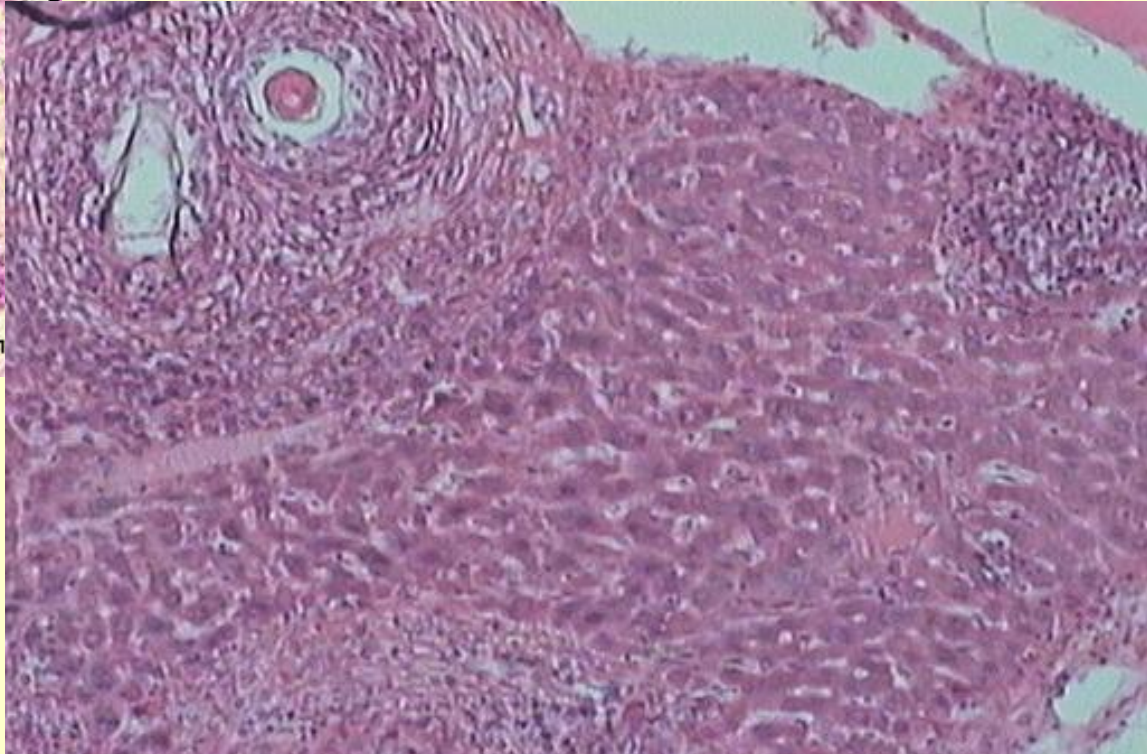
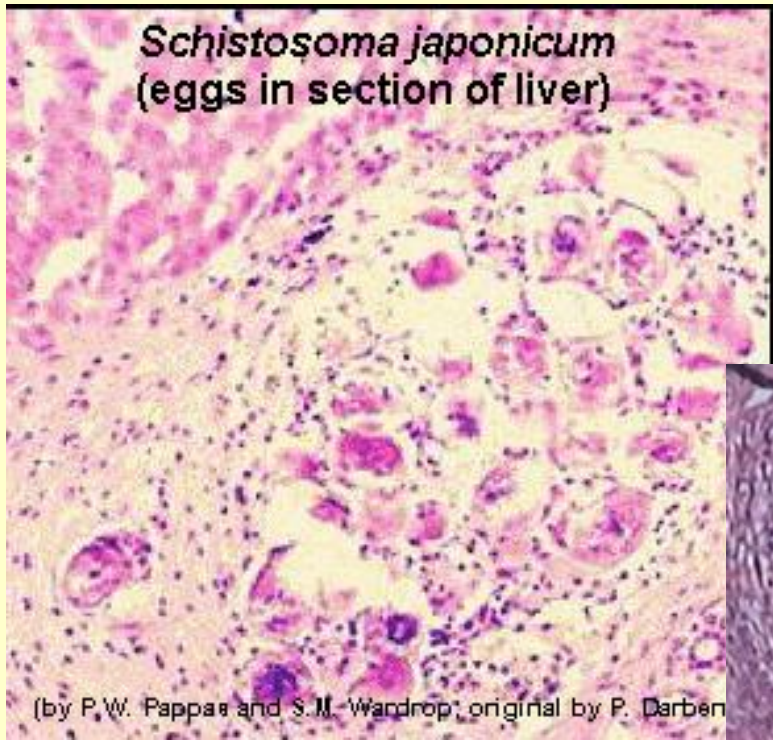


schistosomiasis of pancreas



schistosomiasis of colon

Schistosoma japonicum
(eggs in section of liver)



Granuloma formation in hepar

S. mansoni

Distribution:

Africa, Arabian peninsula, Brasilia, Surinam,
Venezuela, Portorico

Intermediate host: *Biomphalaria*

Final host: **human**; rarely infection of animals

S. mansoni deposited mainly in
V. mesenterica caudalis/inferior

colon, sigmoideum, rectum, hepar

Periportal fibrosis

adenomatous papiloma on mucous surfaces

portal hypertension; venostatic splenomegaly,
fibrosis of pancreas

Eggs discharged in the stool

Symptomatology

Abdominal pain

persistent **diarrhoea** (with blood)

anaemia

Polyp formation

hepato or splenomegaly,

Signs of portal hypertension

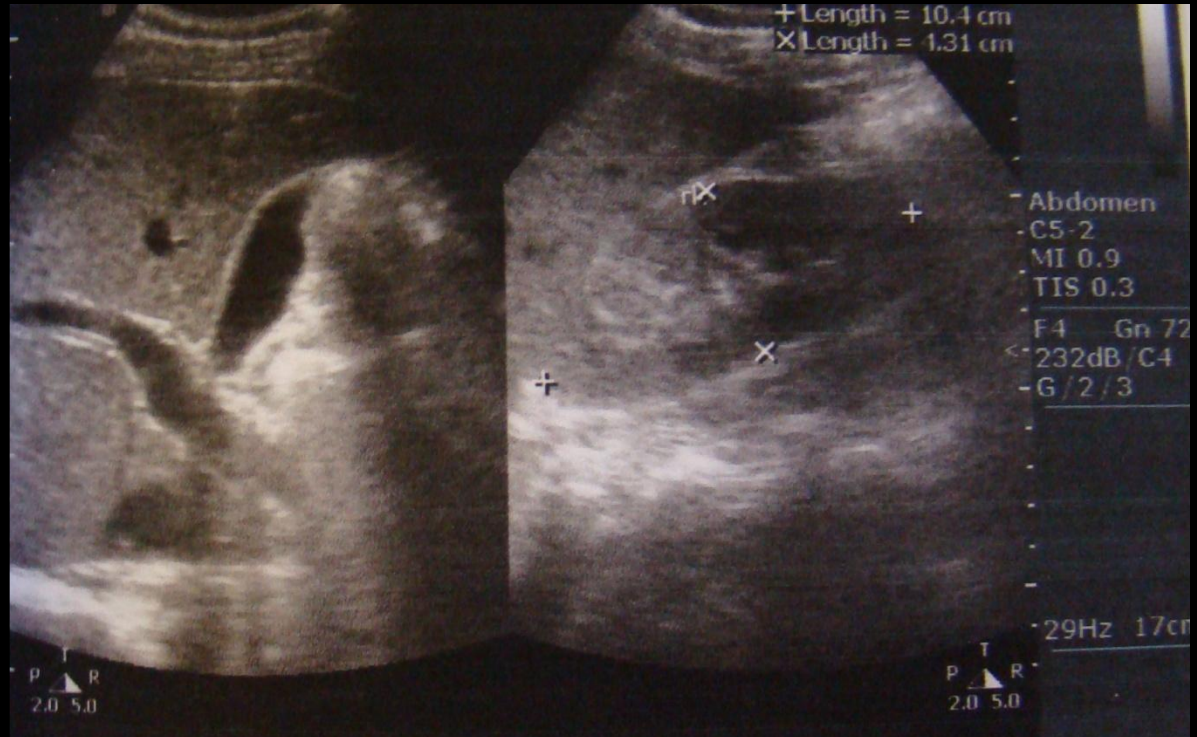
Infection of colon (*S. mansoni*)



Intestinal polyposis



Granuloma formation in hepar



S. japonicum

Distribution:

Taiwan, Japon, China, Indonesia, Philippines

Infective for almost **all species of mammals**

Intermediate host: *Oncomelania*

S. Japonicum is deposited mainly in
v. mesenterica cranialis/superior

v. mesenterica caudalis, vena portae

Shape of the eggs – possible deposition in
whole body, discharge mainly in stool

Affection of **hepar, intestine, portal
hypertension**

Symptomatology: similar to *S. mansoni* inf.

S. haematobium

Distribution:

Nile region, Africa, Asia, Cyprus, south Portugal, Jordan,

Intermediate host: *Bulinus*

Venous plexus vesical and pelvic

Polyposis of urinary bladder, hypertrophy of muscular layer; secondary bacterial infections

Symptomatology

Stenoses a dilatations of ureters

**papilomas, cysts, ulcerations, lithiasis of ren,
hydro a pyonephrosis** (eggs in renal parenchyma)

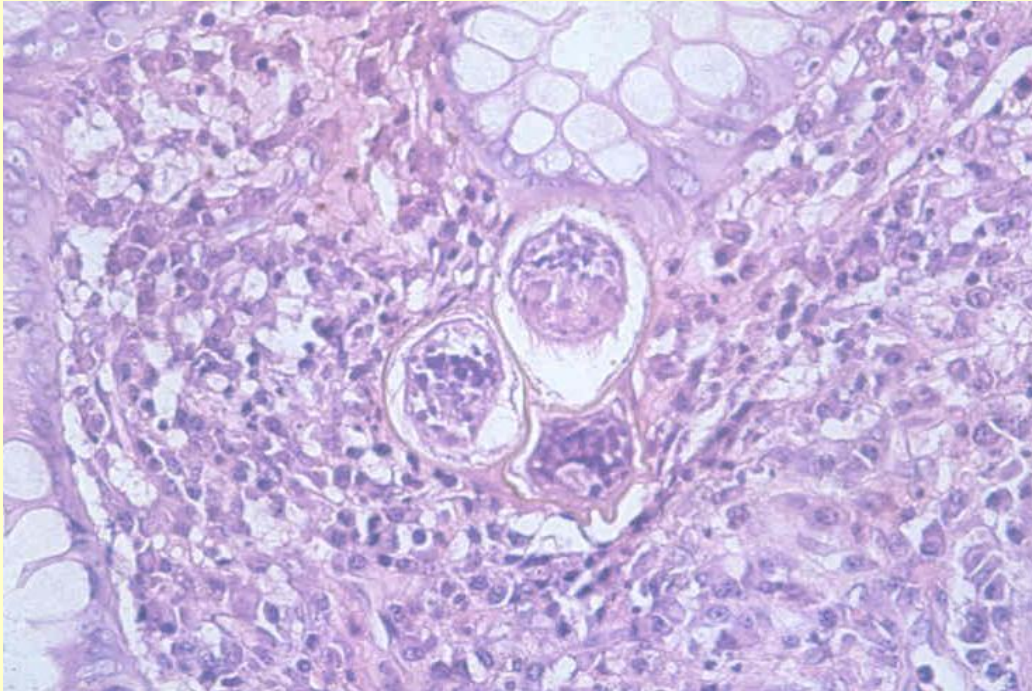
secondary bacterial infection

ca of urinary bladder

**Dysuria, polakisuria, haematuria,
eosinophiluria**

Impotence

S. haematobium



Granuloma in urinary bladder



hydronephrosis



Vesicoureteric reflux in ascending cystography



Bladder calcification in plain x-ray

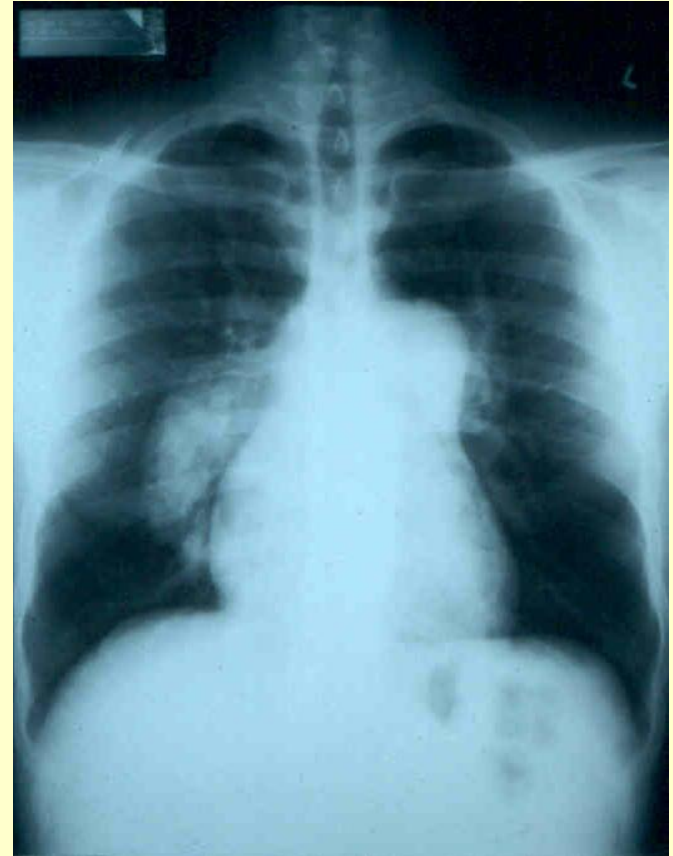
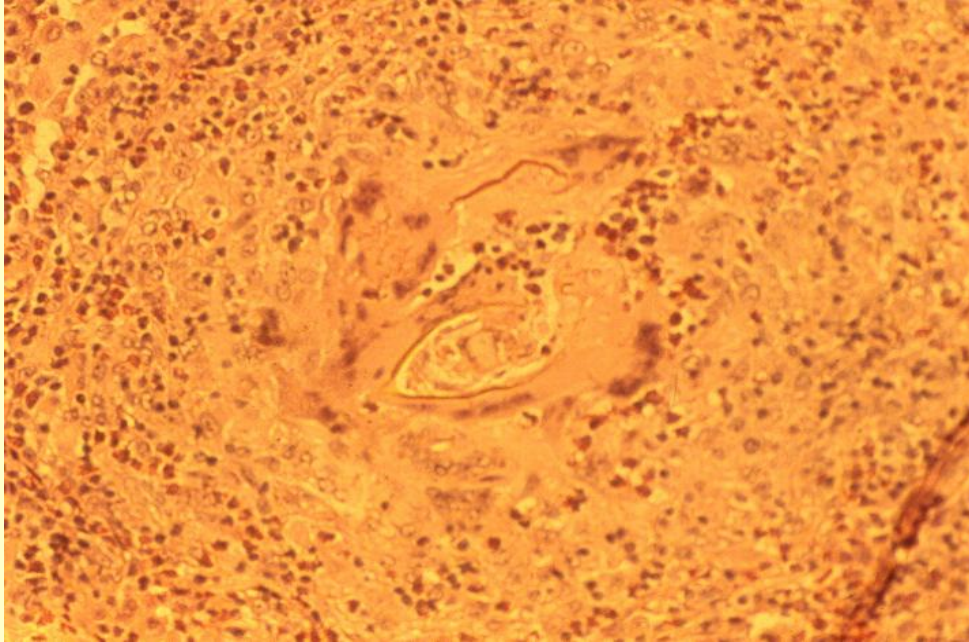
Eggs can be deposited by blood flow to different organs

Lungs: pulmonary hypertension

Joints: arthropathy

Brain: epilepsy, headache, vomiting, visual disturbances

Cor pulmonale, infection with *S. japonicum*

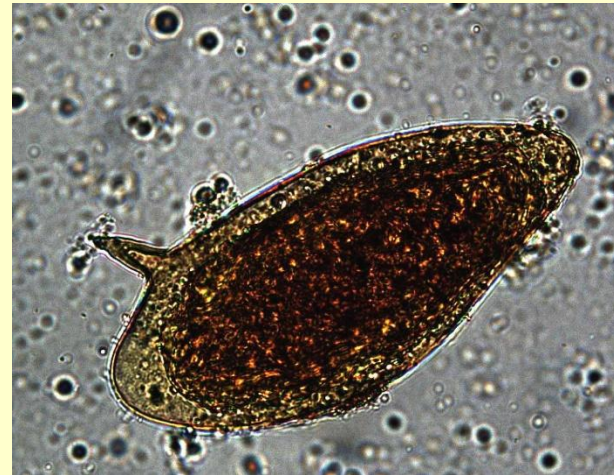
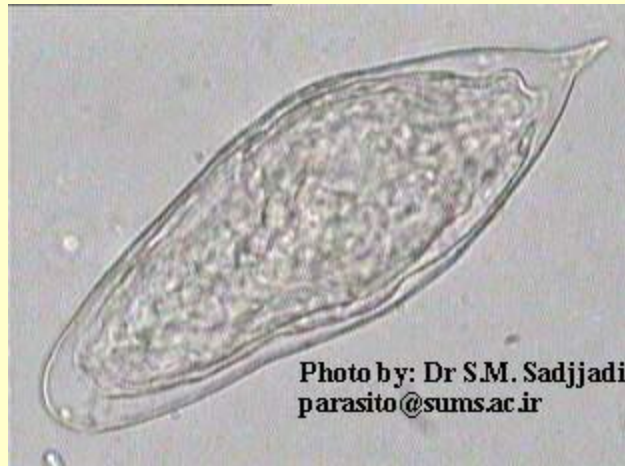


Diagnositics

Direct proof of eggs presence in stool, urine, biopsy

Serology (indirect haematogluttination, ELISA)

The test of viability of miracidia



Therapy

- **Praziquantel:**

- Weak infection: 1 dose po 40 mg/kg
- Severe infection: 2 x 60 mg/kg a 4-6 hrs after meal
- Infection with *S japonicum* 3x á 4 hrs 20mg/kg

- **Niridazol**

- 25mg/kg/day in three doses 7-10 days in combination with diazepam

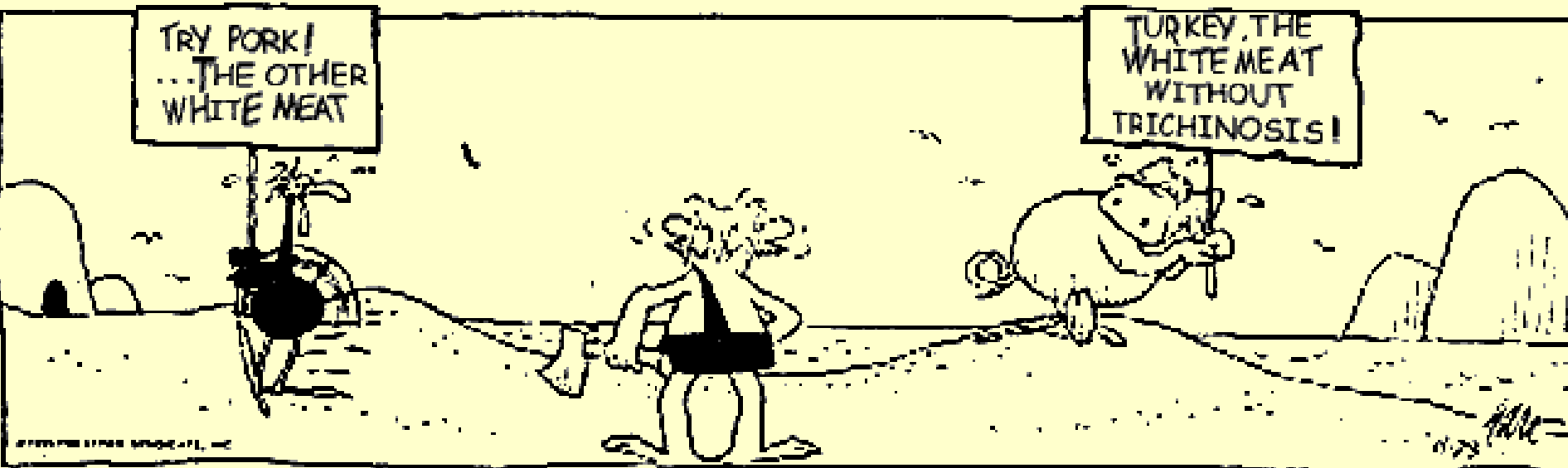
- **Metrifonate**

- Only *S. haematobium* 7,5-10 mg/kg single dose; repeat in three weeks

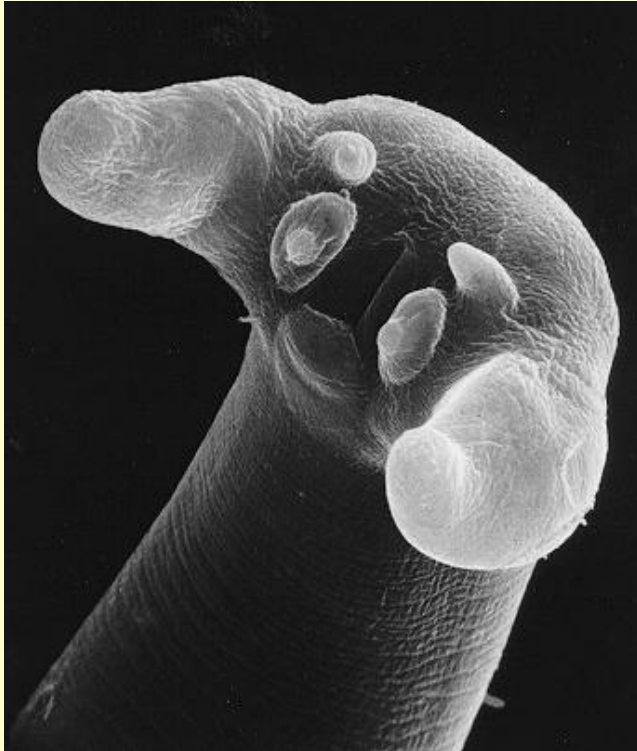
Trichinella spiralis

B. C.

BY JOHNNY HART



© 1978 THE NEW YORK TIMES MAGAZINE, INC.



Nematoda

Distribution: cosmopolite

Transmission:

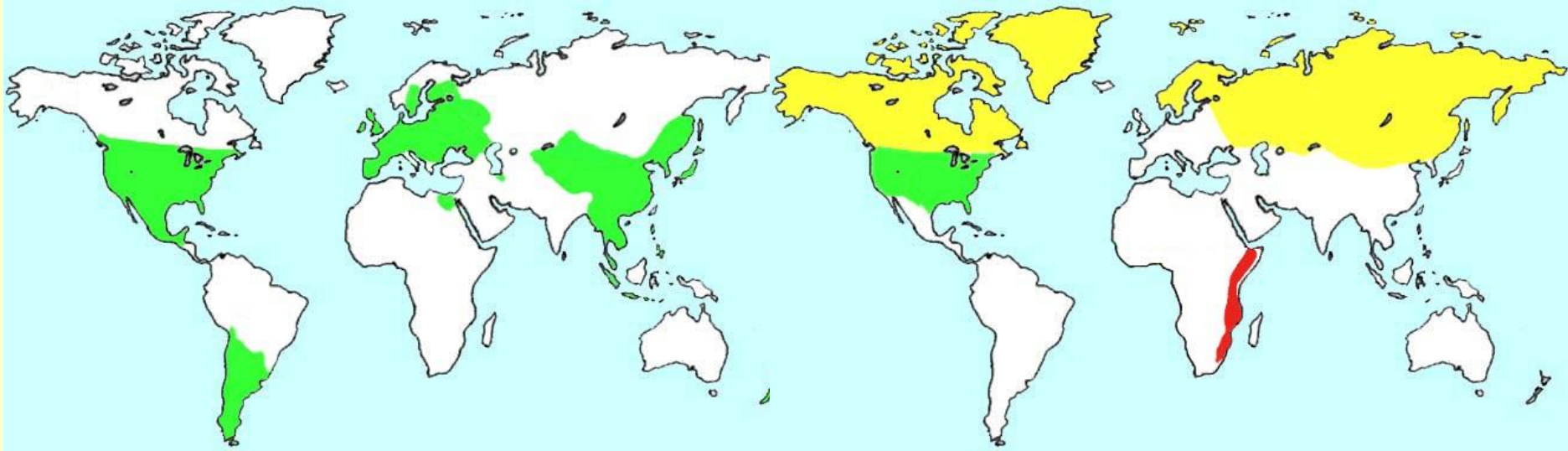
alimentary

(consumation of **undercook
meat** infected by larvae)

Host: human, swine, bear,
wild boar

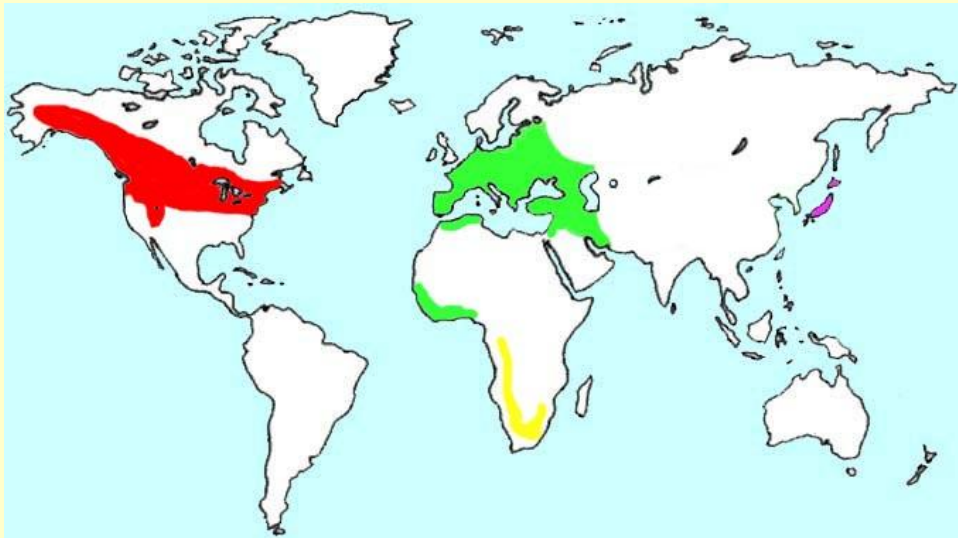
8 species of trichinella

CR - *T. brittovi*

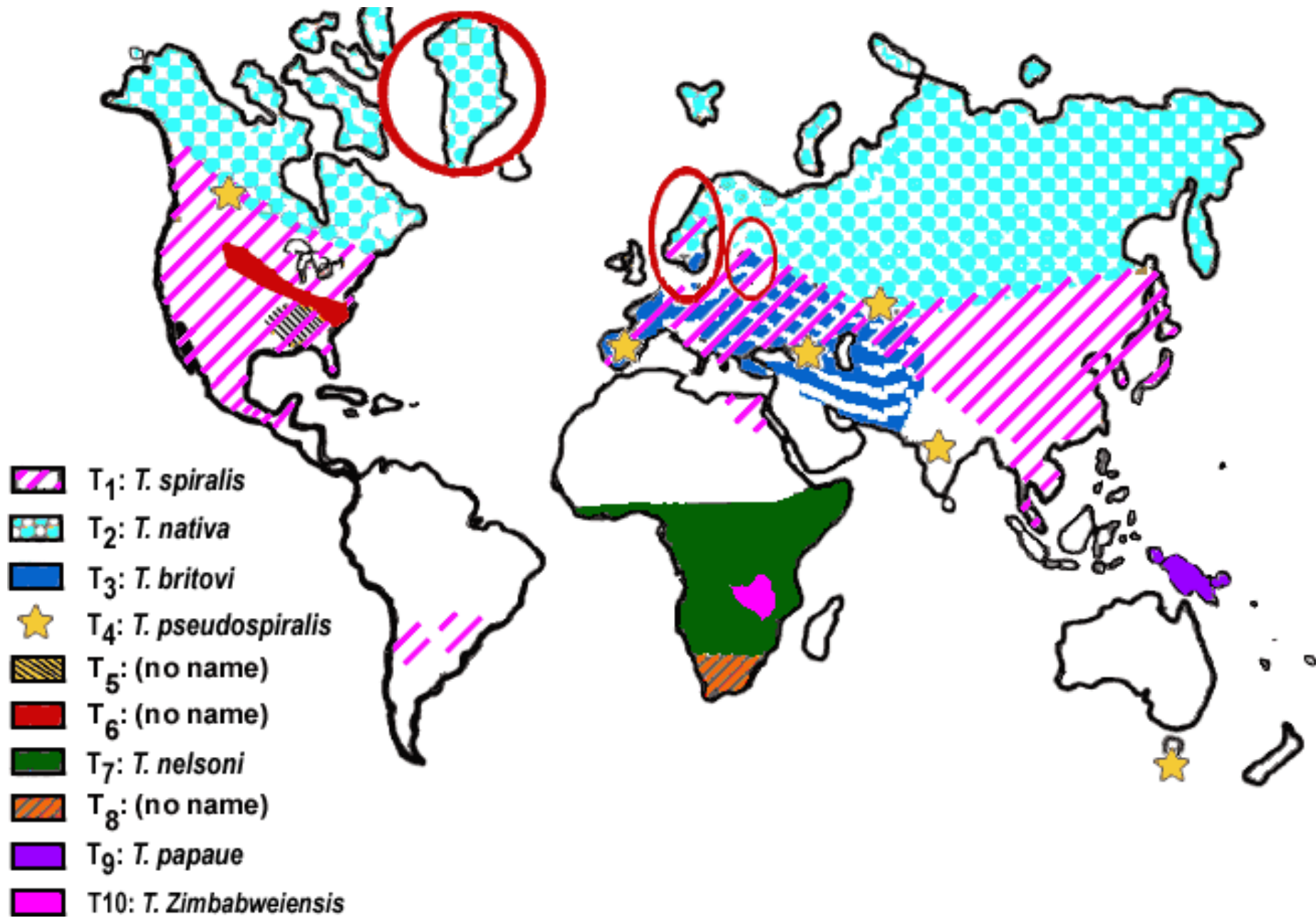


T. spiralis

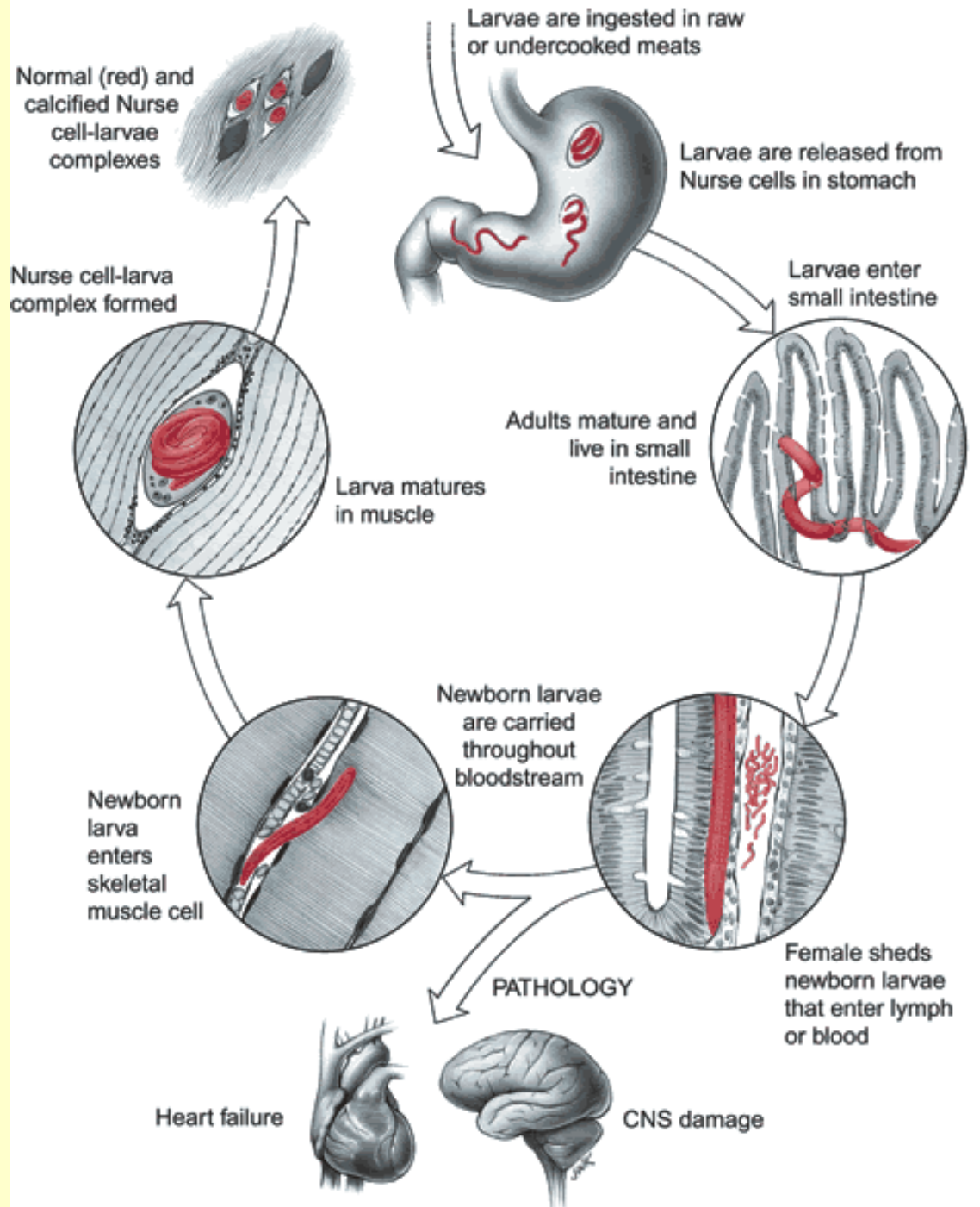
T. nativa

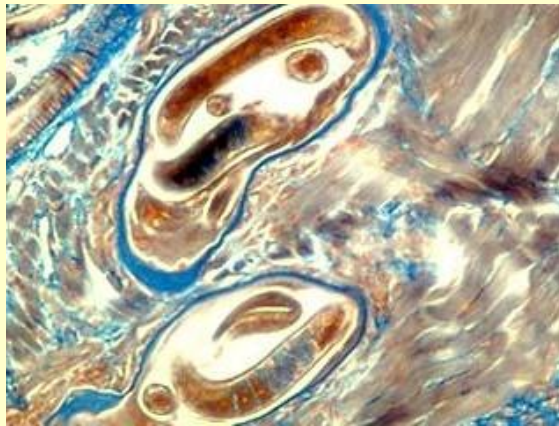


T. spp.; T. britovi



Life cycle





Intestinal phase –
Maturation of larvae about 30 hrs

Females are depositing larvae:

Appr. 4 days pi;

1000-1500 larvae/1,5 month

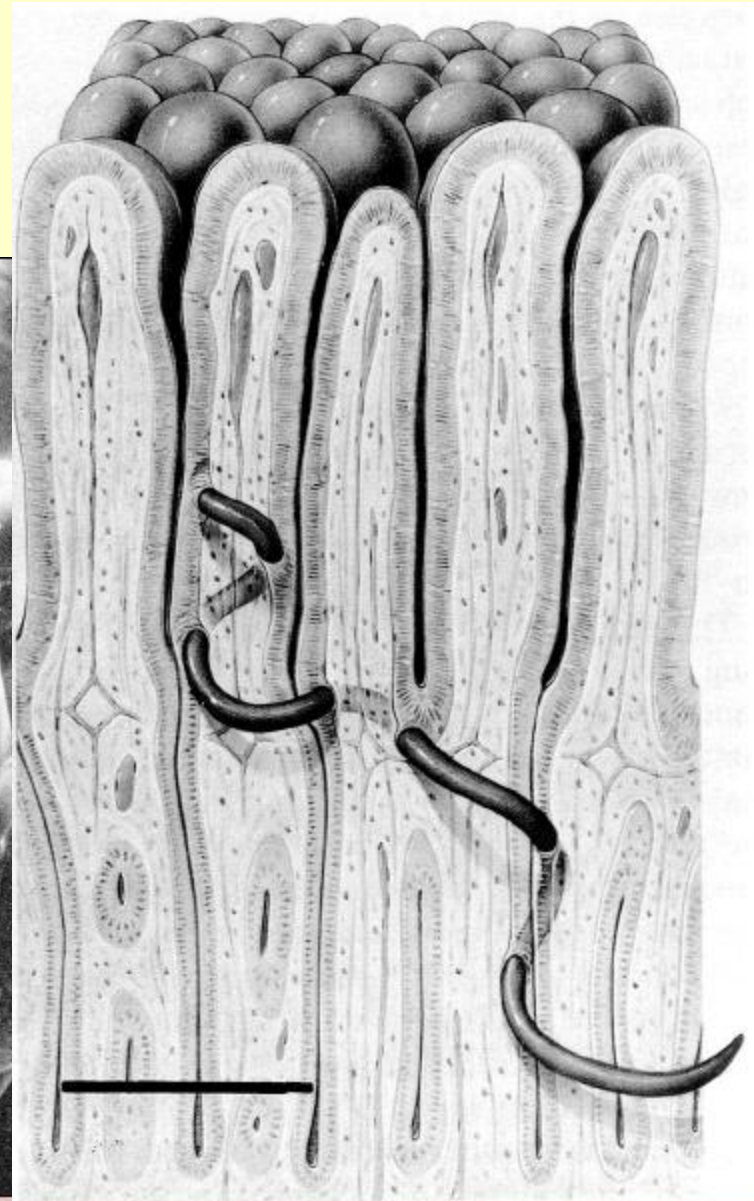
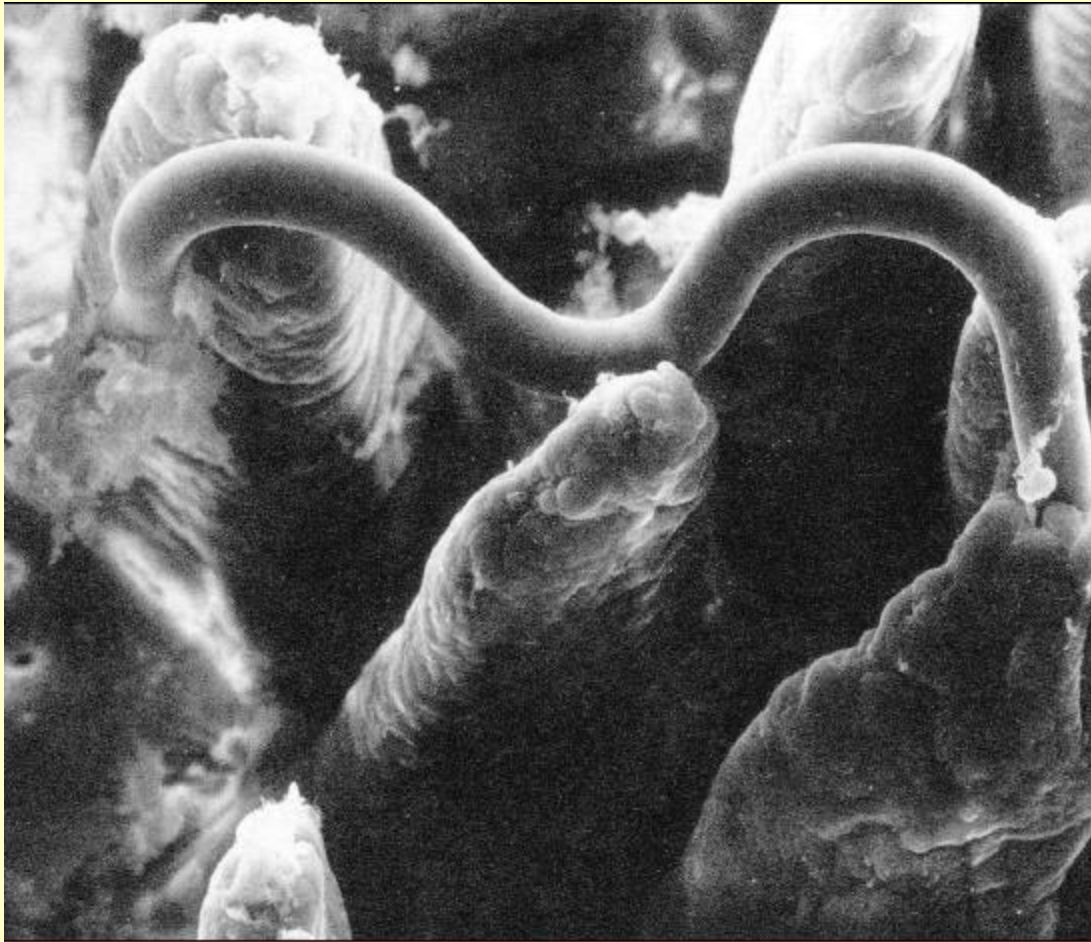
Migration into the muscle –2-3 days
(affection of myocardium – damage to the heart
function)

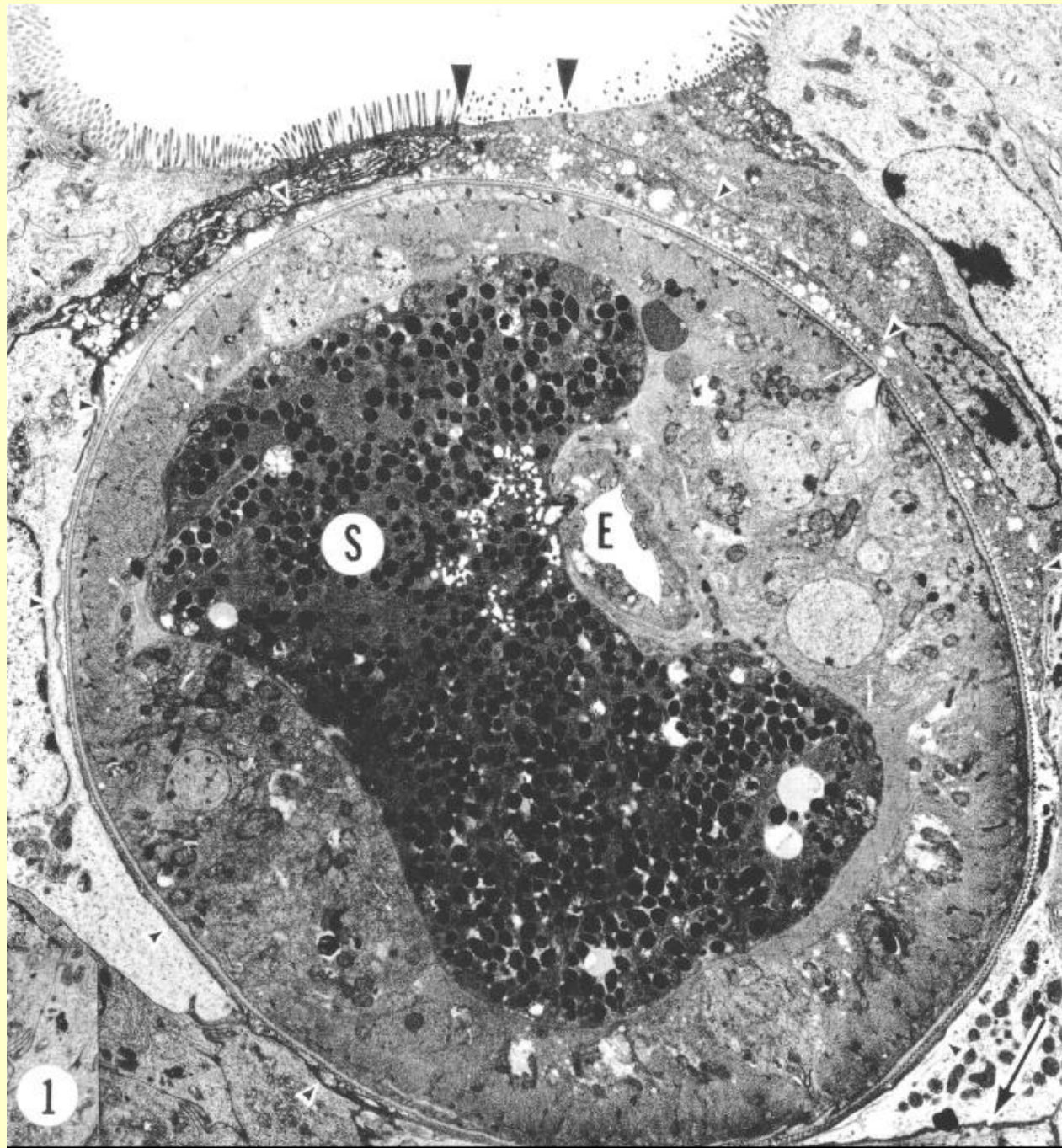
Calcification of larvae – after 6-12
months

Trichinella is **intracellular parasite**

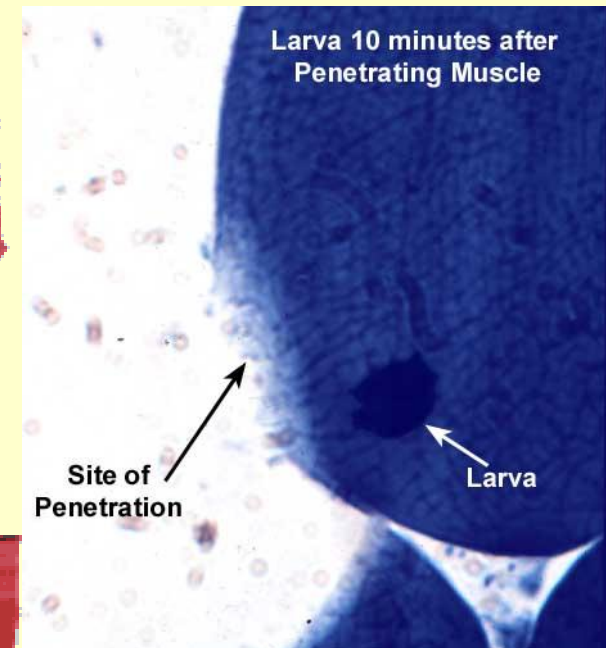
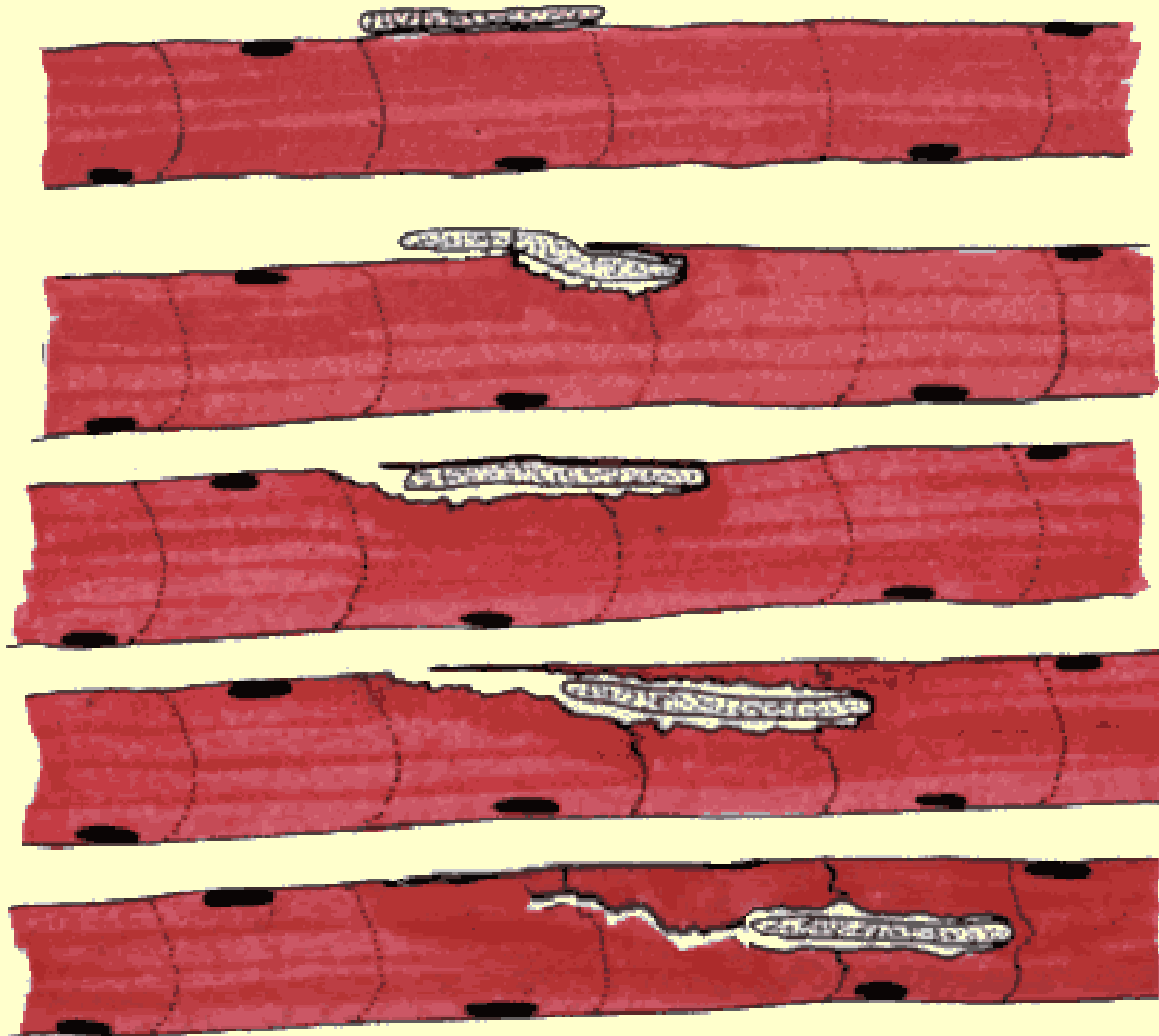
Intestinal phase – formation of
tunels in enterocytes

Female 5 mm, male 2 mm

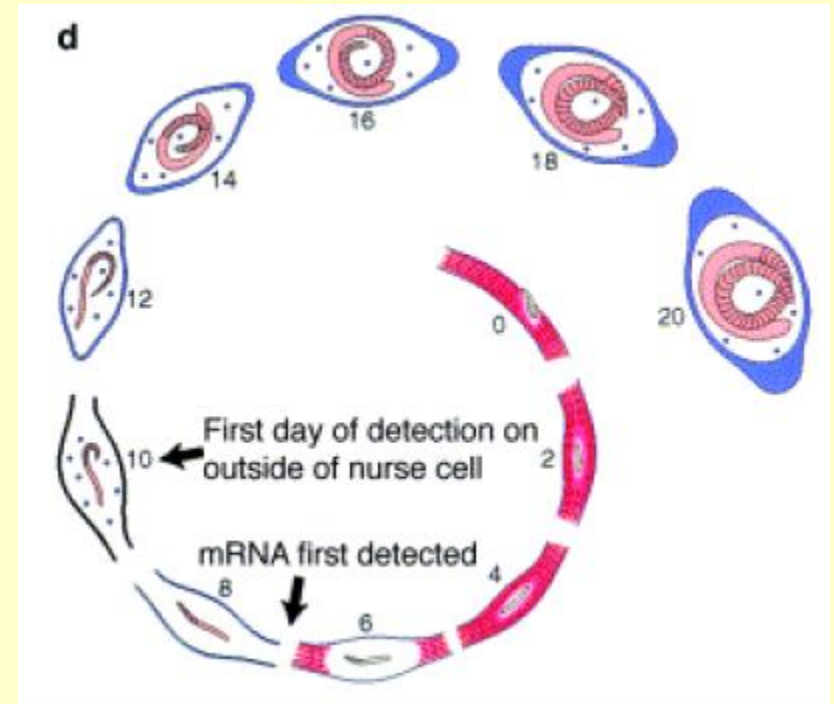
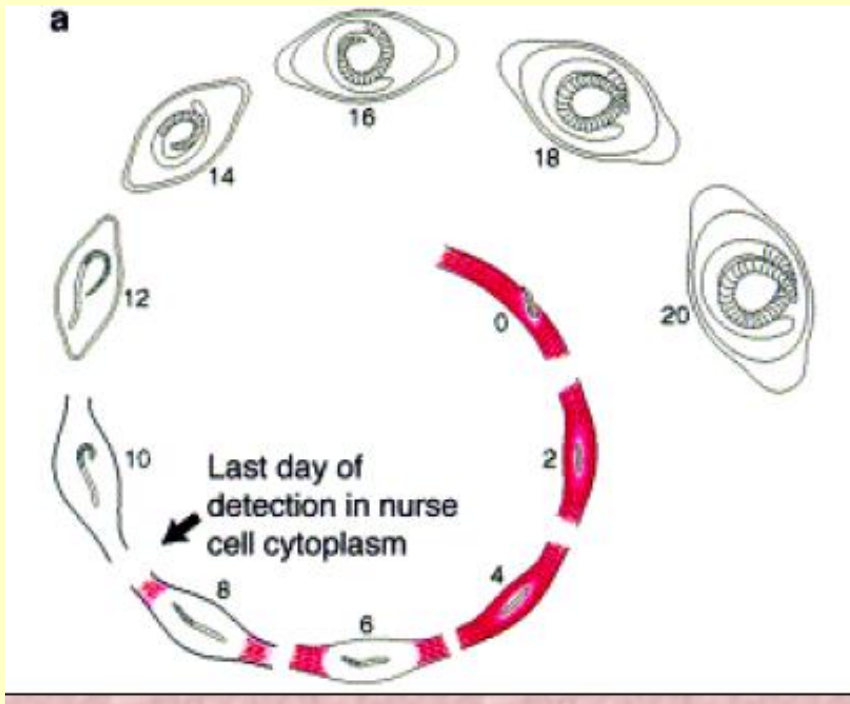




Invasion of muscle cells

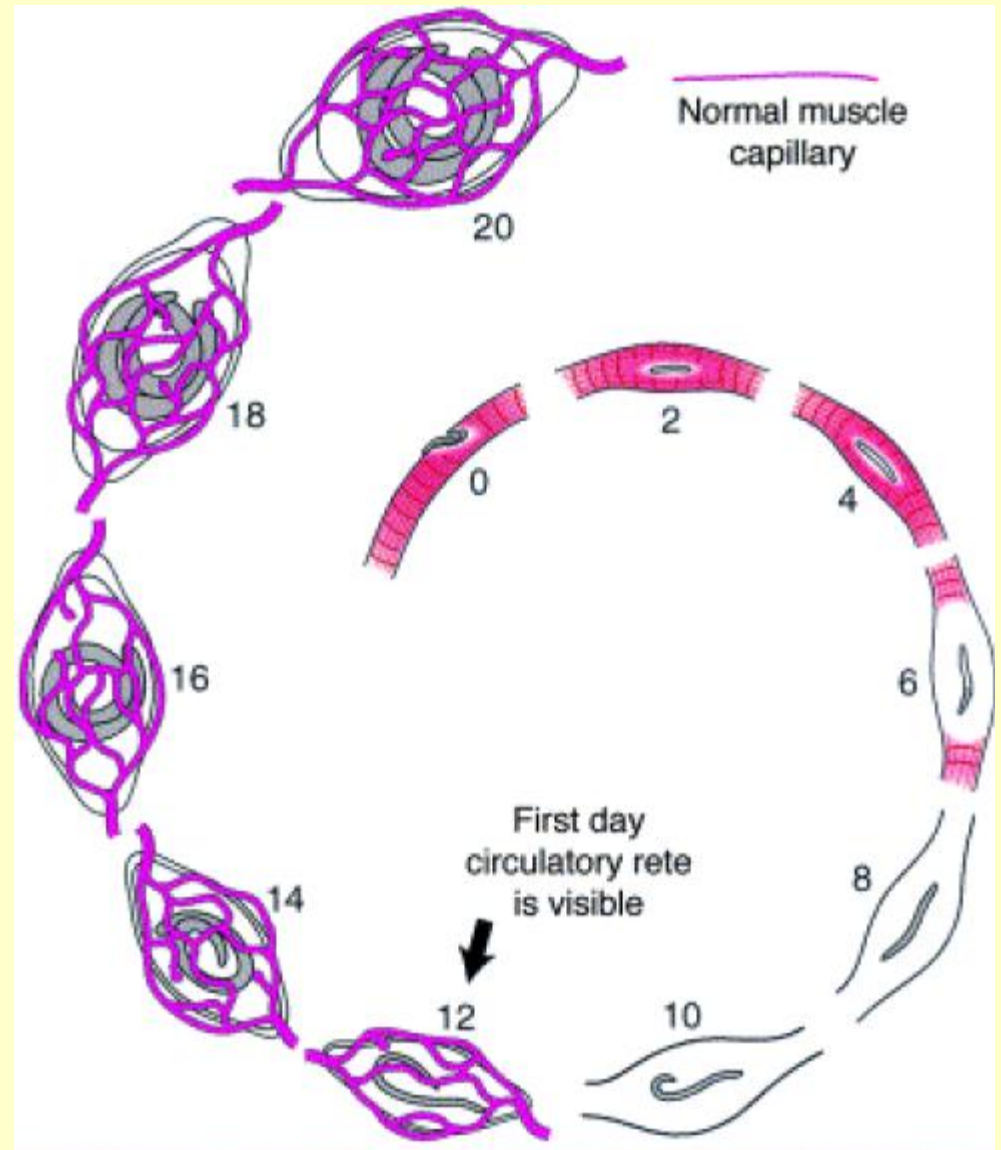
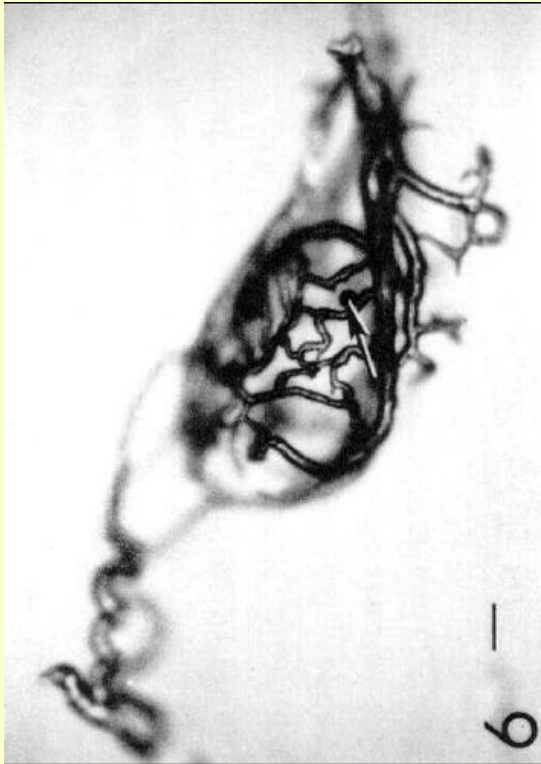


Infection of the muscle cells leads to changes in their function and composition induced by *Trichinella*



Loss of contractile elements and formation of collagenic capsula

Induced angiogenesis in infected cell



Symptomatology

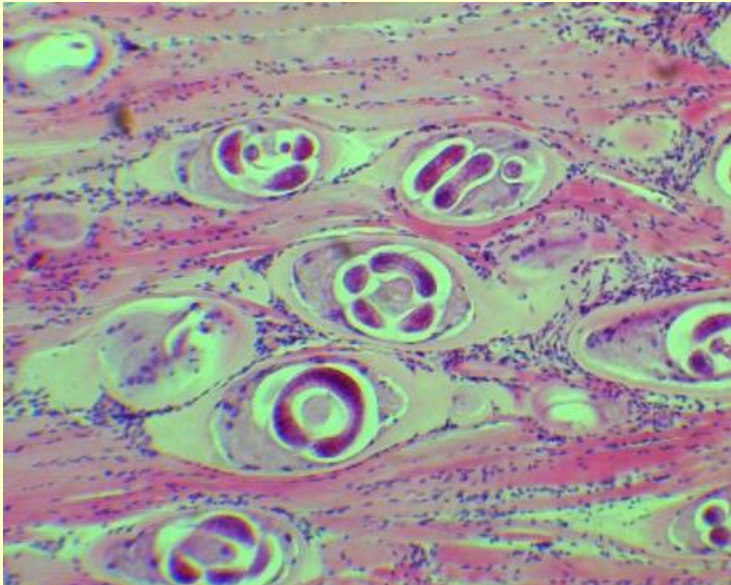
IP: 5-25 days

Intestinal phase:

2-10 days pi;

vomiting

diarrhoea



Muscular phase:

fever (40°C),

myalgia

weakness

tiredness

periorbital edema (80-100%)

cefalea,

konjunktivitis,

Oedemas of limbs

maculopapul. exantema (20-50%)

Lab.: eosinophilia, IgE,

↑ CK, LDH, AST



Periorbital oedema; konjunktivitis



Subungual haemorrhagie



Periorbitel oedema



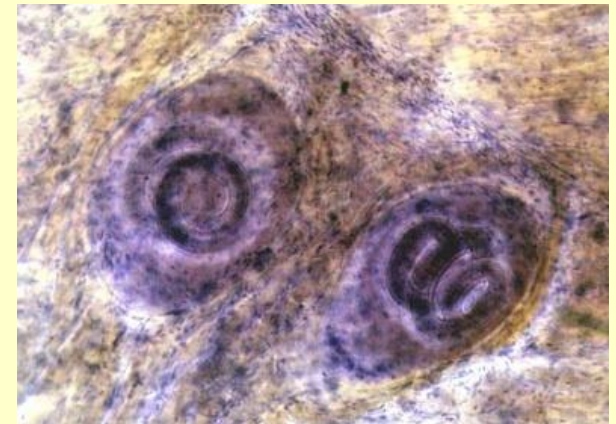
Diagnosics of the infection mainly by serology

Serology: antibodies against E/S antigen

Biopsy

PCR

Veterinary control: trichinoscopy



Therapy

Tiabendazolum

- 25-50 mg/kg/day in 2 doses (max. 3 g/day)
- Within 1 week after infection, affects the adults

Albendazolum

- 1.-3. day: 100-200 mg 3x per day
- 4.-14. day: 400-500 mg 3x per day

Mebendazolum

Albendazolum

- muscle phase
- 400 mg per day in 2 doses for 3 weeks

Cortikosteroids

Symptomatic treatment

In new infection it is possible to use albendazolum and anticonstipacy treatment every other day for 10 days



Taenia saginata

Cestoda

Distribution: geopolit,
typical **food habits**

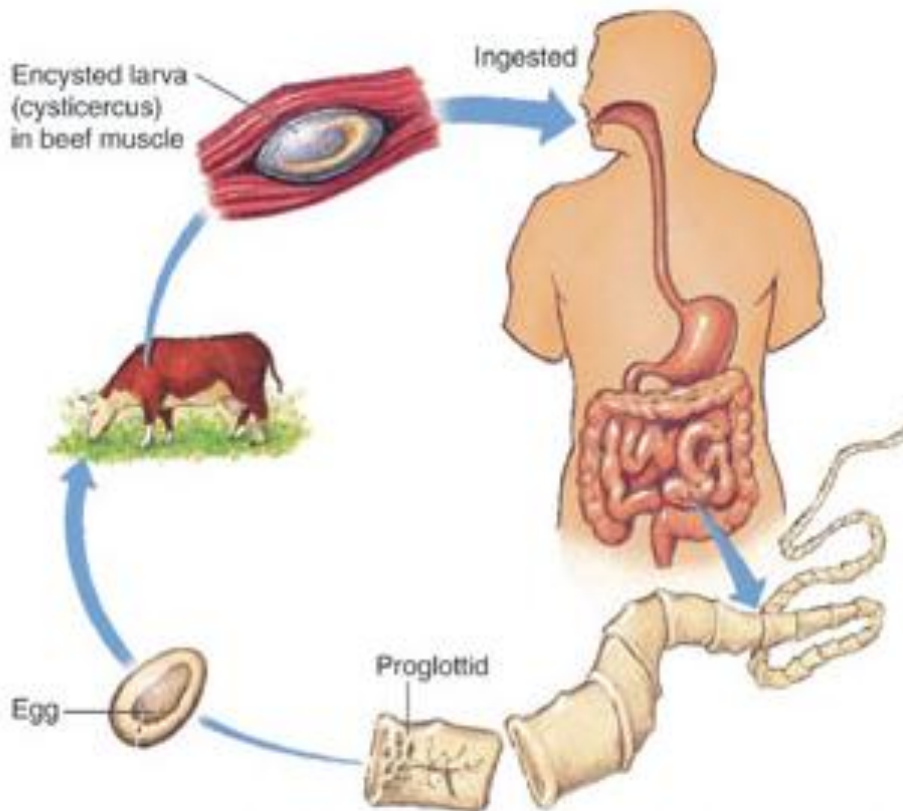
Source of the infection:
raw or uncooked **beef**

Final host: **human**

Intermediate host: **cattle**



Life cycle



In the muscles of cattle:

cysticercus bovis

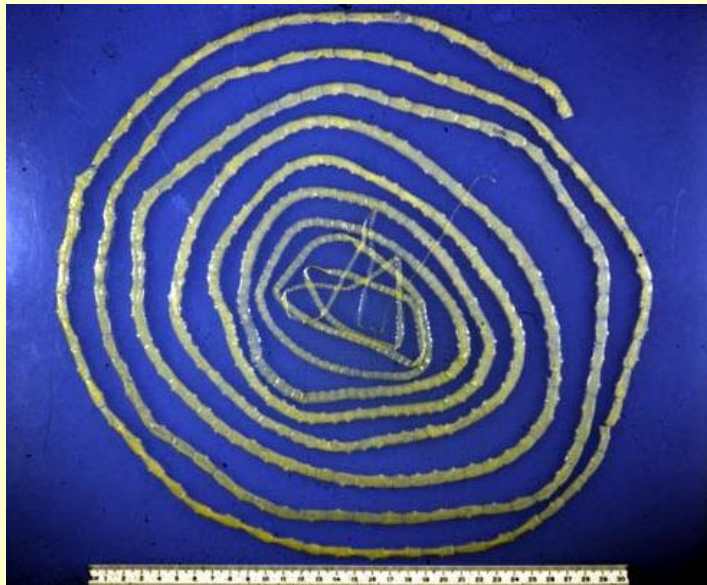
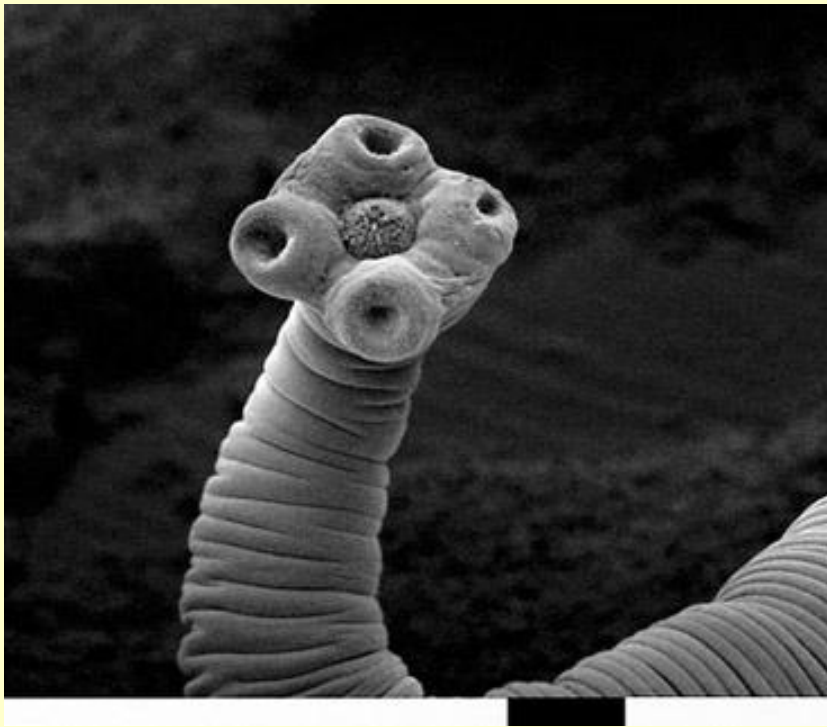
(5-10 mm)

Final host is

discharging

proglottids

containing **eggs**



Adult measures 3-10 m

Prepatent period:

6-12 months

Female releases: 1000-2000
proglottids (80-100 th. Eggs per
day)

Life expectancy: 20 years

Symptomatology

Usually **asymptomatic**
Malnutrition in heavy infection

Atypic migration of proglottids - apendicitis

Cysticercus bovis **only in ruminants**: muscles, myocardium, diaphragm, oesophagus



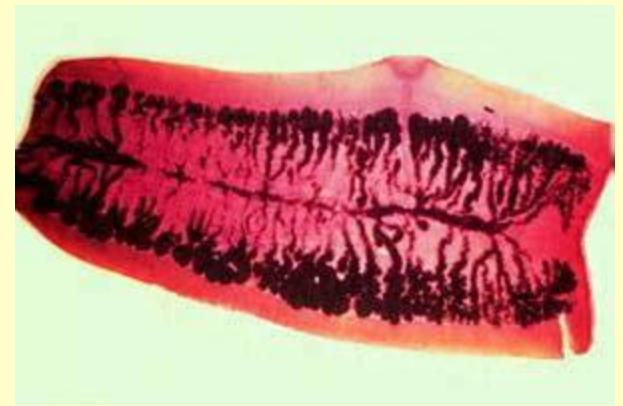
Diagnostics

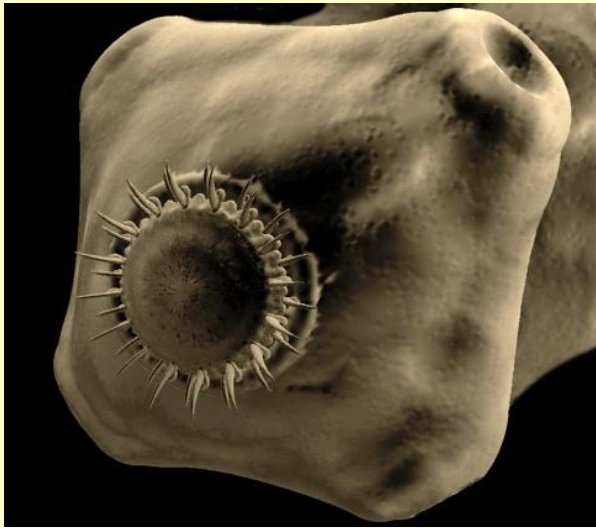


Proglottids in stool

(also discharged with no relation to defecation)

Eggs in anal swabs





Taenia solium

Cestoda

Distribution: **cosmopolite**

Transmission - **alimentary**:

undercooked pork meat –

human as **final host**

food contaminated by eggs –

human as **intermediate host**



Epidemiology

Cysticercosis – 60% CNS; 3% eye
Common asymptomatic infection

Prevalence of neurocysticercosis

Mexico City 1.4 - 3.6% (autopsy)

Bolivia 22% (seropositivity)

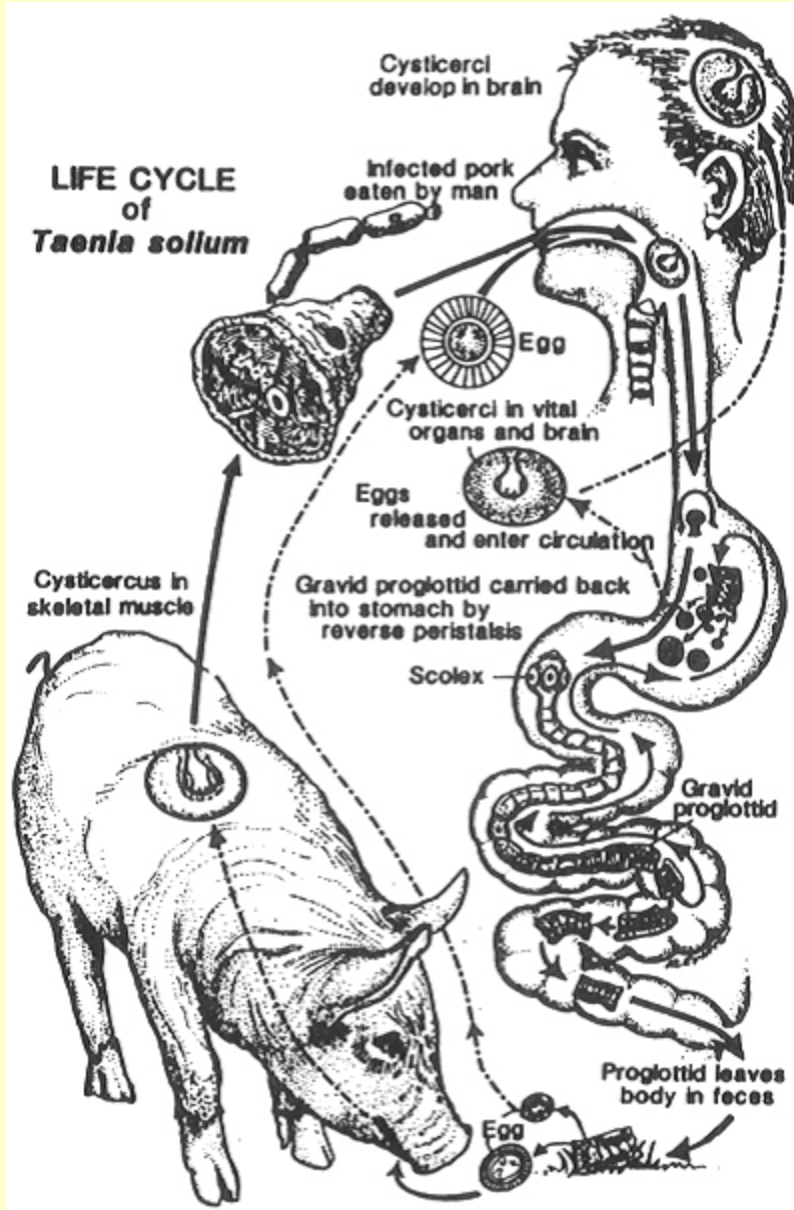
Peru 8%

Rwanda 21%

Bombay 47%

(Seropositivity in orthodox Jews in USA?)

Life cycle



Final host: human
(proglottids)

Intermediate host:
swine (human) –
cysticercus
cellulosae

Inhabitates **the intestine**

Adults measure **2-3 m**

Scolex with suckers and hooks

Prepatent period: 11-12 months



Human as final host: symptomatology

Usually asymptomatic

Irritating movement of parasite, toxins –
unspecific GIT problems

Cysticercosis



© U.L.B. - Louis De Vos



Cysticercus cellulosae:

swine, human

Localisation in host:

**muscles, brain,
subcutaneous infection**

Symptoms are dependent on:

lasting of infection

number of cysticerci

their localisation

immune response of host

Neurocysticercosis (active disease)

Arachnoiditis

Meningeal localisation:

Obstructive hydrocephalus
intrakranial hypertension

Parenchymatic localisation:

asymptomatic;
Brain oedema, seizures, focal neurologic deficiency,
intracranial hypertension

Neurocysticercosis – inactive disease

Most common form of disease

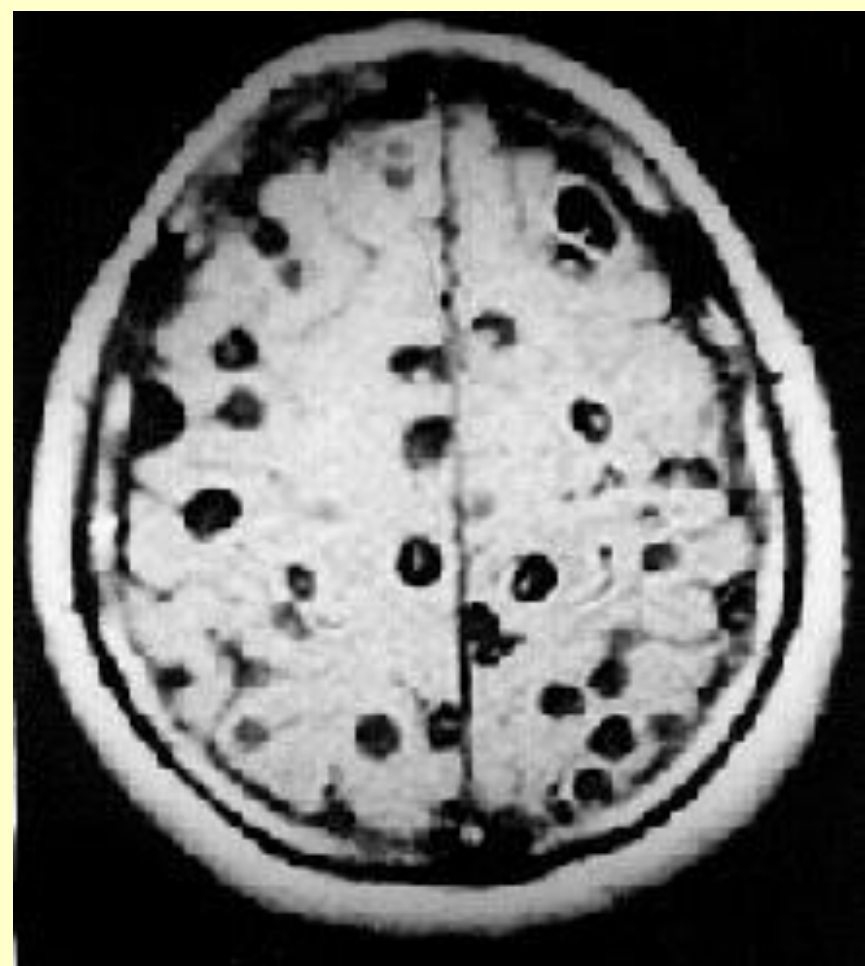
60% of cases parenchymatic localisation

Seizures
Headache
Vomiting

Changes of intellect, ataxy....



Neurocysticercosis



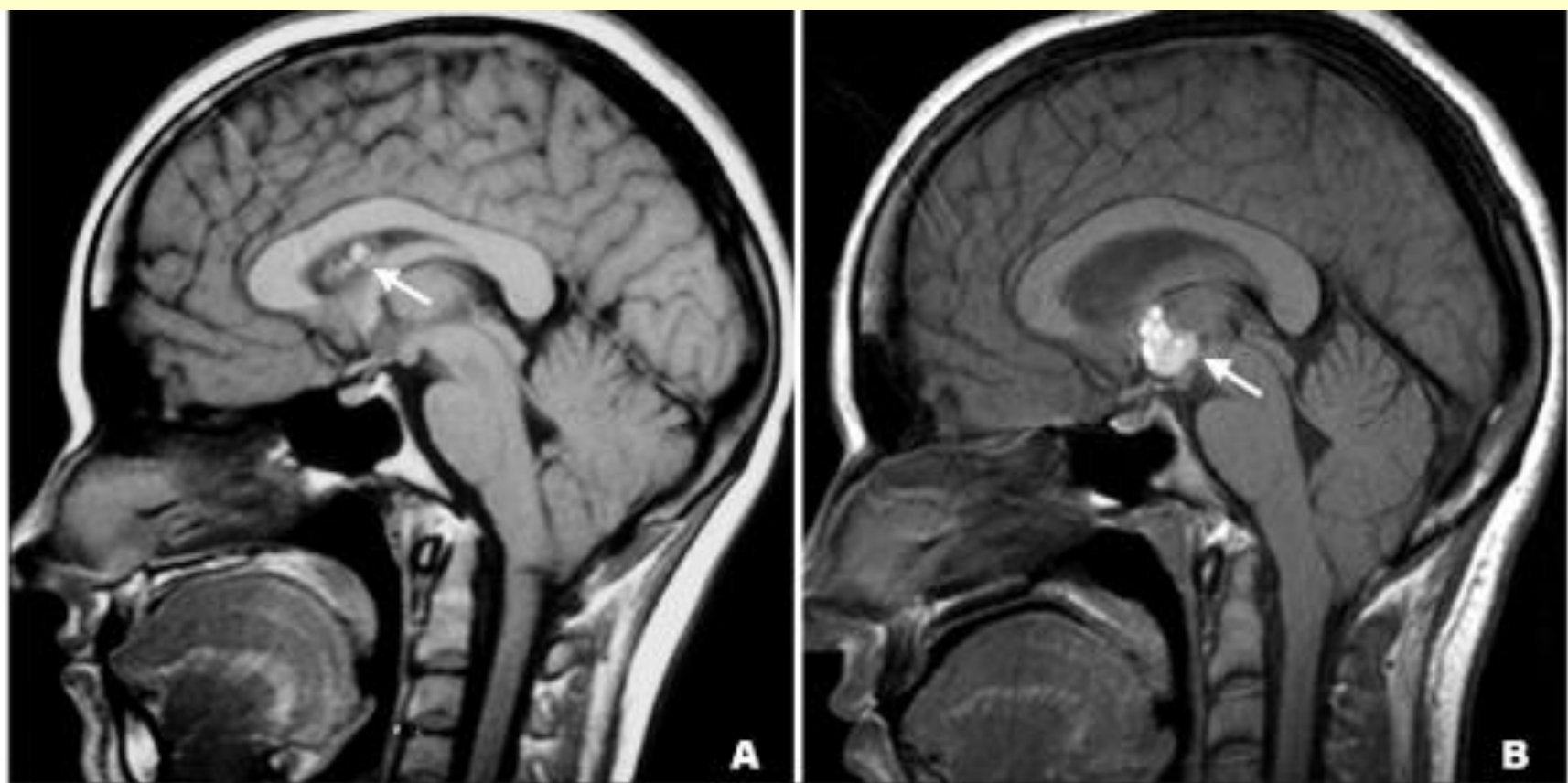


Fig 3. Sagittal T1-weighted MR images demonstrate a multiloculated lesion in the lateral ventricle (arrow) in the first exam (A), and the lesion migrating to the third ventricle (arrow) on the follow-up (B).

Eye – (ophthalmocysticercosis)

- frontal chamber, vitreous humour, below retina:
inflammatory changes, atrophy of retina, chorioretinitis,
iridocyklitis, catarakta

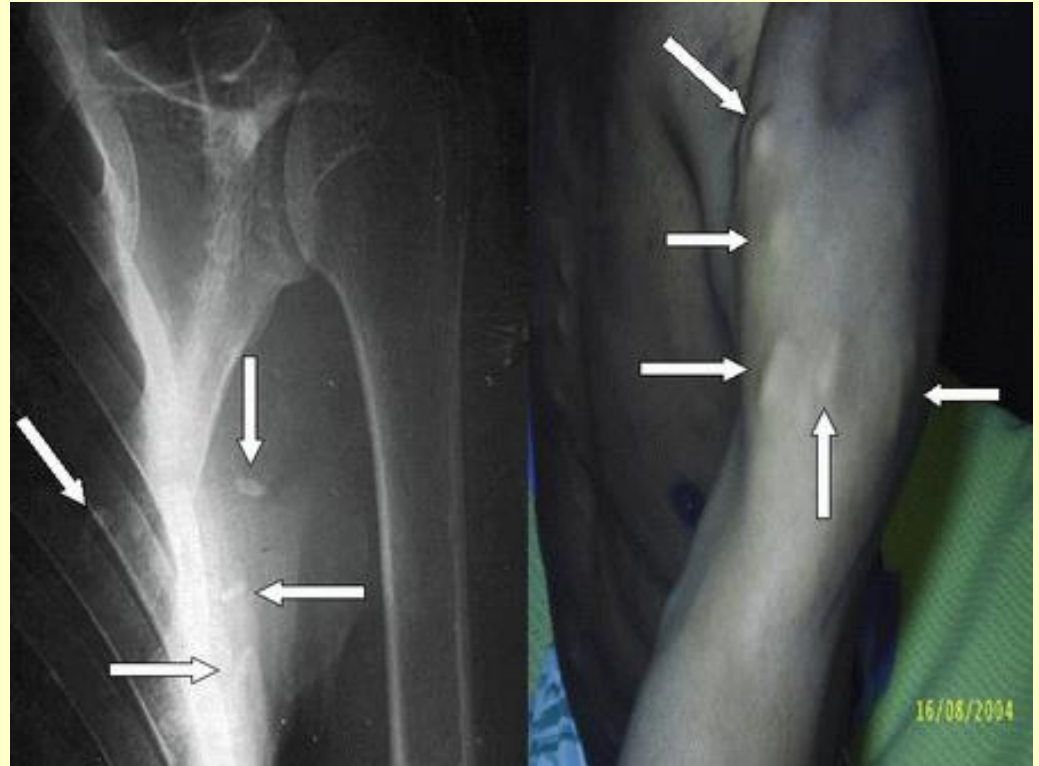
Subcutaneous –

Solitary or multiple; resembling neurofibromatosis

Subcutaneous cysticercus



Cysticercosis of muscles





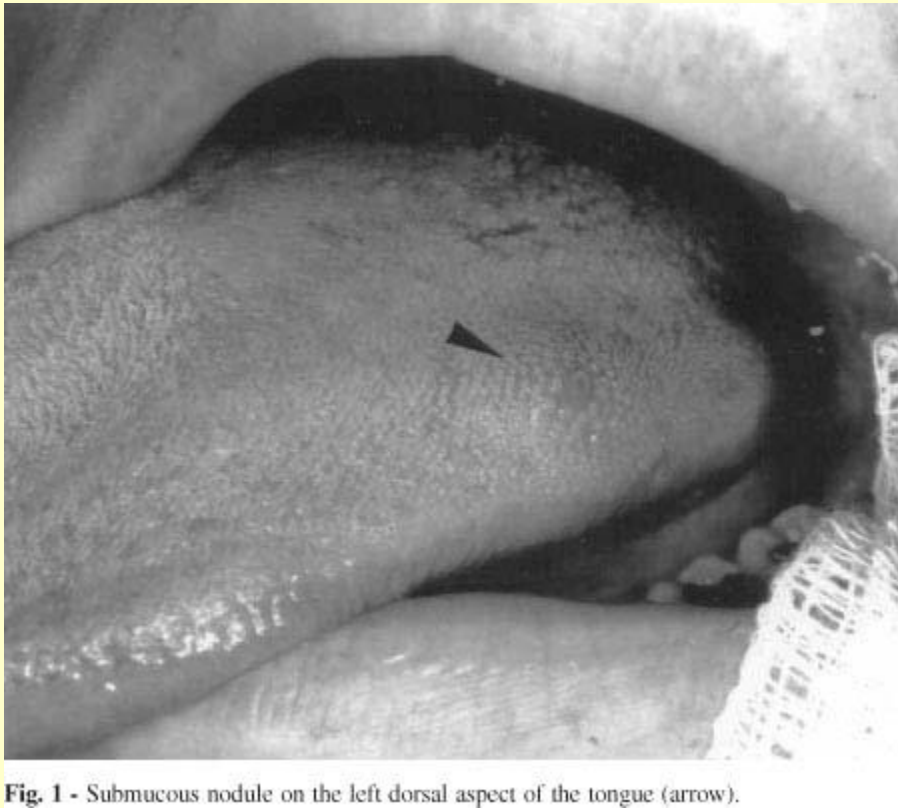


Fig. 1 - Submucous nodule on the left dorsal aspect of the tongue (arrow).

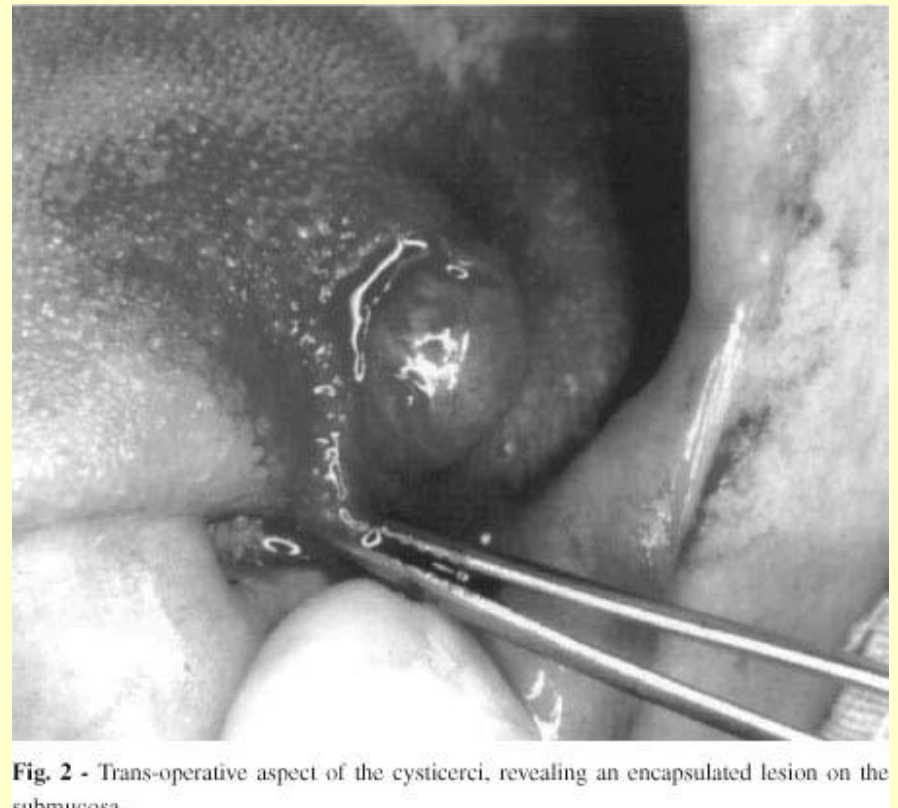


Fig. 2 - Trans-operative aspect of the cysticerci, revealing an encapsulated lesion on the submucosa.

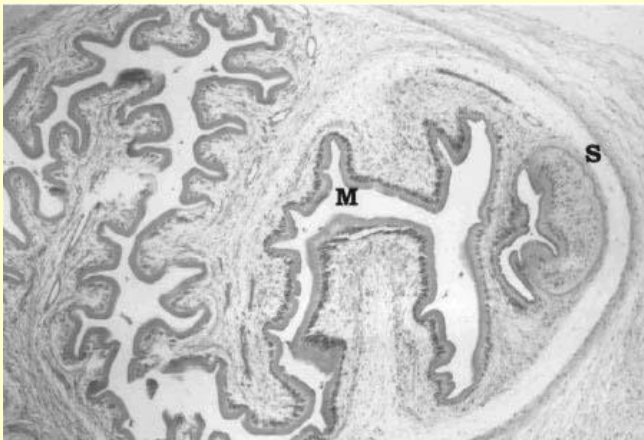
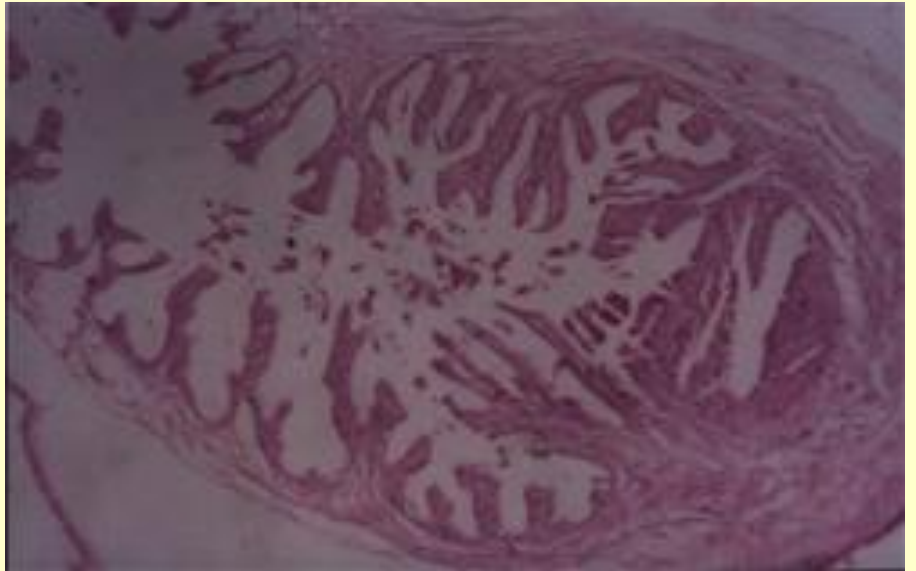
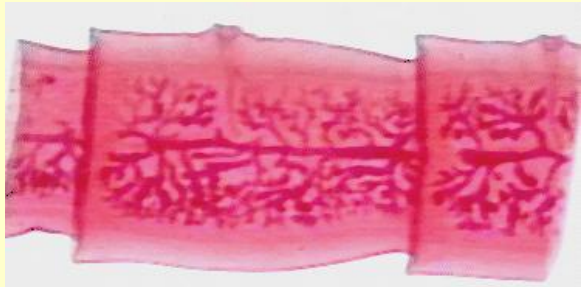


Fig. 4 - The duct-like invagination, which composes the caudal end of the larva, lined by the homogeneous membrane (M). Scolex (S) at the cephalic end. Haematoxylin and eosin x 12.5.





Diagnostics

Taenia
egg



Final host

Proglottids in stool

Eggs in perianal swabs

Intermediate host

Imaging techniques

(US, CT, NMR),
calcificated, hypodense leasions

Serology

ELISA (3 months pi)



Final host therapy

Niklosamid (YOMESAN- Merck, tbl. 500 mg):

Side effects: mild; headache, abdominal pain, fever

Doses: 2 g p.o. in a single dose, after fasting

children: < 11 kg = 0,5 g
11 – 34 kg = 1,0 g
> 34 kg = 1,5 g

Praziquantel (CESOL 150 mg; BILTRICIDE 600 mg)

doses:

– 5-10 mg/kg in a single dose, after meal

Intermediate host therapy:

- **CHEMOTHERAPEUTICS:**
 - Praziquantel 10 – 25 mg/kg 3x per day, 2-3 weeks
 - Albendazol um 7,5 - 15 mg/kg/day (max. 800 mg) in 2 doses., 2-4 weeks
 - Cortikosteroid therapy for supression of oedema and intracranial hypertension
- Chemotherapy is **not indicated** in severe active **neurocysticercosis**, (could lead to life threatening inflammatory reaction), symptomatic therapy
- **Solitary cyst** with symptoms of epilepsy – anticonvulsive therapy
- **Surgery** in **subarachnoideal** and **intraventricular cysts**, causig compression or hydrocephalus
- **Ocular cysts** are treated **surgically** with no chemotherapy

Prevention

- Sufficiently **cooked pork** meat

Freezer:

- 5 C 4 days
- 15 C 3 days
- 24 C 1 day

Toxocara canis/cati

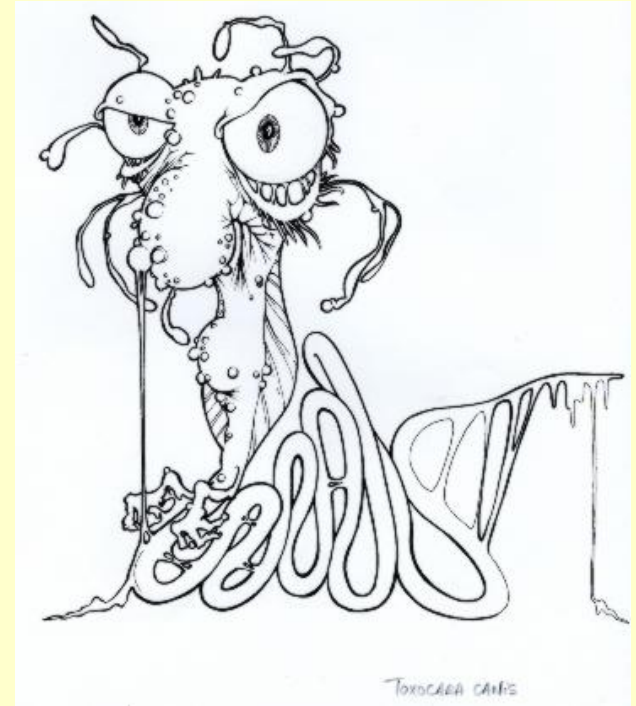
Nematoda

Cosmopolite distribution

Seroprevalence USA: 2-10%

Transmission: — eggs in soil, sand,
larvae in paratenic host

Most common: children



Epidemiology

2-5% positive in cities of developed world

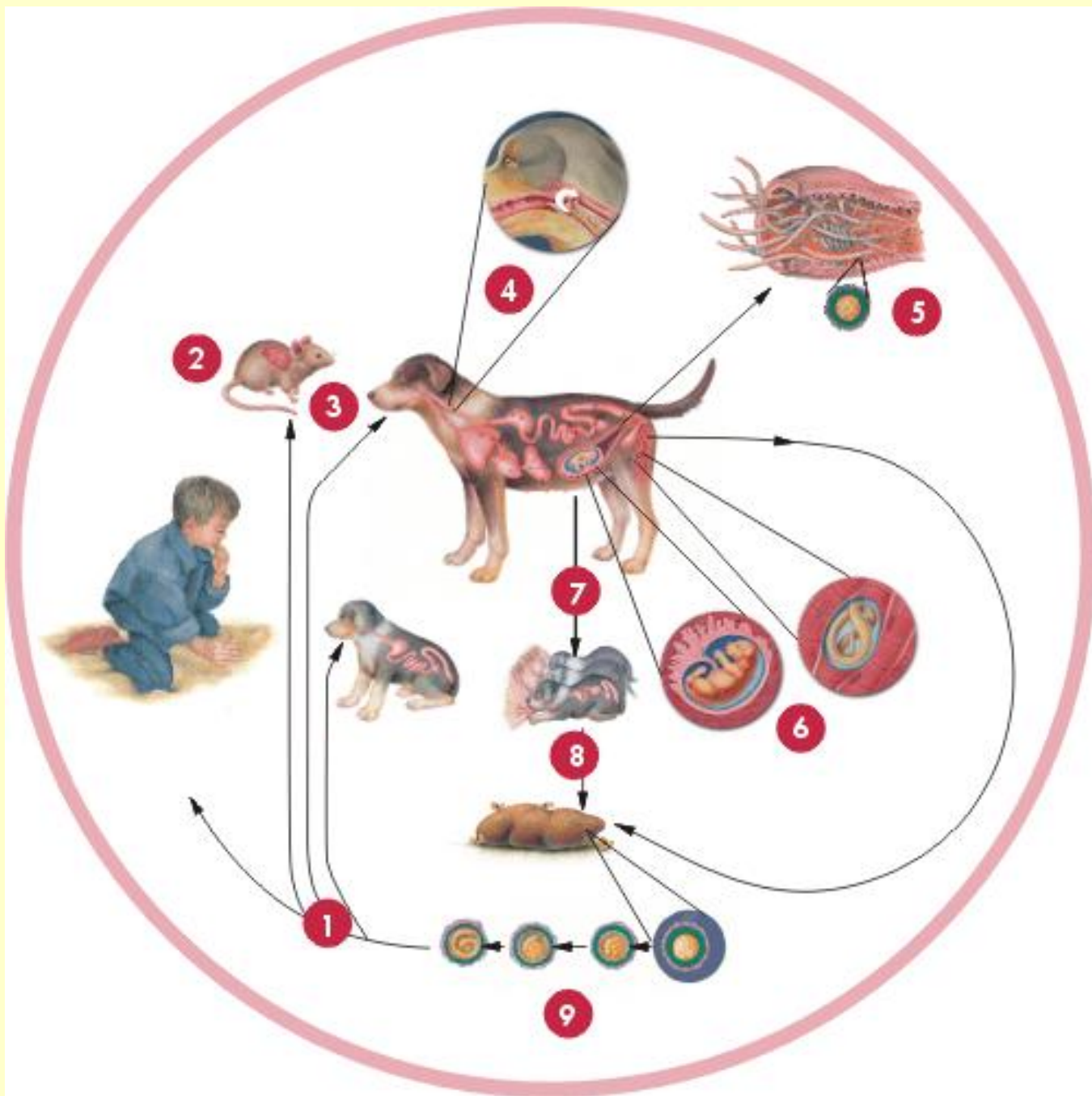
14.2-37% positive in villages in developed world

Tropical countries:

63.2% Bali,

86% Santa Lucia (West Indies)

92.8% La Reunion (French Overseas Territories, Indian Ocean)



Symptomatology

Asymptomatic seroconversion

Larva migrans visceralis

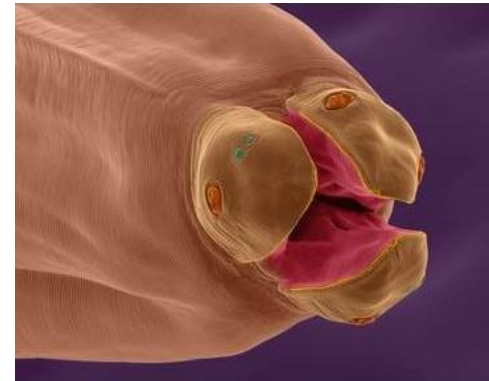
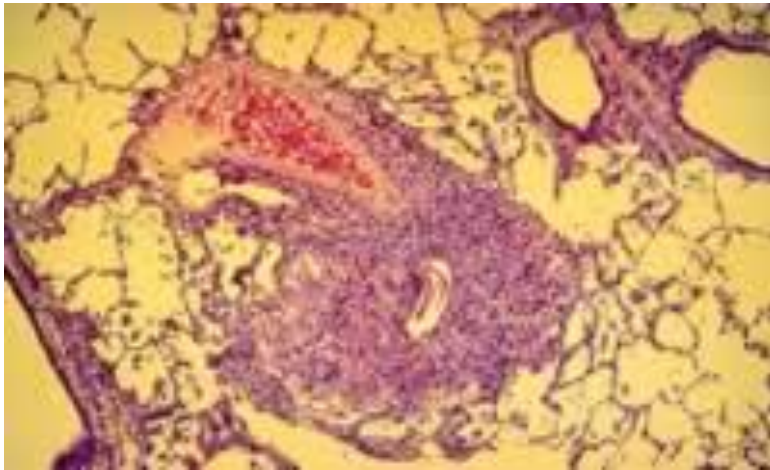
Larva migrans ocularis

Symptomatology is dependent on

Number of larvae in host and immune response

Allergic reaction!!!

Granuloma formation



Larva migrans visceralis – symptomatology
due to the migration of larvae in host

Abdominal pain, nausea, vomiting

Exanthema, pruritus

Hepatomegaly

Pneumonia (cough, fever)

Letargy, difficulties in sleeping

Headache

Myositis

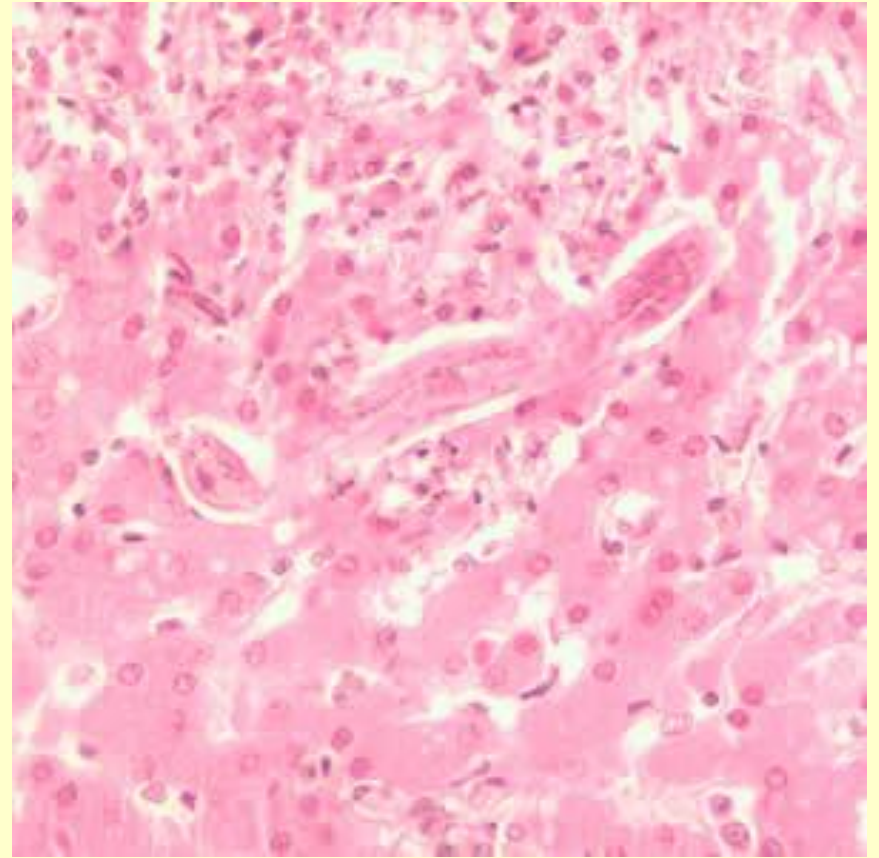
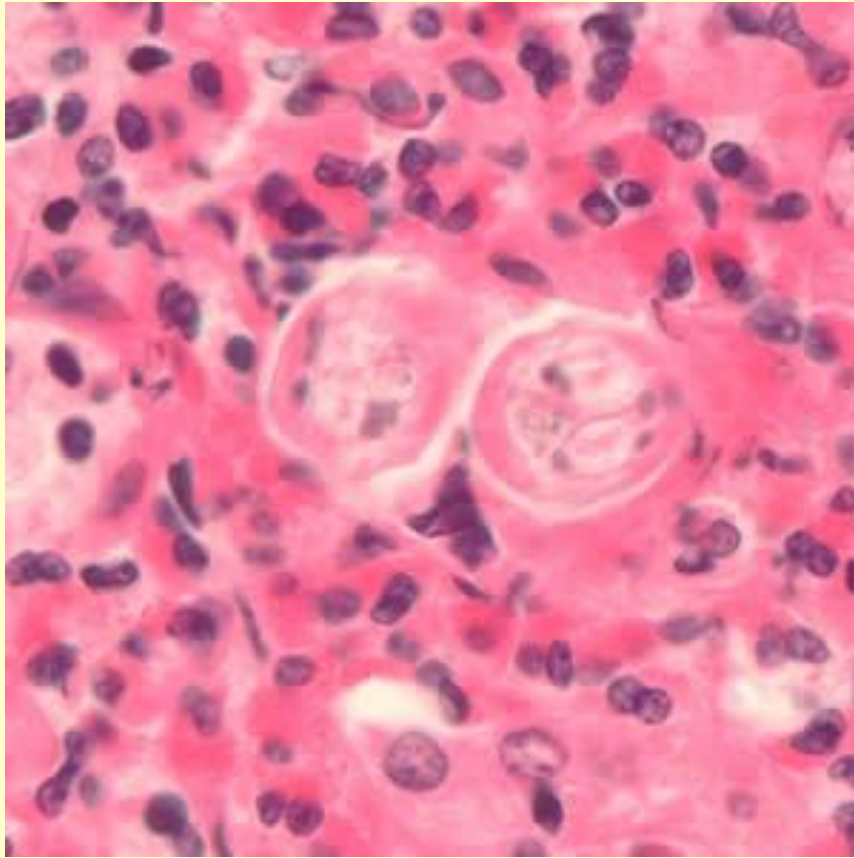
Rarely: seizures, myocarditis

EOSINOPHILIA!!!

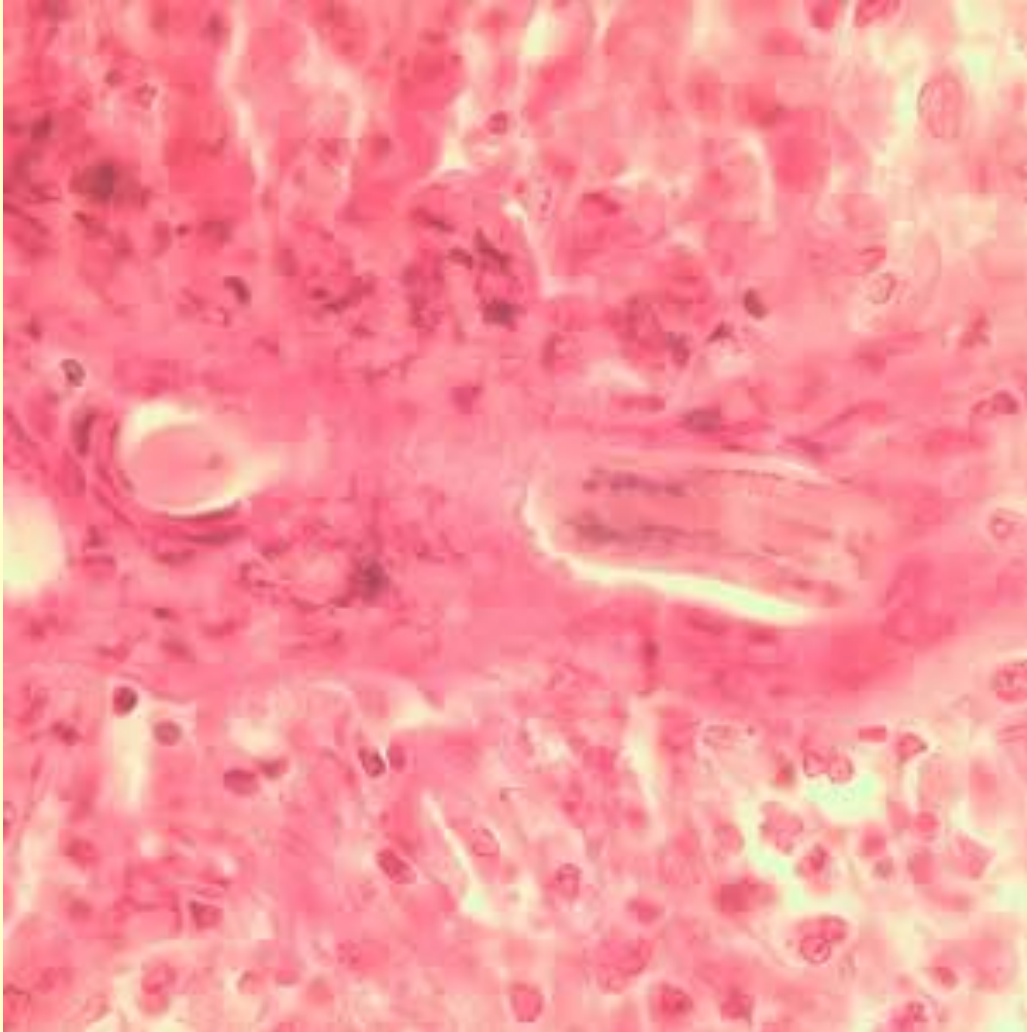
Exanthema; larva migrans visceralis



Larva migrans visceralis - hepar



Larva migrans visceralis - lungs

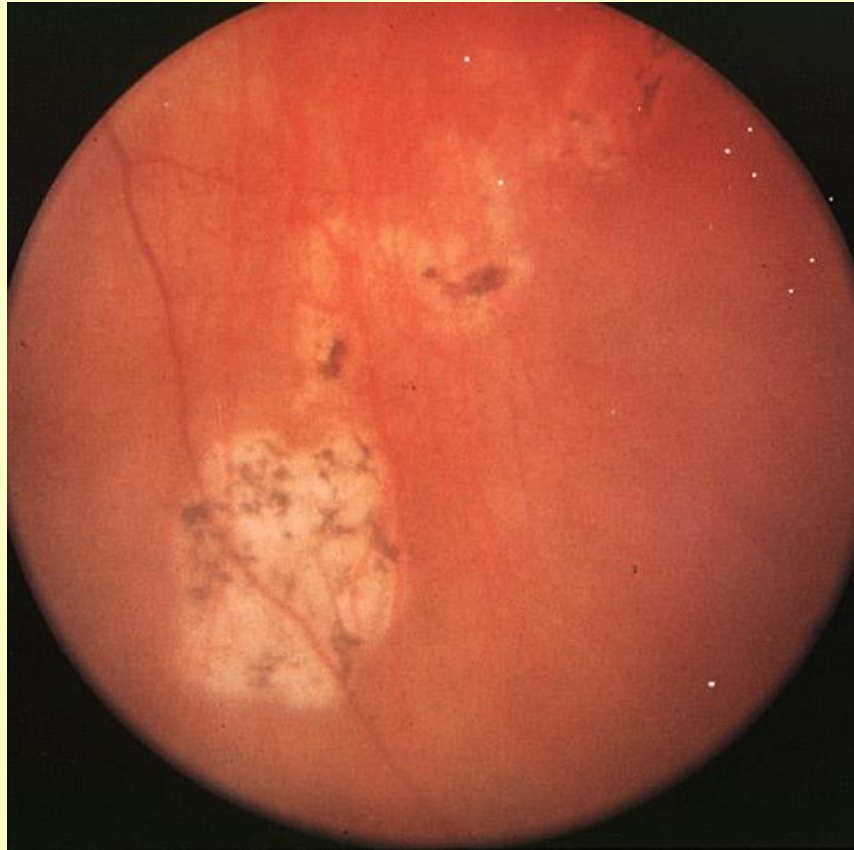


Larva migrans ocularis

Visual disturbances usually unilateral

Strabismus

Leucokoria



Disease in dogs

5-50% seropositive in Europe

Tissue and GIT phase

Sleeping larvae

(transplacental, transmammary infection)

Eggs shed into the environment not mature

Diagnostics

Leukocytosis

Eosinophilia

+ symptomatology

Definitive diagnostics:

Positive serology, **biopsy**

Therapy

Positive serology with no symptomatology dont treat!!!!!!

Larva migrans visceralis – corticoids (Prednison)

Ocular form:

Albendazole (Albenza) - 10 mg/kg/d PO single dose for 4 weeks

Mebendazole (Vermox) - 25 mg/kg/d PO single dose for 4 weeks

Echinococcosis

Trematoda

Echinococcus granulosus, multilocularis, vogeli

Distribution: cosmopolite (Australia, New Zealand)

Definitive host

dog, wolf, coyote (granulosus)

dog, wolf, fox, cat (multilocularis)

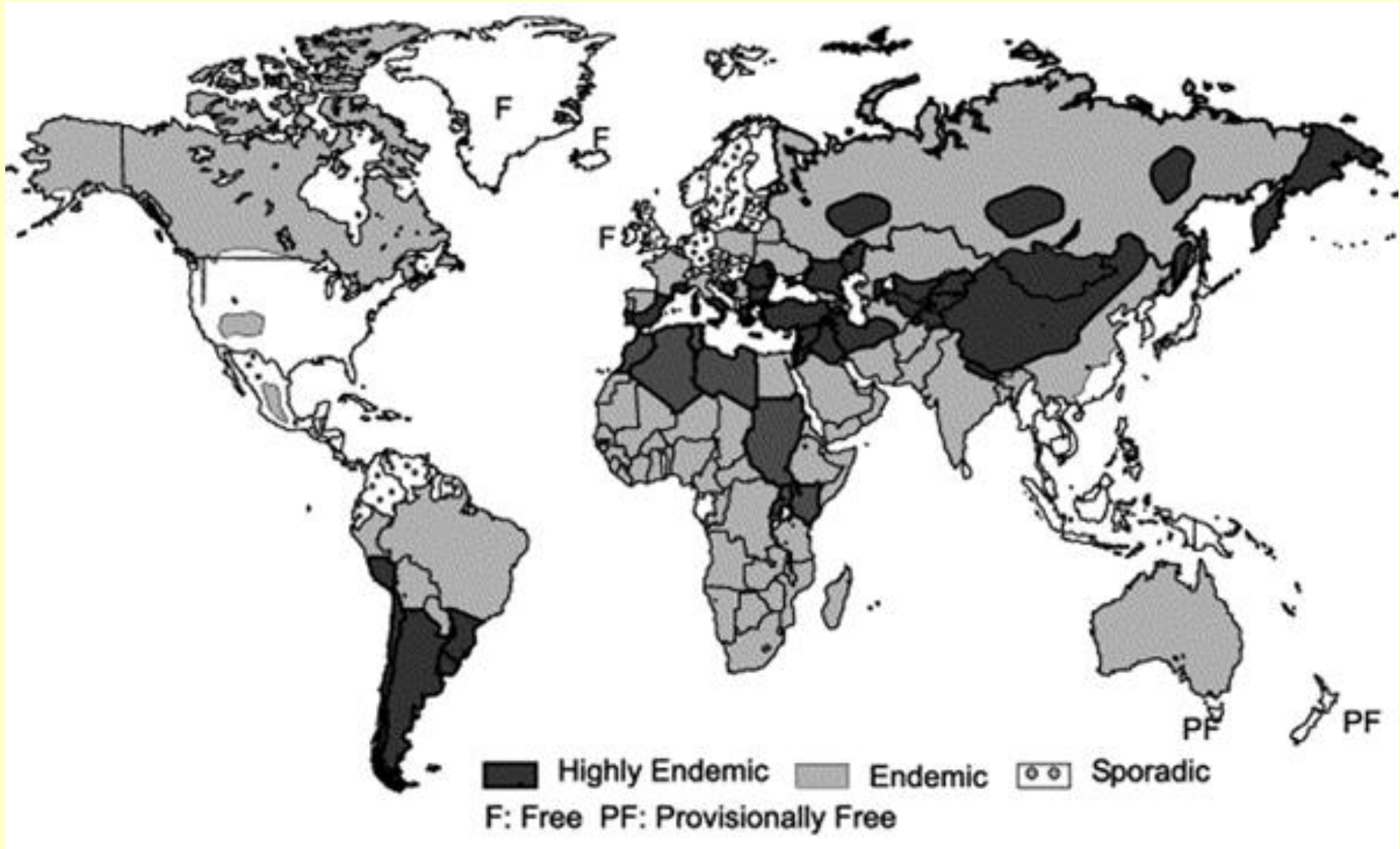
Intermediate host:

sheep, swine, deer (granulosus);

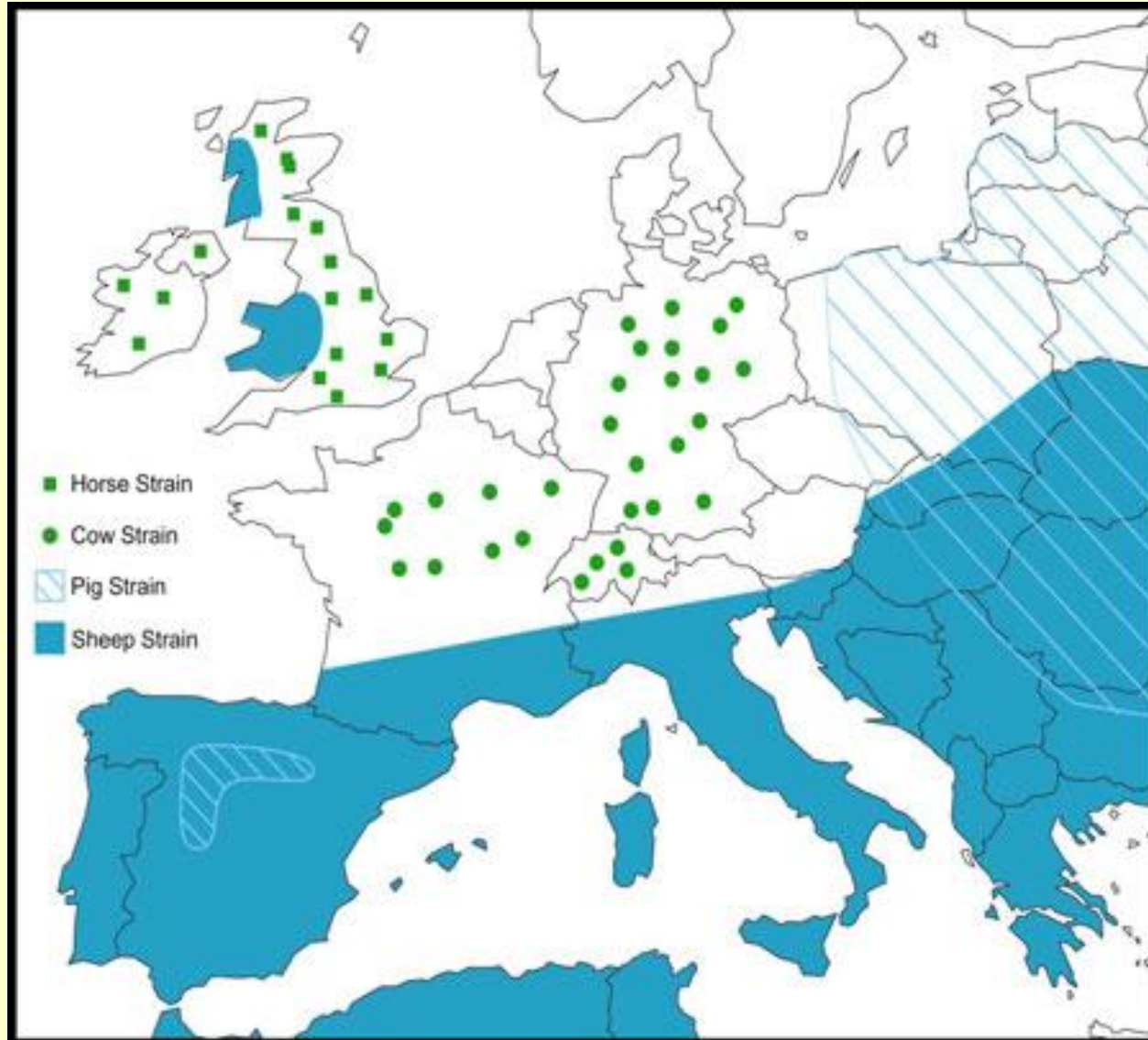
rodents (multilocularis)



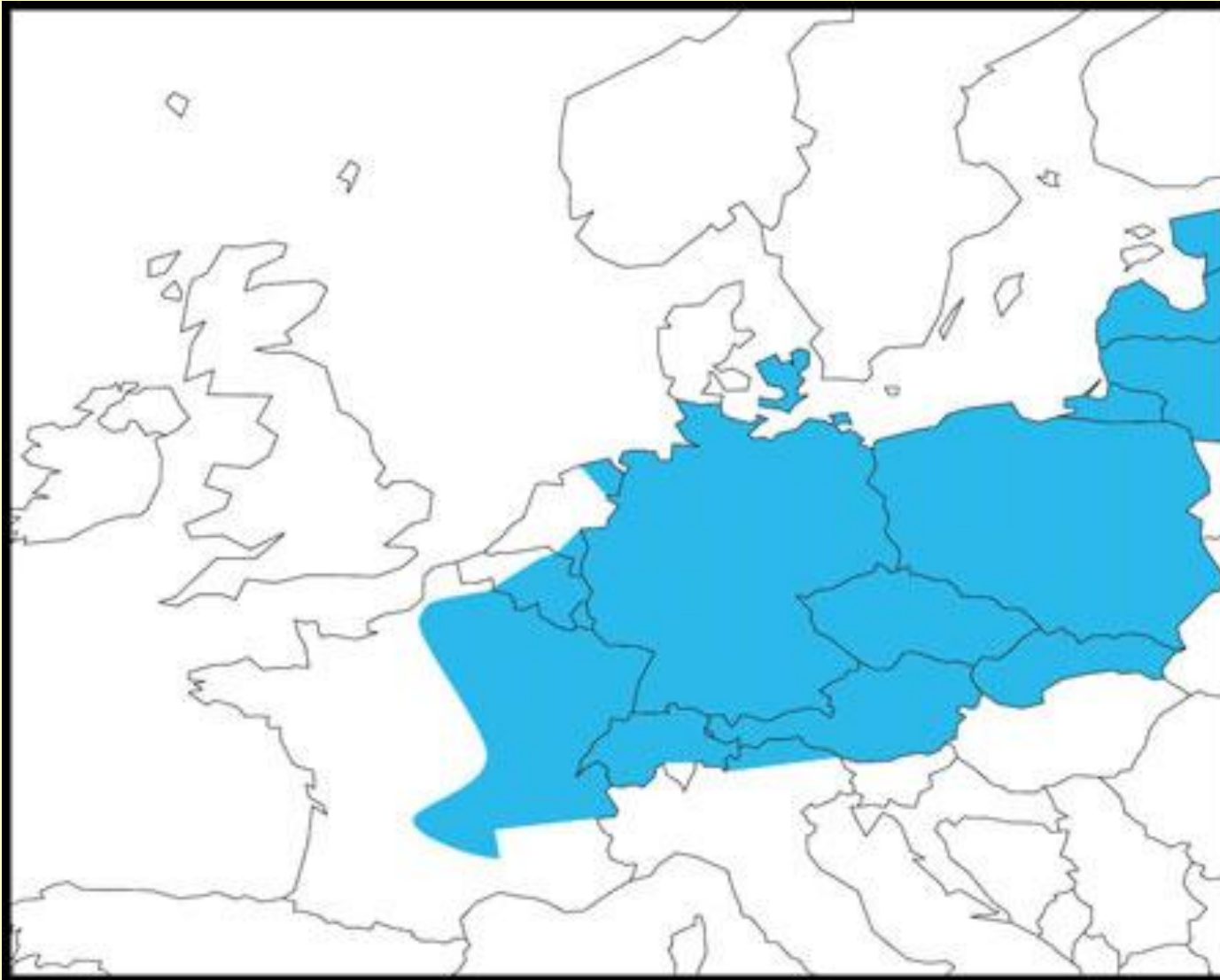
Distribution of *E. granulosus*; 2004, CDC

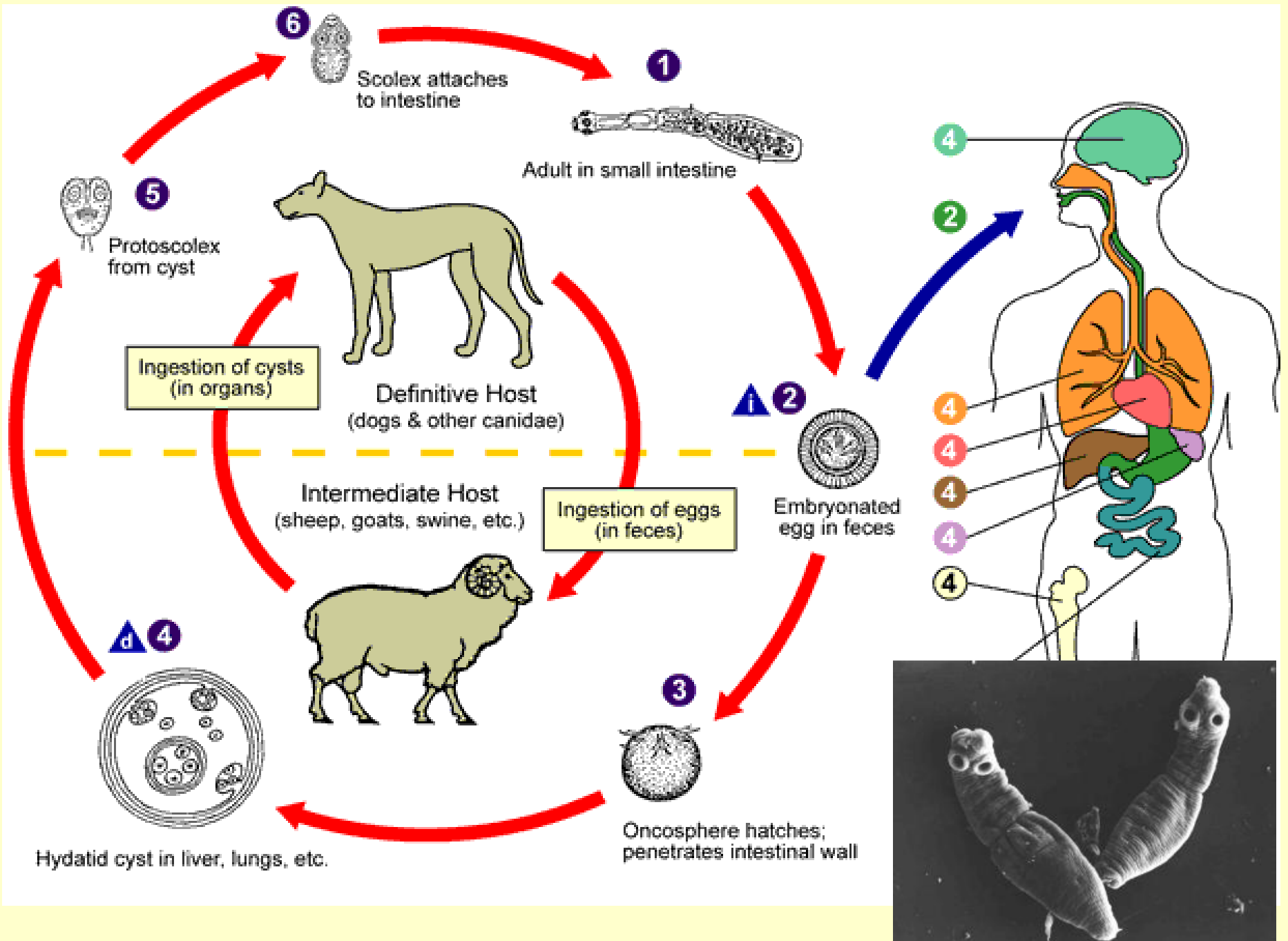


E. Granulosus in Europe

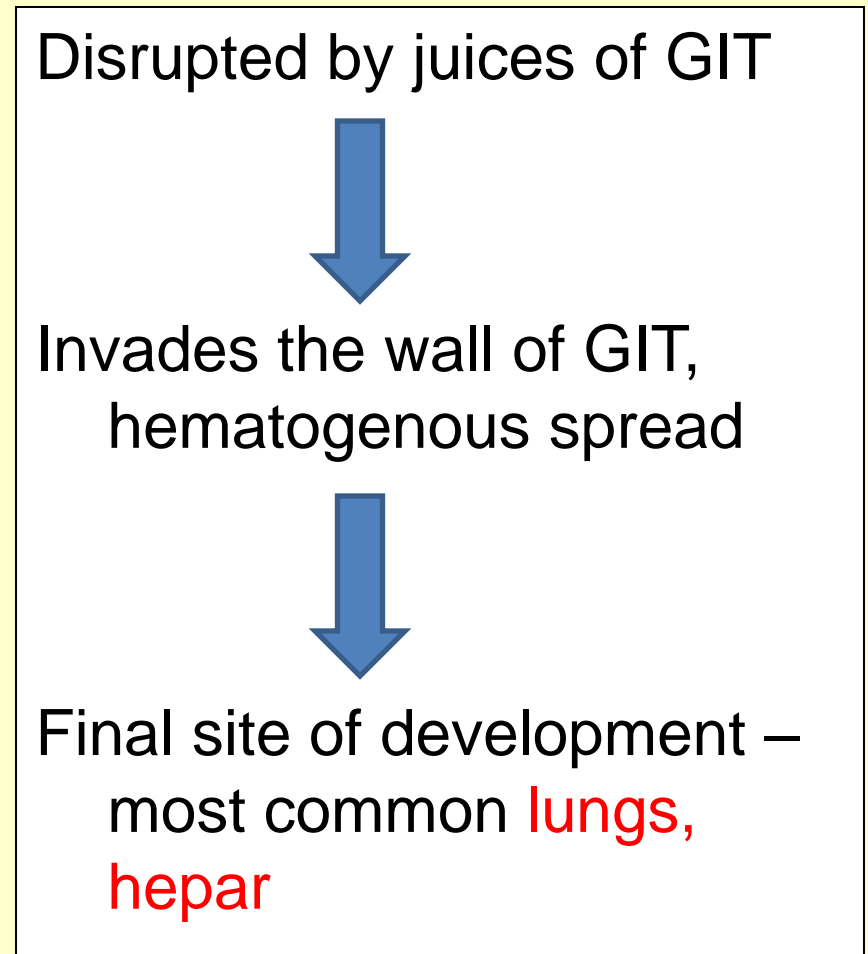
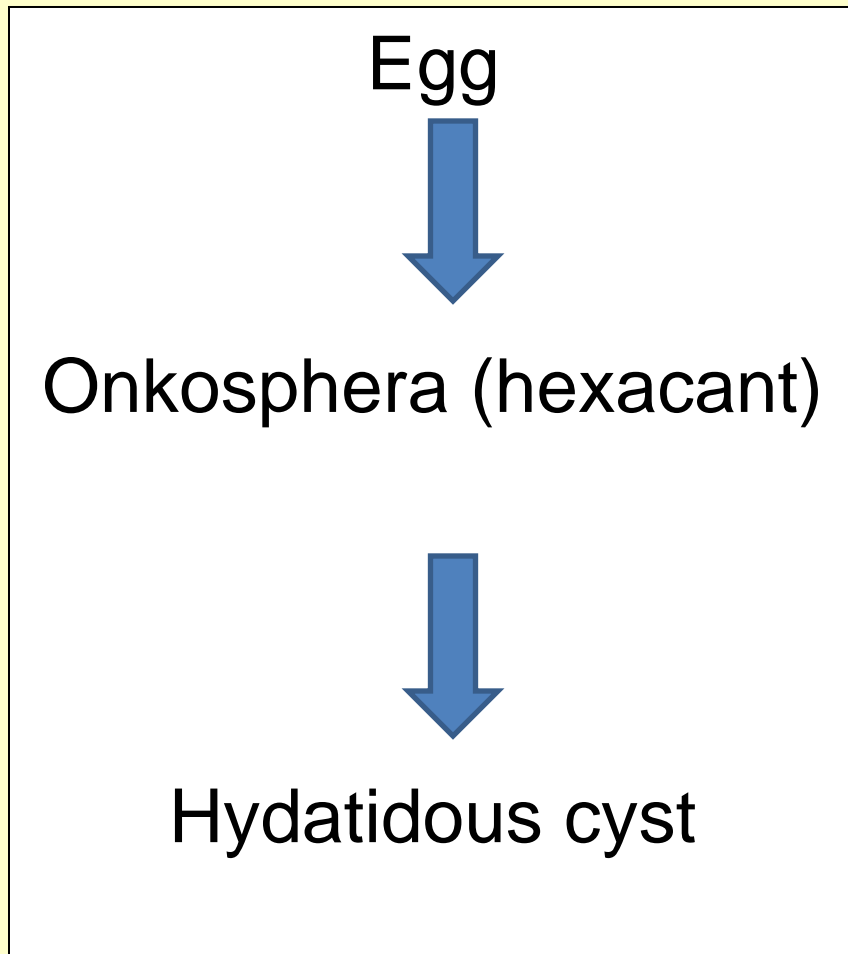


E. Multilocularis in Europe

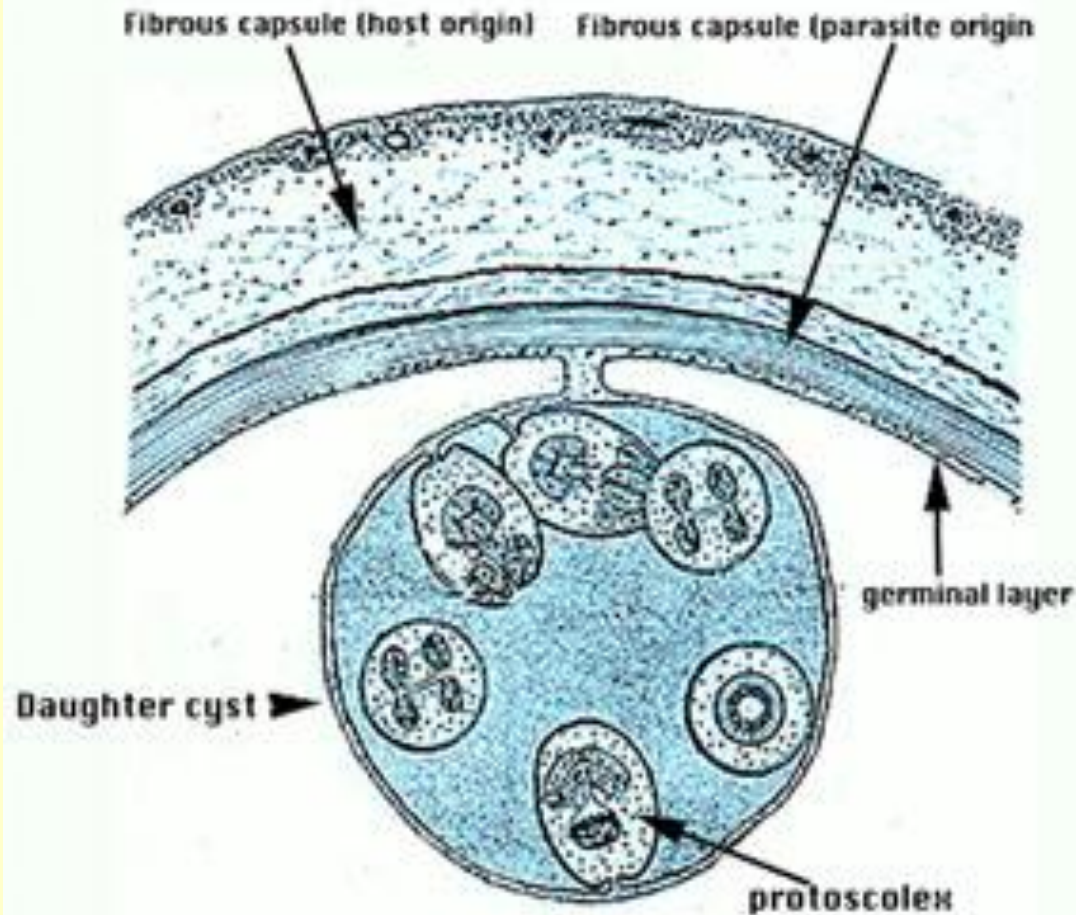




Human serves as accidental intermediate host



The cyst has **three walls**: germinal, fibrous, fibrous (host)



Hydatid cyst

Disease

Cystic echinococcosis (*E. granulosus*)

Alveolar echinococcosis (*E. multilocularis*)

Polycystic echinococcosis (*E. vogeli*)

Cystic echinococcosis

E. granulosus

Grows appr. **1-5 cm/year**
inside

Alveolar echinococcosis

E. multilocularis

grows **appr. mm/year**
outside

Protoscolexes grow from inner wall of the cyst,
the daughter cells and protoscolexes =
Hydatidous sand

Symptomatology

Localisation

Size of cyst

Relationship of expansive cyst to surrounding environment (bile duct, vascular system)

Complications due to the rupture of cyst

Secondary bacterial infection

Reaction of immune system

(asthma, anaphylactic shock, nephropathy)

Infection: primary and secondary

Hepatic involvement

- Usually **asymptomatic for long time**
- Accidental findings
- Abdominal discomfort, pain, decreased appetite
- **Hepatomegaly**
- **Icterus**
- Biliary colic

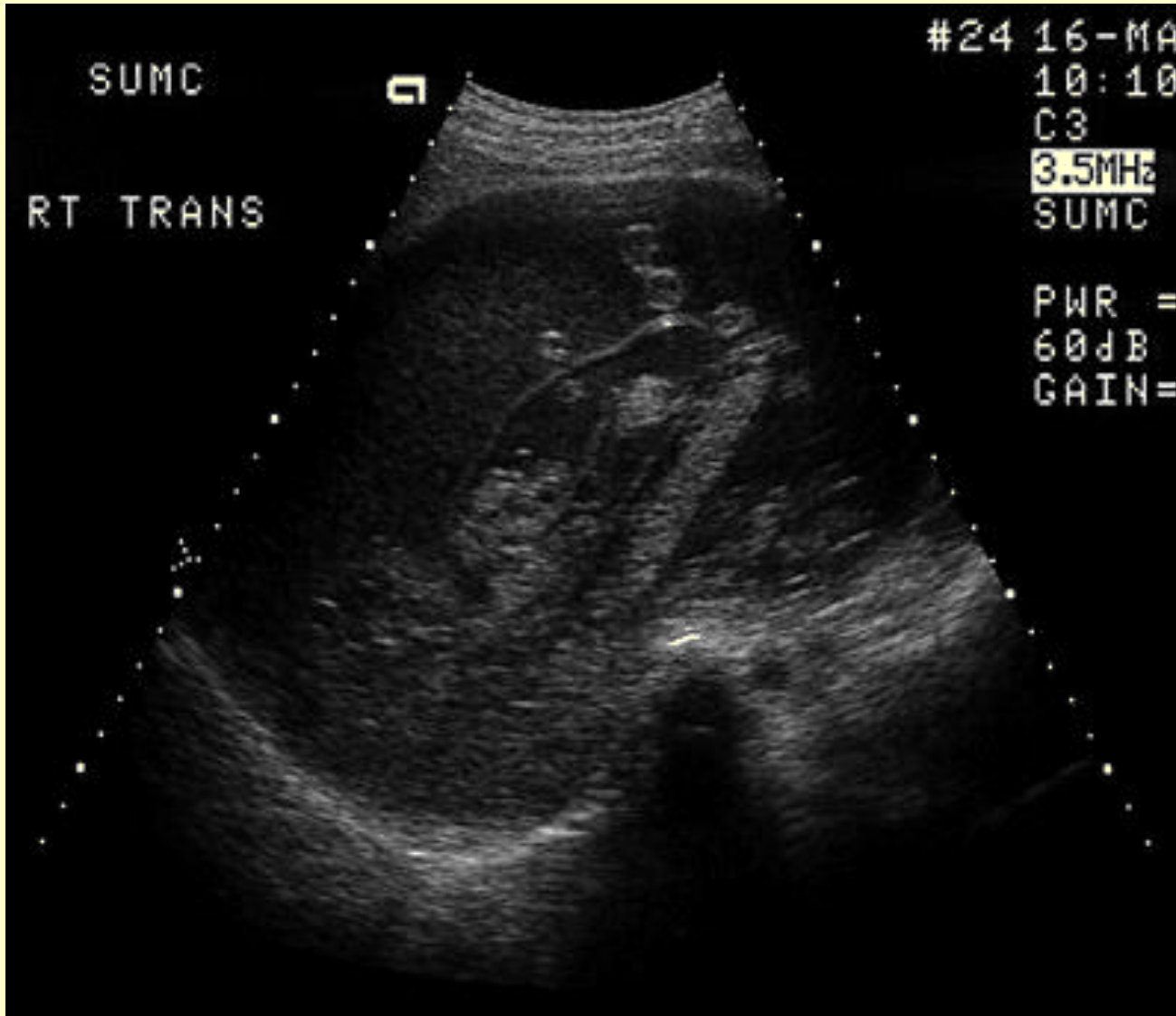
- **Biliary colic**
- Cholangiitis
- Pancreatitis
- **Abscessus**
- Portal hypertension
- Ascites
- Compression; thrombosis of *v. cava inferior*
- Budd-Chiari syndrome
- Rupture



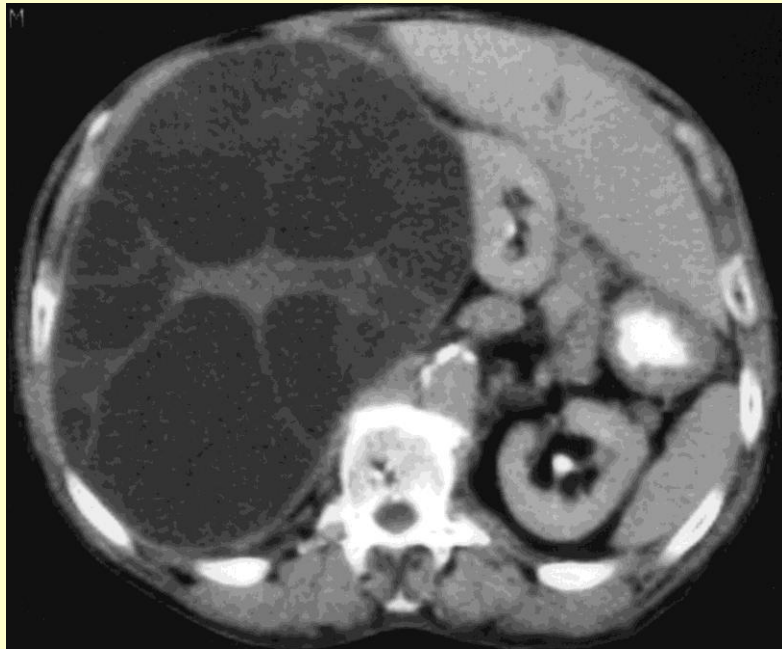




E. granulosus; hydatid sand



E. Multilocularis – cyst is divided by septae, it grows outside, inside necrotic tissue



Obr. 2. CT – stejný nález v sagitální rovině
Fig. 2. The same picture in sagittal plane



E. multilocularis, central necrosis

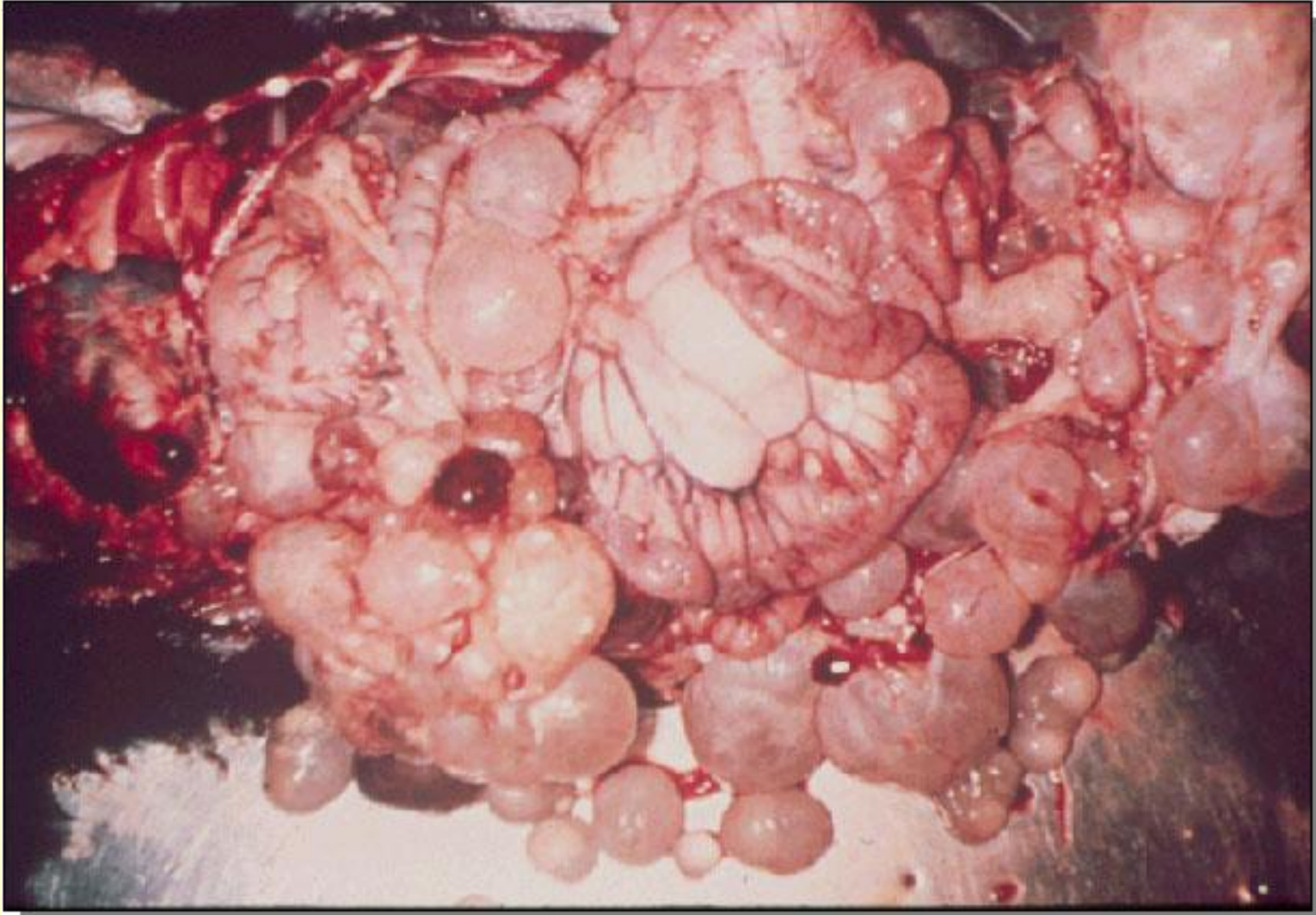


Obr. 4. Resekát jater s hydatidovou cystou larválního stadia *Echinococcus multilocularis*

Fig. 4. Resected liver specimen with hydatid cyst of larval stadium of *Echinococcus multilocularis*



Hydatid cyst



Involvement of lungs

„Tumour“ of lungs

Chest pain

Chronic cough, expectoration, dyspnoe

Pneumothorax

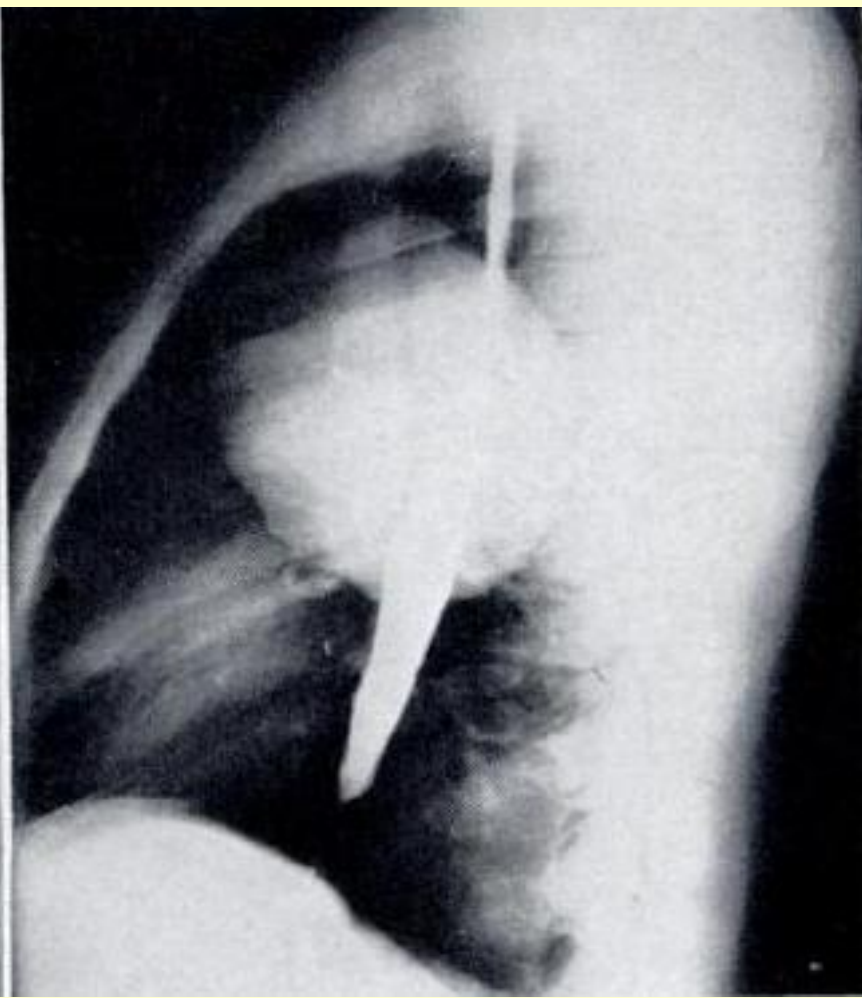
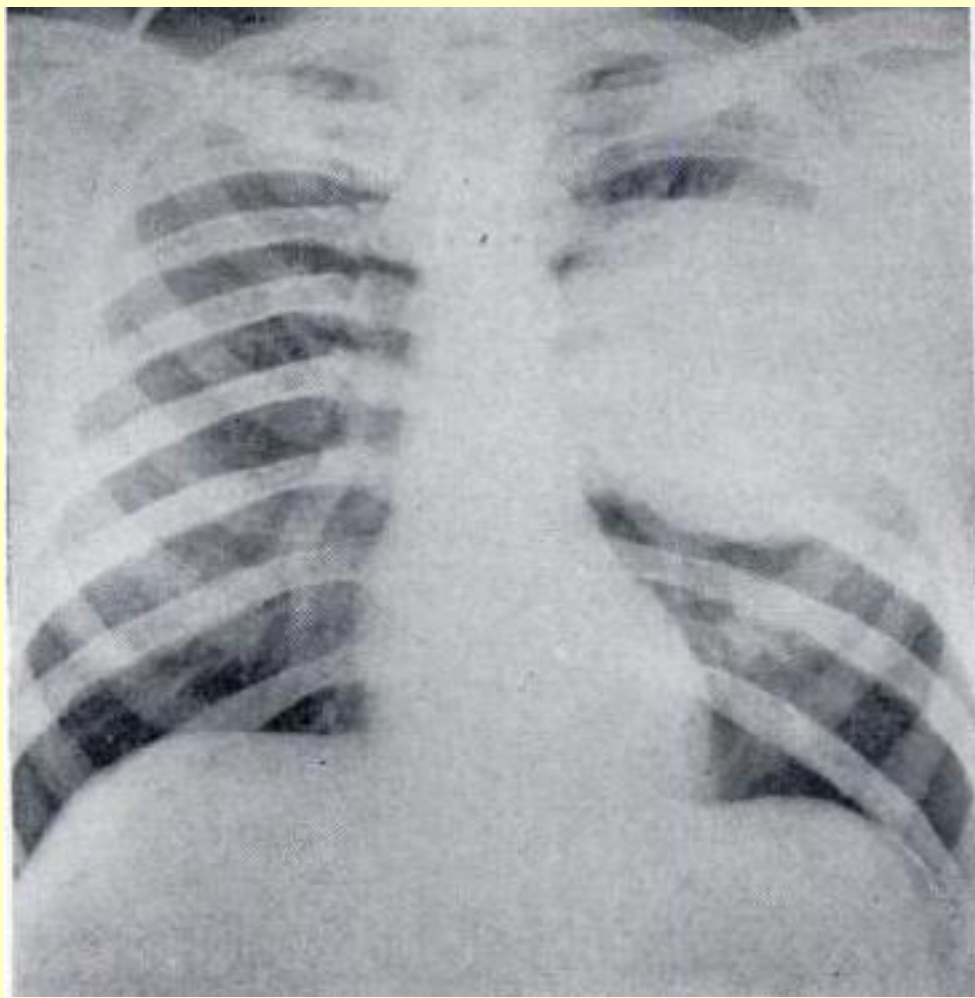
Eosinophilic pneumonitis

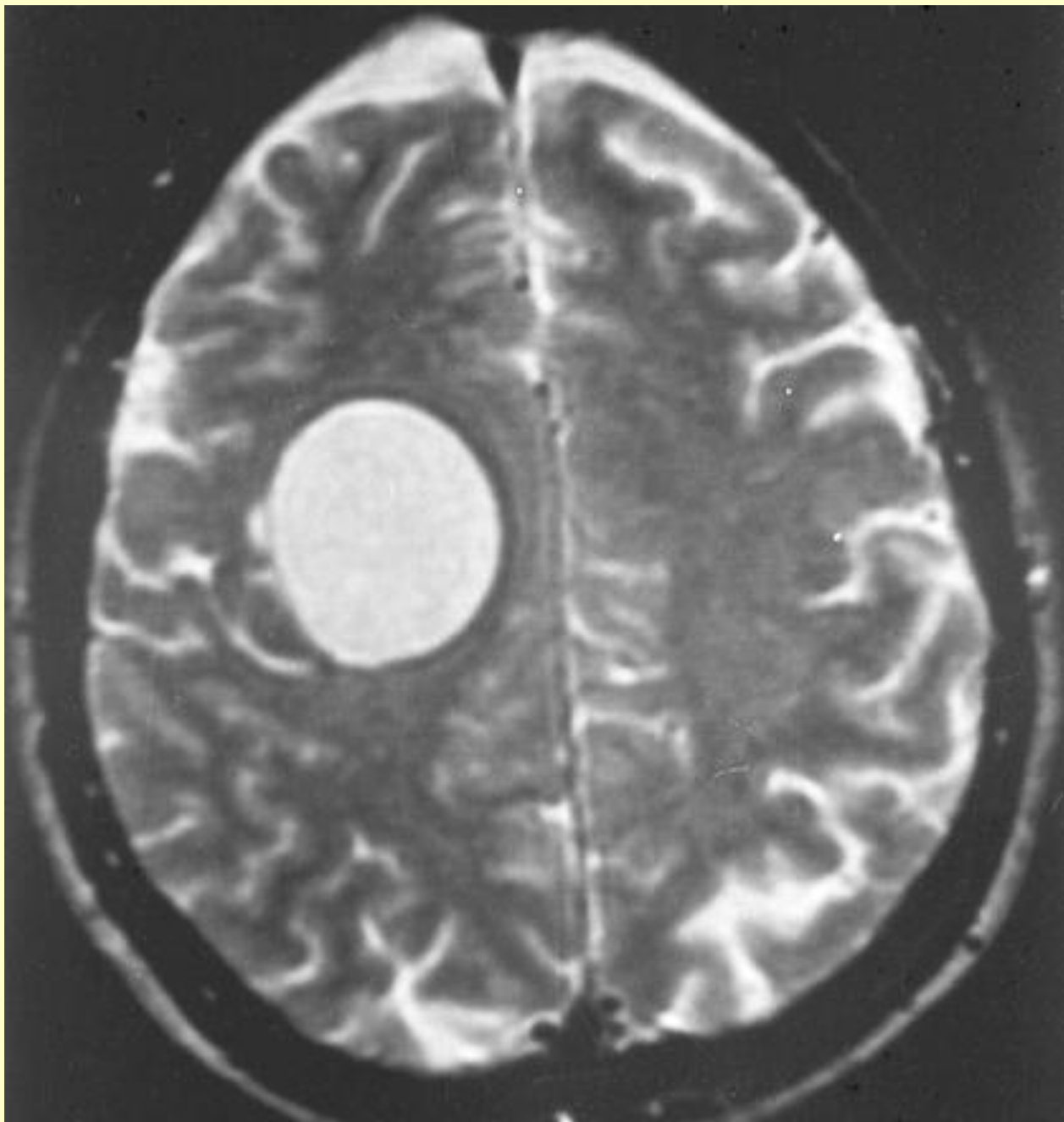
Pleural effusion

Parasitic pulmonary embolus

Hemoptysis

Biliptysis



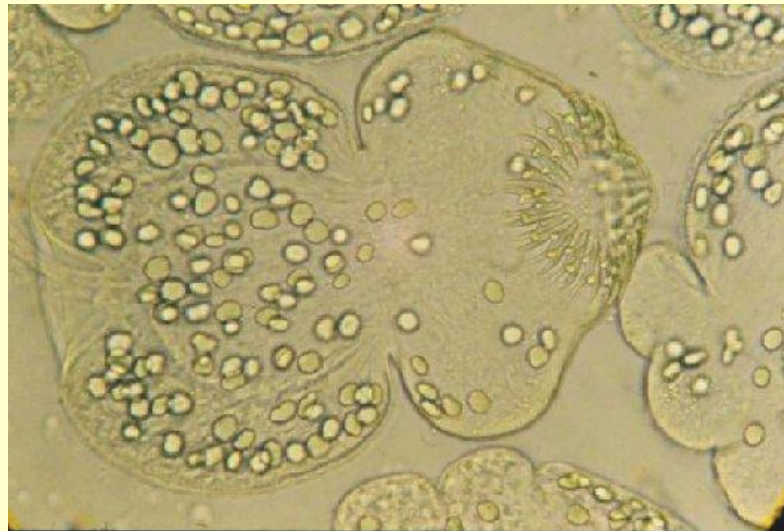


Rupture of cyst

Spontaneous vs evoked (trauma)



Dissemination of infection; anaphylactic shock



Diagnostics

- Eosinophilia not remarkable: up to 15%
- **Serology:**

Limitations:

- 10% patients with hepatic cyst and 60% with pulmonary cyst – false negative result
- Children up to 3 years – false negative result

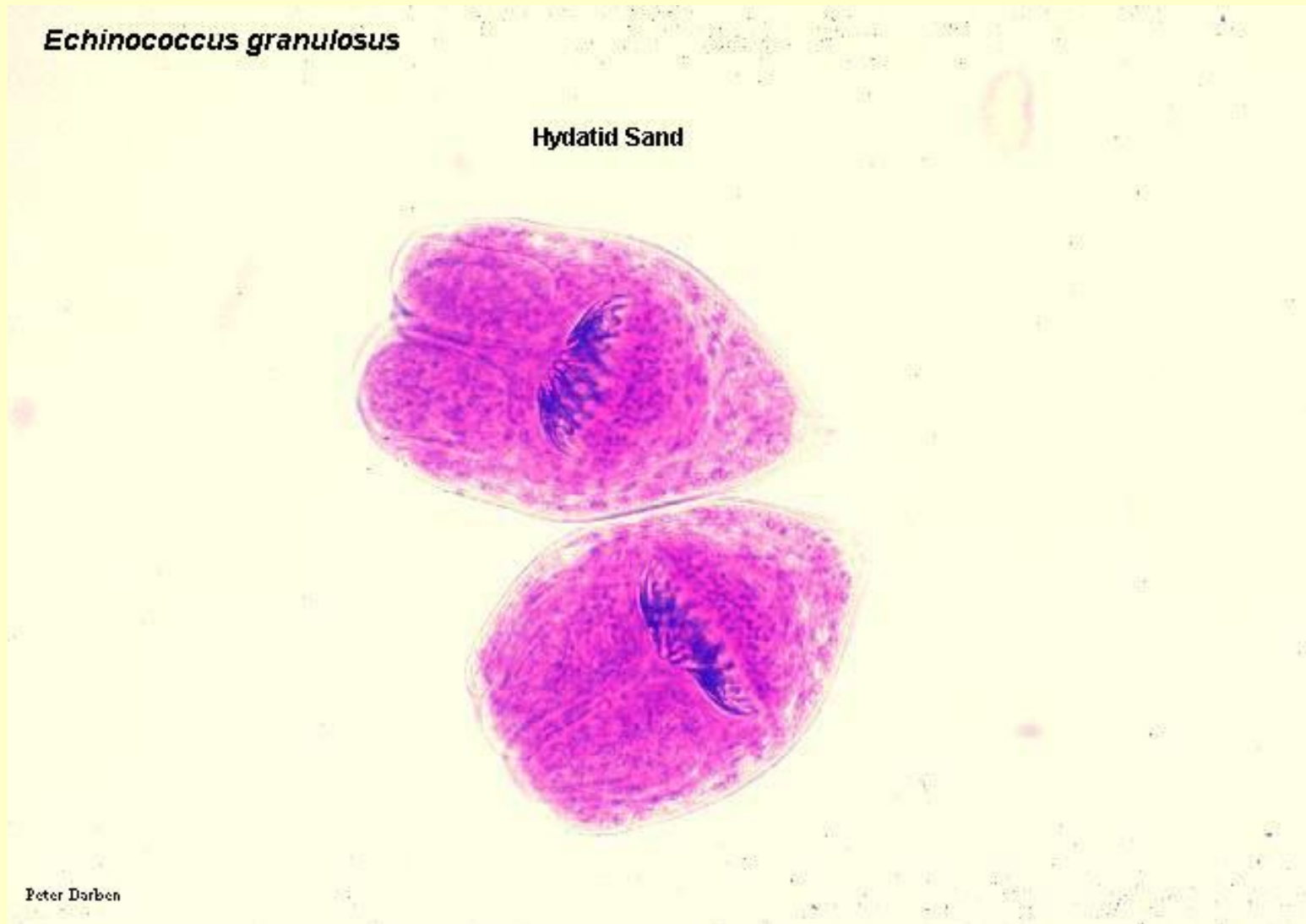
Imaging techniques – interpretation

- Simple cyst with clearly defined wall and uniform anechogenous composition - unlikely
- Cysts with remarkable **different structure of wall** - likely
- Cysts with **septae** – likely
- Solid heterogenous mass difficult to distinguish from granuloma or tumour, **calcification** – points out to echinococcosis

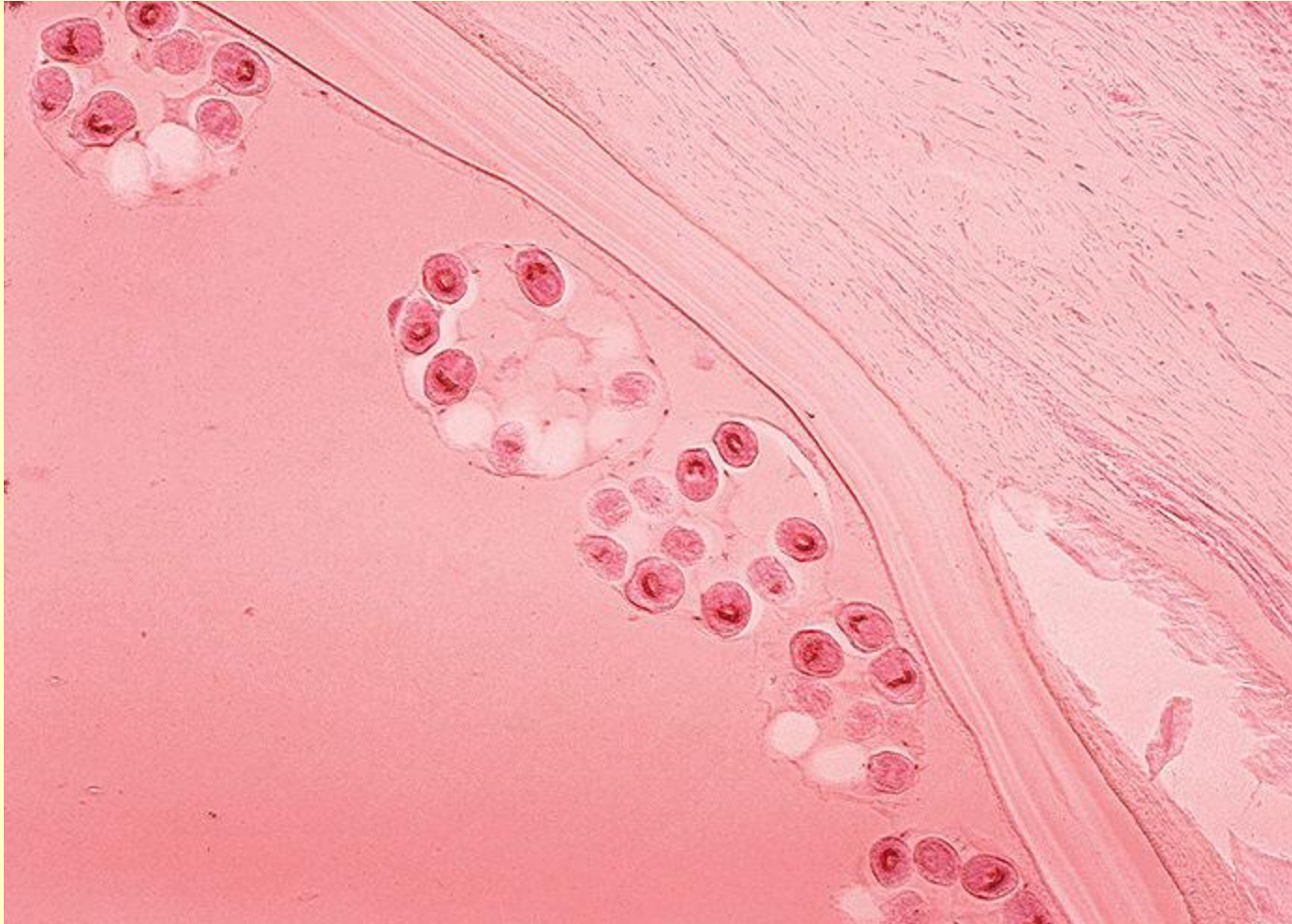
Casoni skin test, replaced currently by serology



Hydatic sand (movement of protoscolexes in cyst)



Hydatid cyst, histology



Therapy

Albendazole — 10 mg/kg 4 weeks, 12 cycles with 2 weeks breaks
(sono, CT)

Praziquantel

Surgical removal

PAIR — Percutaneous Aspiration Injection (hypertonic saline, scolical solution, alcohol)

