

Welcome to ***Parasitic Diseases***  
Fall 2008

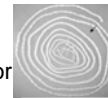
Dickson Despommier, Ph.D.  
Charles Knirsch, MPH, MD  
Josh Stillman, MD

**Helminths (Worms)**

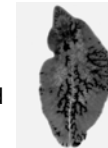
Nematodes - non-segmented round worms



Cestodes - segmented flat worms



Trematodes - non-segmented flat worms



**Parasite**

Any organism that takes metabolic advantage of another organism

Viruses  
Rickettsiae  
bacteria  
Fungi  
Protozoa\*  
Helminths\*

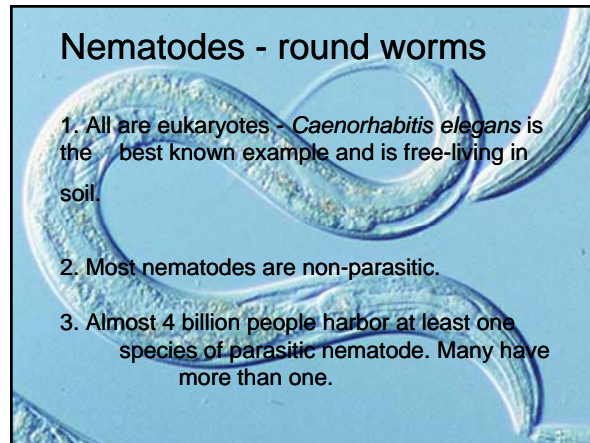
Nematodes - round worms  
Cestodes - segmented flat worms  
Trematodes - non-segmented flat worms

Arthropods\* - six and eight-legged critters

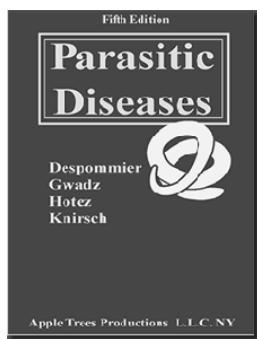
\* Covered in *Parasitic Diseases*

**Nematodes - round worms**

1. All are eukaryotes - *Caenorhabditis elegans* is the best known example and is free-living in soil.
2. Most nematodes are non-parasitic.
3. Almost 4 billion people harbor at least one species of parasitic nematode. Many have more than one.



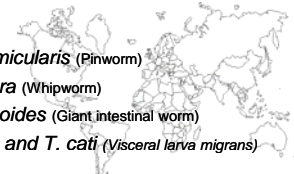
Highly Recommended textbook\*  
Available at bookstore.  
Still only \$69.95



\* Lots of really gross pictures!!!

**Geohelminths**

*Enterobius vermicularis* (Pinworm)  
*Trichuris trichiura* (Whipworm)  
*Ascaris lumbricoides* (Giant intestinal worm)  
*Toxocara canis* and *T. cati* (*Visceral larva migrans*)  
Hookworms  
*Ancylostoma duodenale*  
*Necator americanus*  
*Strongyloides stercoralis*



## Site Location For Helminth Infections Of Humans

Alimentary tract	197 species
Cavities, organs, and tissues	107 species
Circulatory system	21 species
Skin and tissues	56 species

## Going Green !

(with brown and yellow)



## Helminths That Infect Humans

Acanthocephala	7 species
Nematoda	138 species
Nematomorpha	24 species
Platyhelminthes	173 species
Digenea	113 species
Eucestoda	57 species
Turbellaria	3 species
<b>TOTAL</b>	<b>342 species</b>

## Food Plus



Why so many parasite species

We live everywhere.

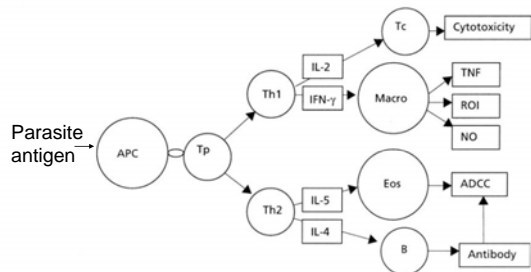
We eat everything.



This should never happen!

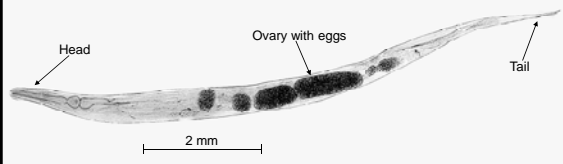


## Immunity and Parasitism



Worm infections elicit Th2 protective immune responses.  
Protozoan infections elicit Th1 protective immune responses.

## Adult Female *Enterobius vermicularis*

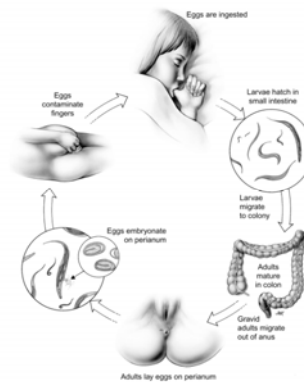


## Cytokines and Immunity to Parasites

Th1 cytokines	Th2 cytokines	Pro-inflammatory cytokines	Counter-inflammatory cytokines	Cytokines that can lead to pathology (e.g. increased vascular permeability, tissue damage, circulatory collapse, multi-organ failure etc.)
IFN- $\gamma$ *	IL-4*	IL-12	IL-4	IL-1
IL-2	IL-5*	IL-15	IL-10	IL-6
IL-3	IL-3	IL-18	TGF- $\beta$	IL-8
TNF- $\alpha$	IL-13	IFN- $\gamma$		IL-12
TNF- $\beta$	IL-6			TNF- $\alpha$
GM-CSF	IL-10			MIF
	TGF- $\beta$			

\*most important in immune expulsion of protozoa and worms

## *Enterobius vermicularis*



## Helminths: Nematoda

*Enterobius vermicularis*  
(Pinworm)

## Heavy Infection of *Enterobius vermicularis*

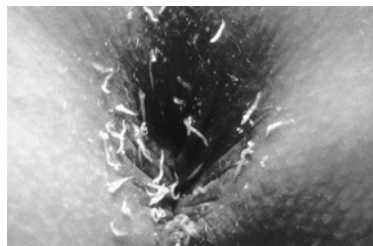
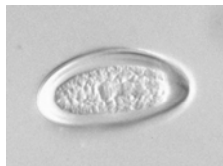


Photo: Martin Weber, MD, Children's Hospital, Hannover Medical School Hannover, Germany

### Eggs of *Enterobius vermicularis*



Unembryonated



Embryonated

Larva

### Diagnosis:

Eggs found on microscopic examination of clear sticky tape.



larva

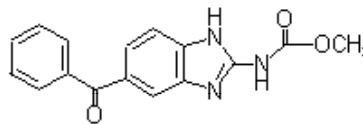
### *Enterobius vermicularis* in appendix



Alae

### Drug of Choice:

#### Mebendazole



### Mode of Action:

De-polymerizes invertebrate tubulins, only.

### Clinical Disease:

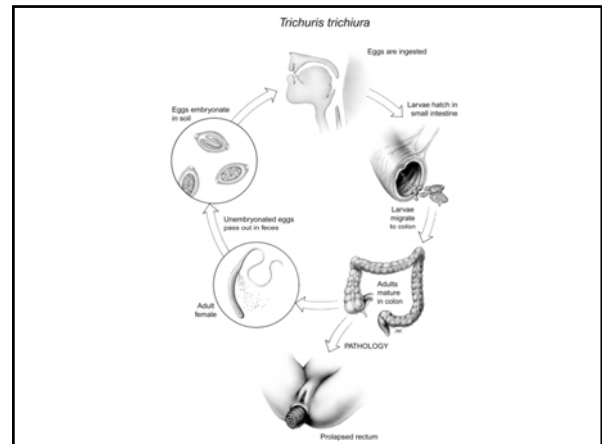
None

### Prevention and Control:

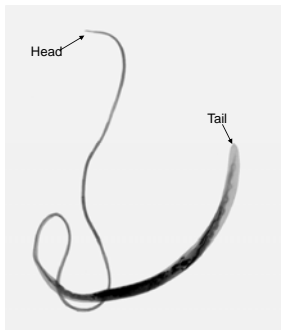
Prevention is difficult among children, especially those attending day care facilities and lower grades in school. We "out-grow" our pinworm infections once we reach puberty.

Helminths:  
Nematoda

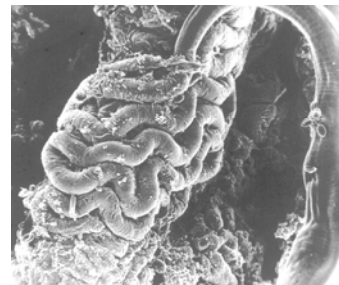
*Trichuris trichiura*  
(Whipworm)



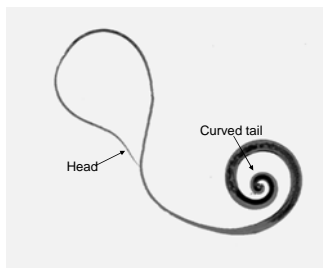
Female adult *Trichuris trichiura*



Adult *Trichuris muris* in situ (SEM)



Adult male *Trichuris trichiura*



Pathogenesis:

*Trichuris spp.* secrete a pore-forming protein that may play a role in anemia and diarrhea. Adult worms **do not** feed directly on blood or other host tissues. Mechanism of anemia still unknown.

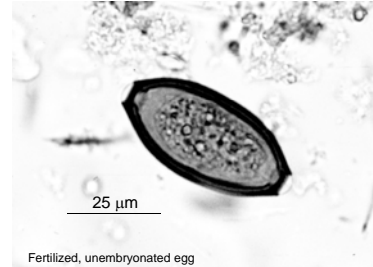
## Approaches to helminth-based therapy of IBD

World J Gastroenterol. 2008 Sep 7;14(33):5125-32.  
Helminth infections and intestinal inflammation.  
Wang LJ, Cao Y, Shi HN.  
Mucosal Immunology Laboratory, Massachusetts General Hospital, Building 114 16th Street,  
Room 3504, Charlestown, Massachusetts 02129, United States. shiha@helix.mgh.harvard.edu

Evidence from epidemiological studies indicates an inverse correlation between the incidence of certain immune-mediated diseases, including inflammatory bowel diseases (IBD), and exposure to helminths. Helminth parasites are the classic inducers of Th2 responses. The Th2-polarized T cell response driven by helminth infection has been linked to the attenuation of some damaged Th1 driven inflammatory responses, preventing some Th1-mediated autoimmune diseases in host, including experimentally induced colitis. Helminth parasites (the porcine whipworm, *Trichuris suis*) have been tested for treating IBD patients, resulting in clinical amelioration of the disease. As a result, there is a great deal of interest in the research community in exploring the therapeutic use of helminth parasites for the control of immune-mediated diseases, including IBD. However, recent studies have provided evidence indicating the exacerbating effects of helminths on bacterial as well as non-infectious colitis in animal models. Therefore, a better understanding of mechanisms by which helminths modulate host immune responses in the gut reveal novel, more effective and safer.

## Diagnosis:

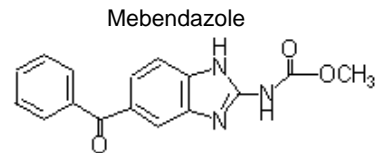
Microscopic examination of feces for eggs



## Prolapsed rectum with adult *Trichuris trichiura*



## Drug of choice:



Mode of Action:  
De-polymerizes invertebrate microtubules, only

## Clinical Disease:

1. Diarrhea
2. Anemia
3. Malnutrition (protein calorie deficiency?)

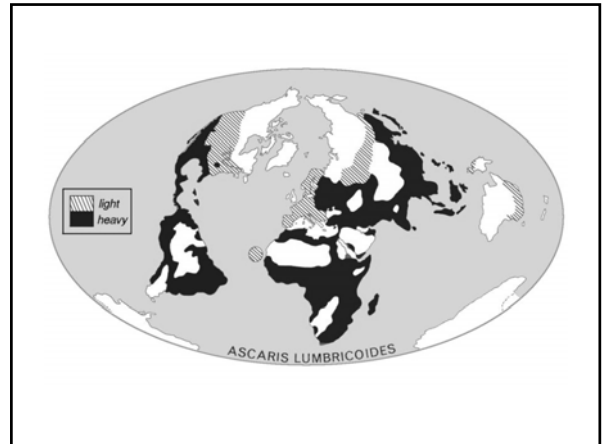
## Prevention and Control:

Sanitary disposal of feces

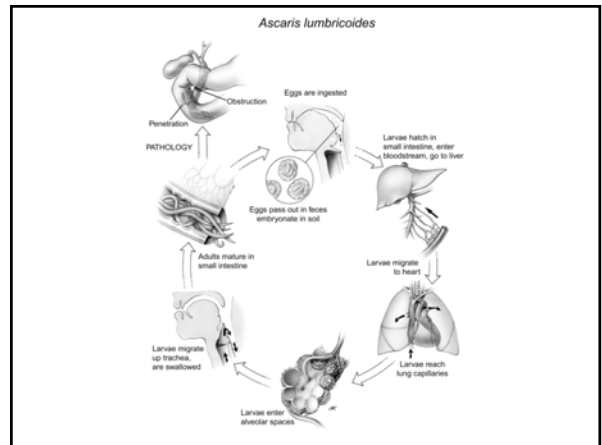
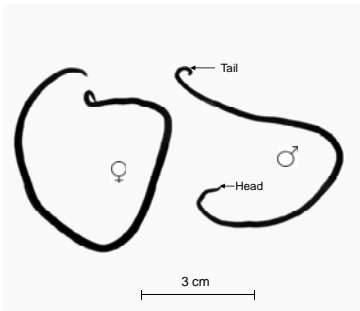


Helminths:  
Nematoda

*Ascaris lumbricoides*  
(Giant intestinal worm)



Adult *Ascaris lumbricoides*



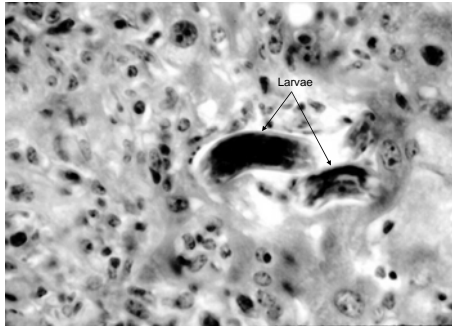
**“Jar-O-Worms”**  
Collected from one rural village in Bangladesh in a single day!



Cross section of adult *Ascaris lumbricoides*



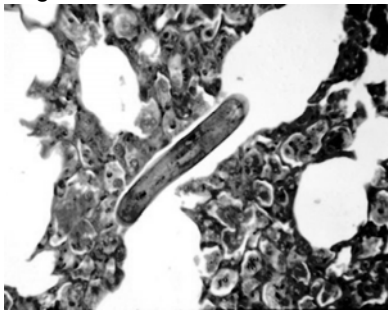
Larvae of *Ascaris lumbricoides* in liver



## Clinical Disease:

1. Light infections are asymptomatic as long as the adult worms do not migrate.
2. Heavy infection leads to:
  - a. protein calorie malnutrition - "failure to thrive" syndrome.
  - b. bowel obstruction.
  - c. aberrant migratory events.

Larva of *Ascaris lumbricoides* in lung



Child with heavy *Ascaris lumbricoides* infection



## Pathogenesis:

1. "Verminous" pneumonia, lung tissue damage due to migratory larvae.
2. Bowel obstruction - too many adult worms.
3. Parasite secretes trypsin inhibitor, prevents host from digesting proteins.
4. Aberrant migration of "irritated" adult worms to:
  - a. Ampulla of Vater
  - b. Common duct
  - c. Liver
  - d. Pharynx
  - e. Peritoneum

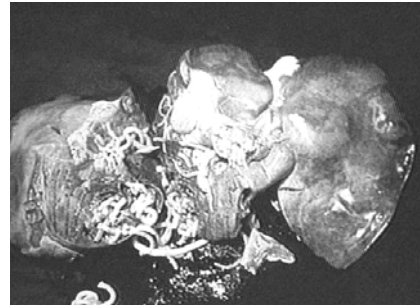
The result of successful treatment!



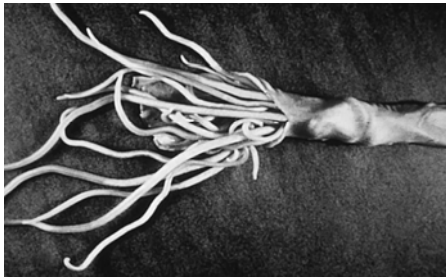
Infant with heavy *Ascaris* infection



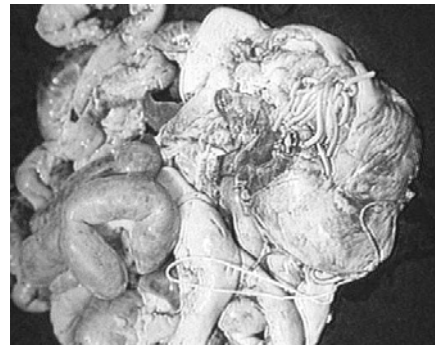
*Ascaris* adults in liver (fatal case)



Bolus of *Ascaris lumbricoides* in small intestine



Bolus of *Ascaris lumbricoides* (fatal case)

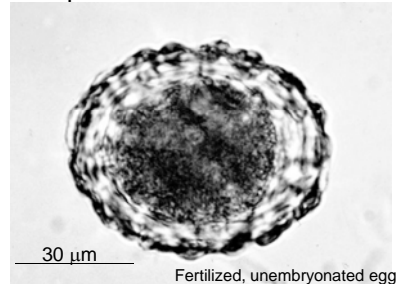


*Ascaris lumbricoides* adult in appendix



### Diagnosis:

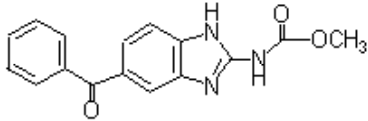
Microscopic examination of feces for eggs



Fertilized, unembryonated egg

## Drug of choice:

Mebendazole



Mode of Action:  
De-polymerizes invertebrate microtubules, only

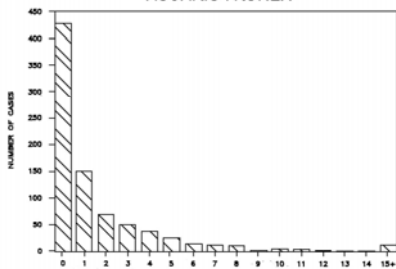
## Helminths: Nematoda



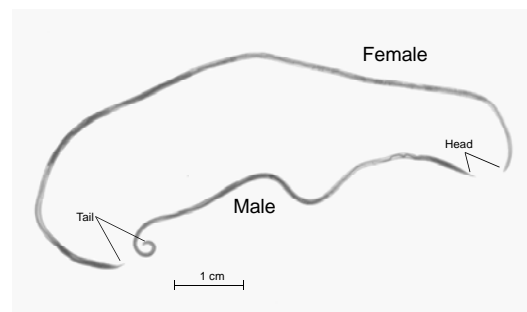
*Toxocara canis*  
*Toxocara cati*  
Visceral and ocular larva migrans

## Medical Ecology

ASCARIS : KOREA



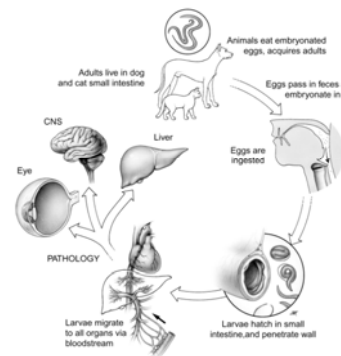
Adult *Toxocara canis*



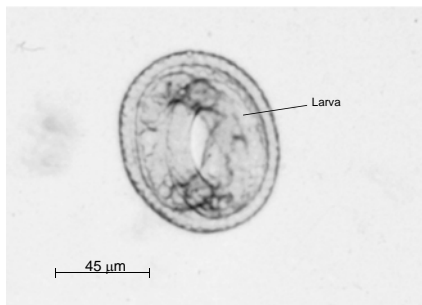
## Prevention and Control: Sanitary disposal of feces



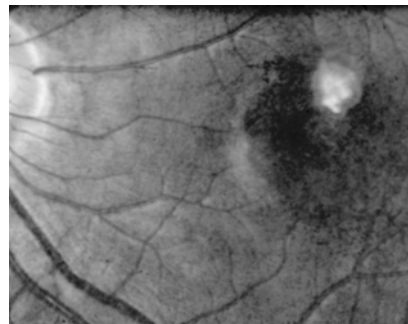
*Toxocara canis* and  
*Toxocara cati*



Embryonated egg of *Toxocara canis*



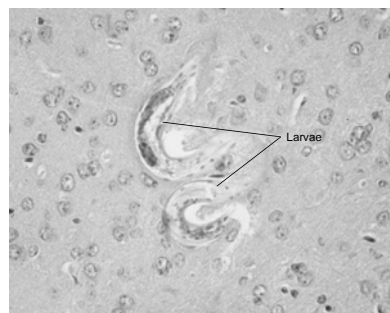
Granuloma in retina due to *Toxocara canis*



### Pathogenesis:

Tissue damage (systemic) due to migratory 3rd stage larva

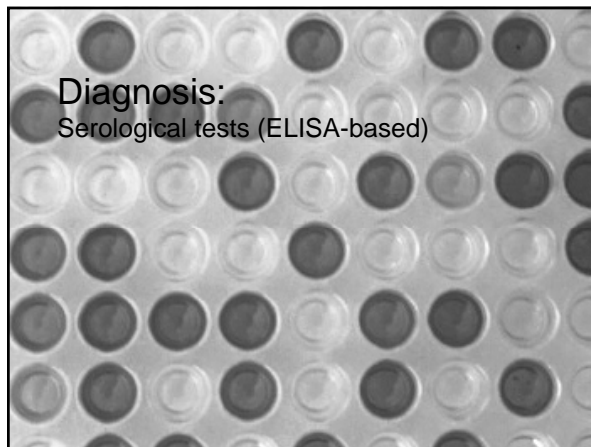
Larvae of *Toxocara canis* in mouse brain



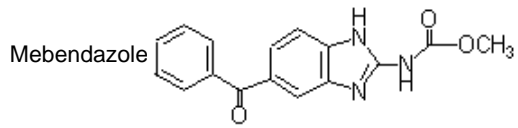
### Clinical Disease:

1. Fever
2. Loss of visual acuity
3. Blindness
4. Learning disabilities

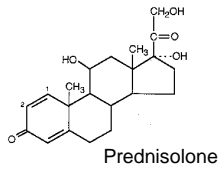
Diagnosis:  
Serological tests (ELISA-based)



## Drugs of choice:



## Steroids



## Prevention and Control:

Sanitary disposal of dog and cat feces



Not practical



Pooper-scooper



Potty-trained cat!

## Prevention and Control (cont'd):

2. Periodically de-worm pets.
3. Cover public sand boxes at night.

