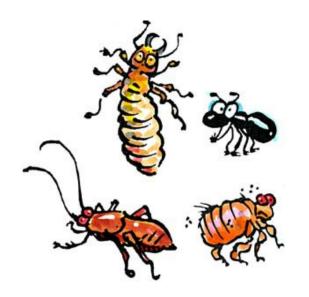
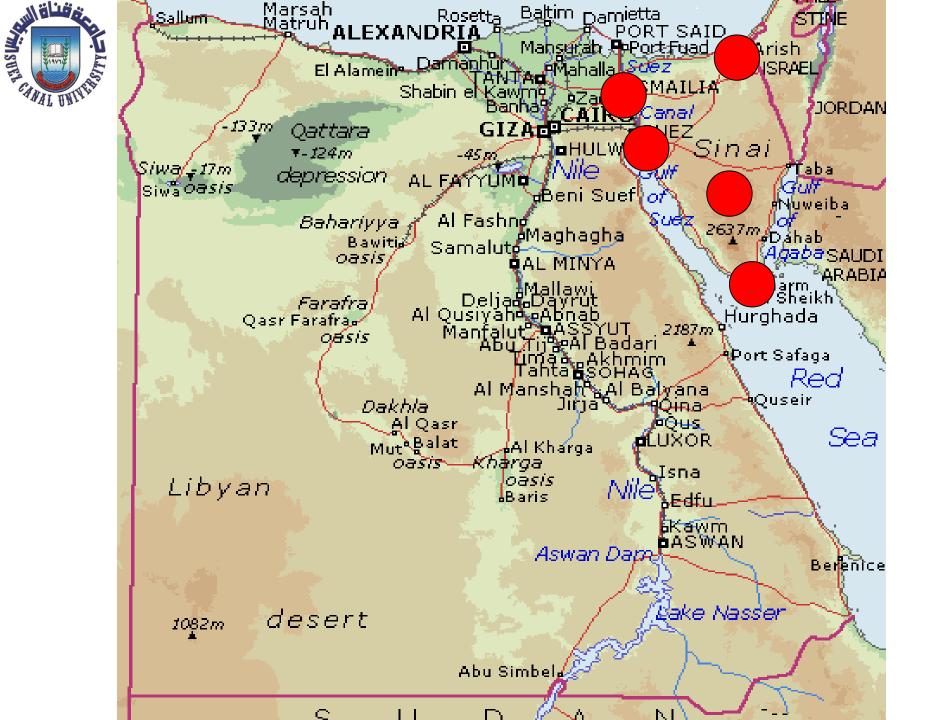


# New trends in veterinary insecticides



# By Prof. Dr. Mostafa Fayez

Professor of Pharmacology Faculty of Vet. Medicine – Suez Canal University





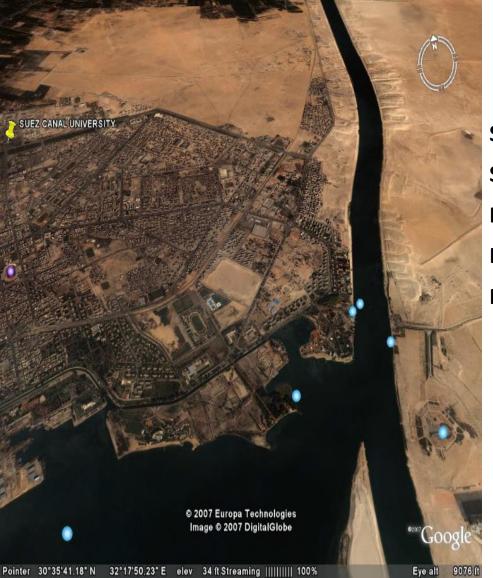
## Suez Canal University

# **A Community-Based Education Institu**





## **SUEZ CANAL UNIVERSITY**



Students	40.000,0
Staff Members	4.000,0
Employees	7.700,0
Faculties	22
Provinces	4

## The Japan International Cooperation Agency (JICA)

- Workshops for sub-Sahara African physicians at SCU and funded by JICA.
- **Combating Infectious Diseases in Sub-Sahara Africa.**
- Medical crisis management

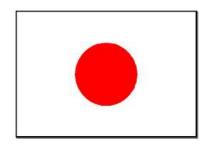


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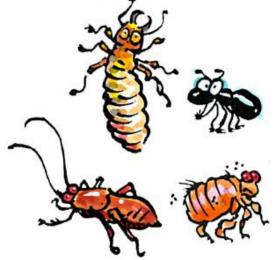
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# Insecticide

Any substance or mixture of substances intended for preventing, destroying, repelling, or mitigating insects.

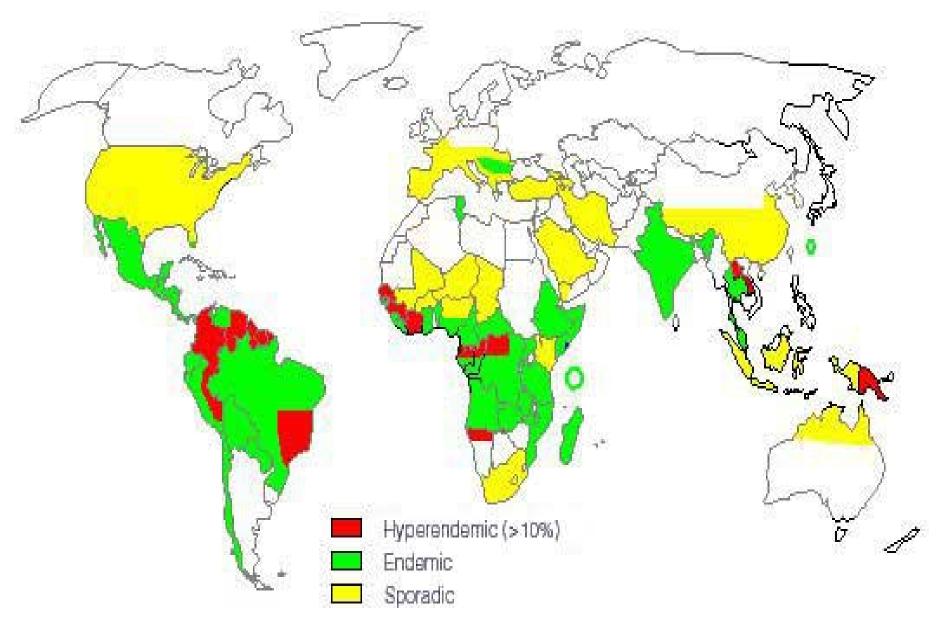


# Insecticides represent an \$8 billion industry





# Epidemiology











# **Classes of Insecticides**

#### **Botanical Insecticides**

- · Sabadilla
- · Nicotine
- · Quassia
- Unsaturated isobutylamides
- · Ryanodine
- Naphtoquinones
- Rotenone

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- · Sweet flag
- · Marigolds
- · Pyrethrum
  - Azadirachtin
- **Essential oils**

#### Synthetic Insecticides

Organochlorines

Organophosphate

S

Carbamates

Pyrethroids

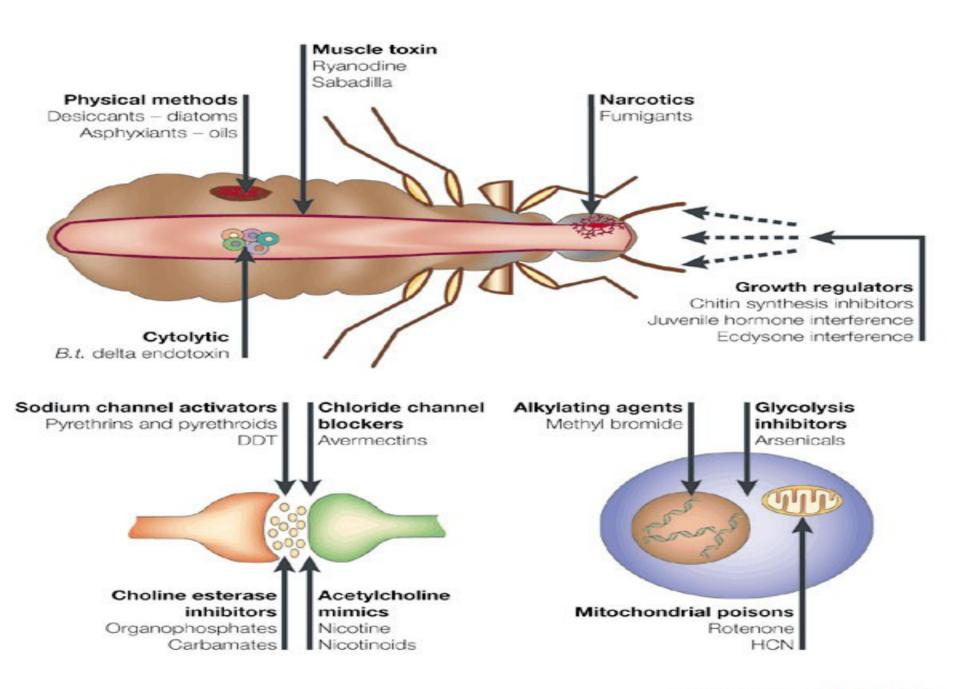
Neonicotinoids

#### **Biorational Formulations**

Growth regulators Pheromones Microbial formulations









Insecticides are one of the four building blocks of IPM

Chemica

Biologica



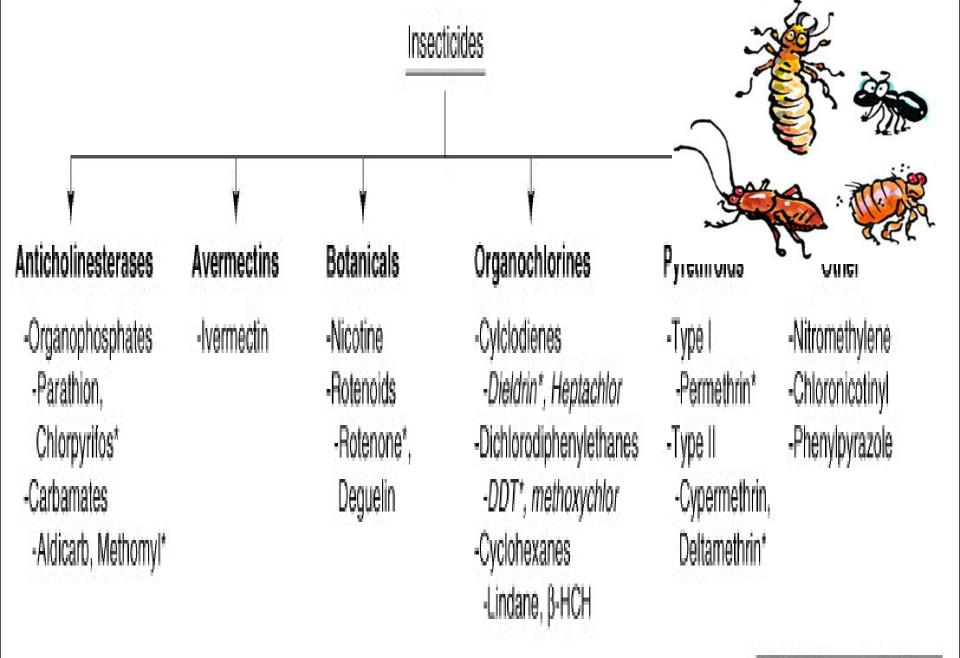
# Chemical Control: Insecticides











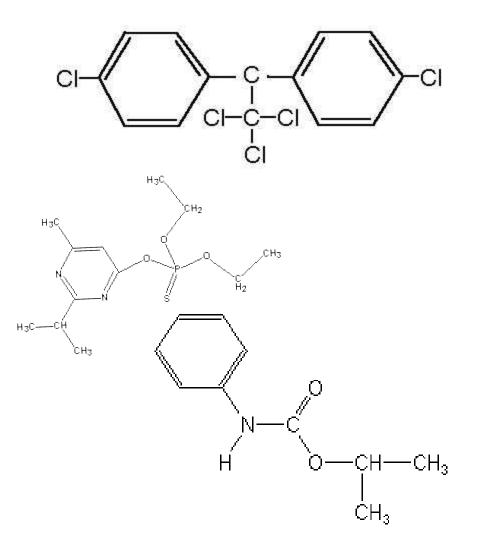
TRENDS in Pharmacological Sciences

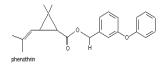
## Synthetic Insecticides

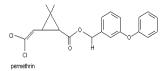
#### Synthetic Insecticides

Organochlorines Organophosphates Carbamates Pyrethroids

Neonicotinoids





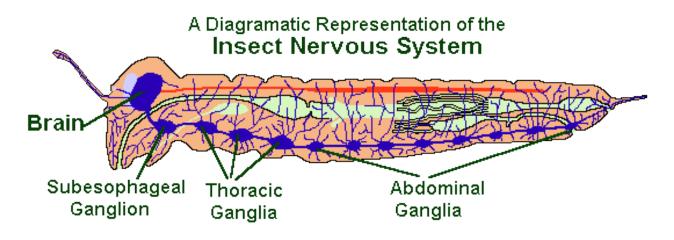


## **Nervous System**

• Why does this happen?

· Let's look at a "normal" nerve impulse.

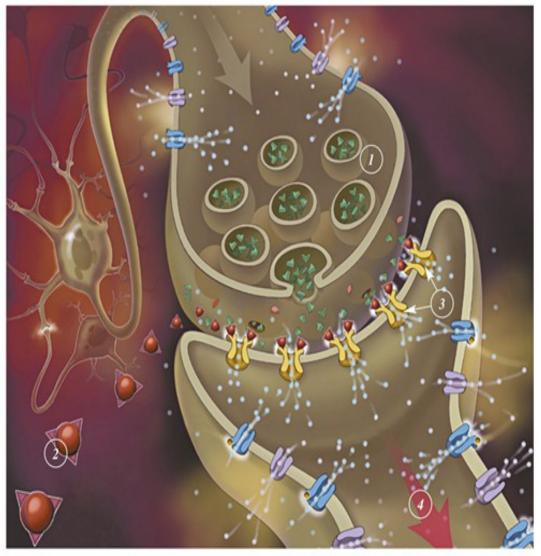
# Nervous system



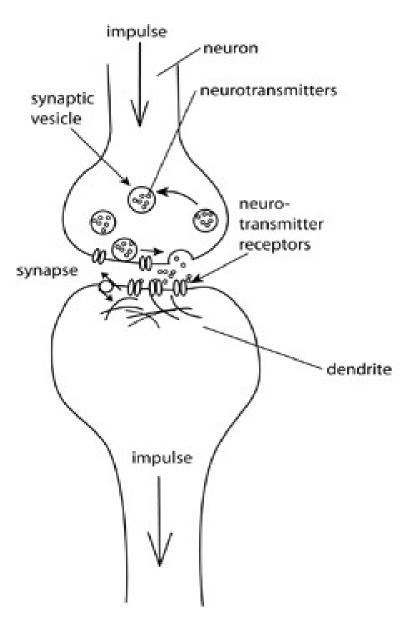


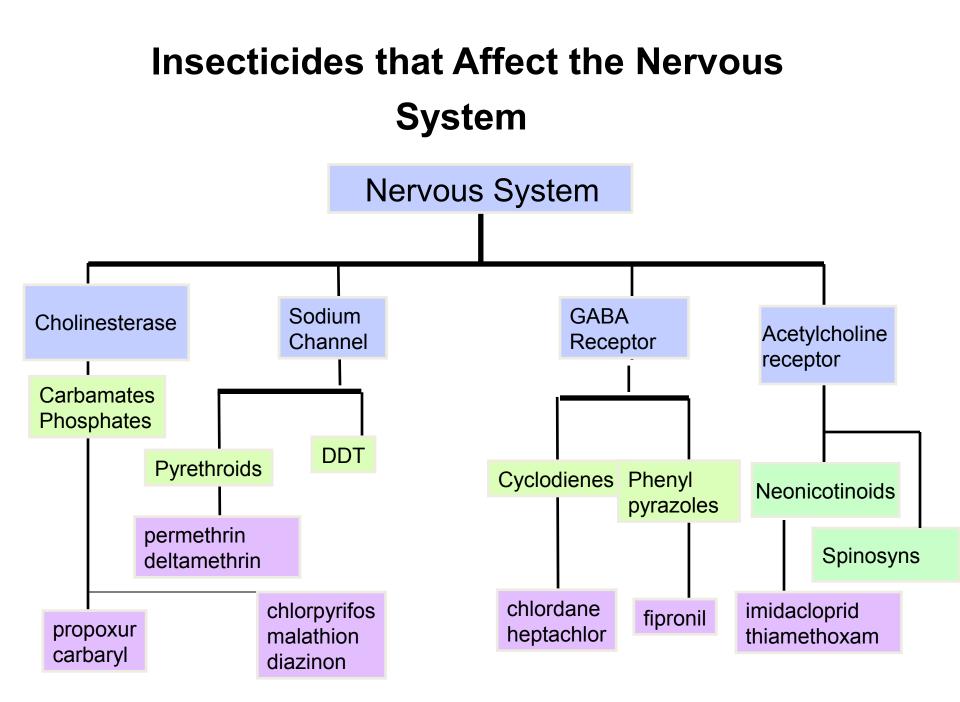
JEFF JOHNSON BIOLOGICAL & MEDICAL VISUALS

Affect synaptic transmission

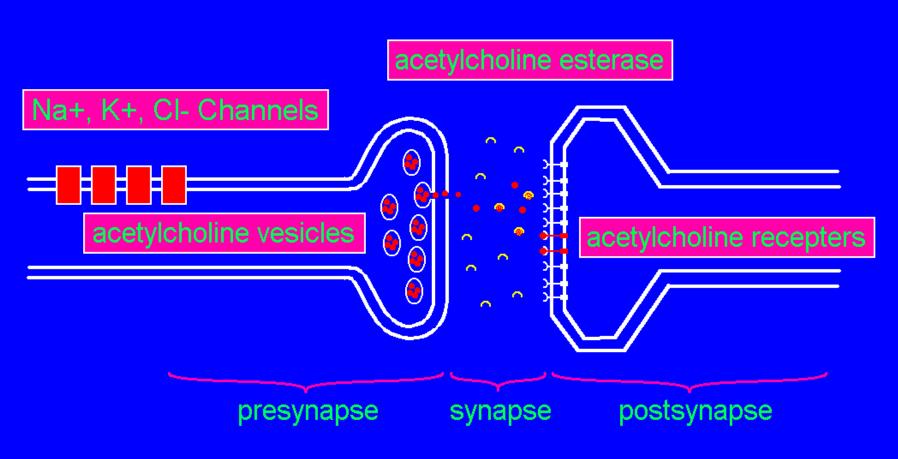


1 Vesicles of acetylcholine 2 Imidacloprid 3 Nicotinic receptors blocked open 4 Constant neuromuscular stimulation





### **Affects on Nerves**



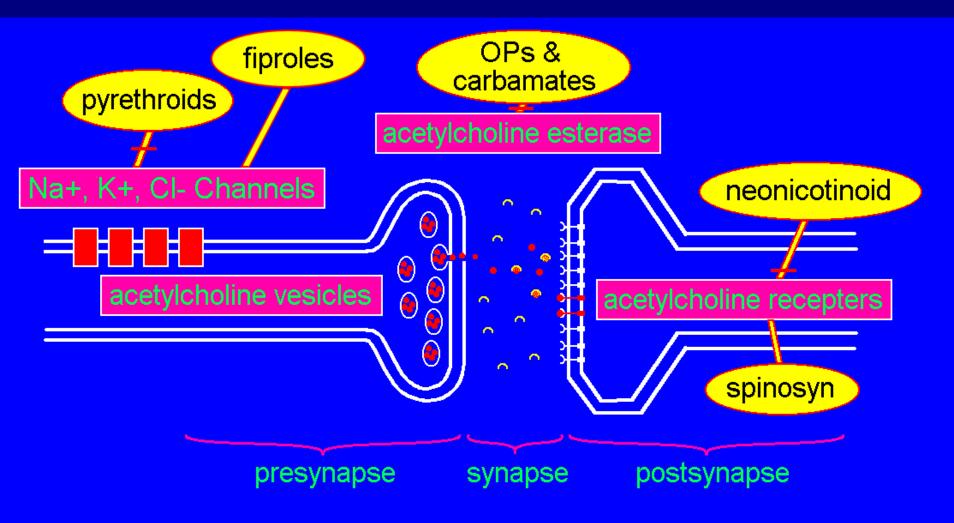
#### Notes:

Neural transmission is a complicated process that includes an actual electrical impulse that travels up and down nerve cords (axons) and chemical transmission between nerve cells (synapse).

The electrical transmission is accomplished by nerve cells maintaining an ion imbalance inside and outside the cell membrane. This is done through movement of sodium (Na+), potassium (K+) and chloride (Cl-) ions to opposite sides of the cell wall (sodium channel and GABA channels). When the electrical impulse passes, this electrical charge is switched, but almost immediately it's restored.

At the synapse, when the electrical impulse arrives, small packets (vacuoles) of acetylcholine are released within the synapse gap and almost instantaneously move to the receiving nerve (acetylcholine receptors) which starts another electrical impulse. To keep the acetylcholine from continually firing the receiving nerve, there is an enzyme (acetylcholine-esterase) that quickly destroys the acetylcholine, thereby resetting the synapse for the next nerve impulse.

### **Affects on Nerves**



#### Notes:

Organochlorine insecticides, pyrethoids as well as the new fiproles disrupt the sodium/potassium/chloride channel systems that maintains the electrical charge of nerve cell membranes. When this is disrupted, nerves can not properly transmit the electrical impulse.

Organophosphate and carbamate insecticides block the acetylchlineesterase enzyme which causes the receiving nerve to keep firing. This causes the affected animals to virtually twitch to death!

Neonicotinoids fill up the acetylcholine receptors, actually the nicotinicadetylcholine receptors (which insects have), thereby blocking neural transmission. Affected insects simply stop activity especially feeding, grooming and protective behaviors.

Spinosyns appear to act like acetylcholine, stimulating the receptors of the receiving nerve. The result is much the same as organophosphate or carbamate activity, but insects, again, are differentially affected by spinosyns.

# Modes of Action Nervous Systems

## Neural membrane disruption (ion transport disruption)

sodium/potassium ions <--> chloride ions (ion pumps) (organochlorines & pyrethroids) (fiproles = phenylpyrazoles)

# Modes of Action Nervous System

## Neural synapse disruption

acetylcholine (ACh)

(neural transmitter)



cholinesterase (ChE)

(neural transmitter eraser)

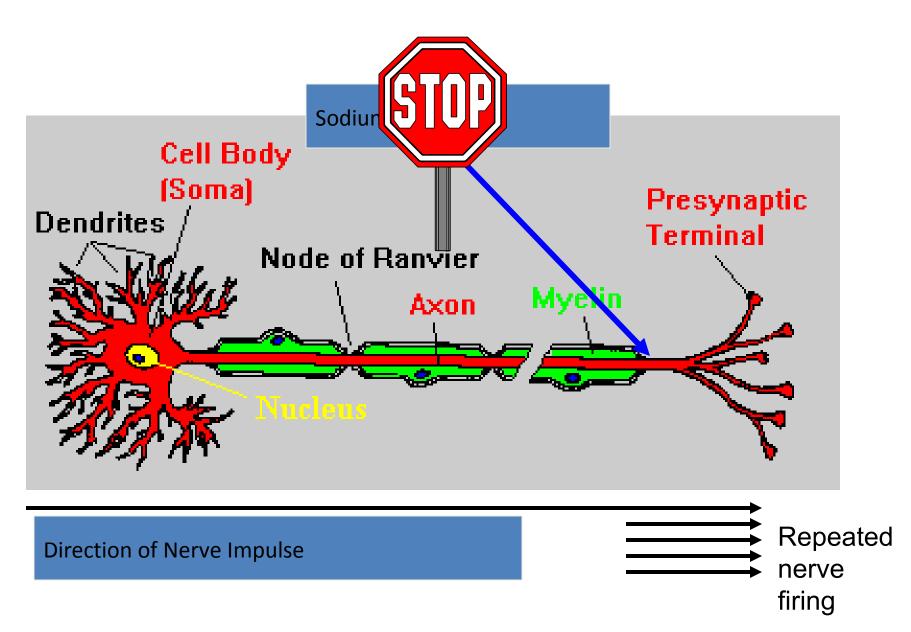
(organophosphates & carbamates - inhibit ChE) (alkaloids, like nicotine, mimic ACh)

# Modes of Action Nervous Systems

(stimulates)

(neonicotinoids and spinosyns)

## Nerves and Pyrethroids



# MOA: Pyrethroids

- · Axonic poisons
- Bind to a protein called the voltage-gated sodium channel
- Normally, the sodium channel opens, causing stimulation of the nerve and closes to end the nerve signal
- Pyrethroids find to the socium channel and prevent it from closing normally
- · Result: continuous nerve stimulation

• Symptoms: Tremors, uncoordinated movement From Valles and Koehler. 1998. Insecticides Used in the Urban Environment: Mode of Action. ENY-282

# Pyrethrins and Pyrethroids

Pyrethroids -- synthetically produced chemicals with similar chemistry and mode of action as pyrethrins, axonic poisons

- Type I
  - Short residual,
  - Fast flushing, and
  - Quick knockdown,
  - Negative temperature correlation: more effective at low temps
  - Ex.: Allethrin, d-phenothrin, resemethrin

- Type II
  - Long residual,
  - Slower flushing, and
  - Slower knockdown,
  - Better killing power than type 1
  - Positive temperature correlation
  - Ex.: Permethrin,
- Synergists -- used to deactivate mixed function oxidases (MFOs) within insects

## MOA: Carbamates and OPs

- Synaptic poisons
- Carbamates and organophosphorus insecticides bind to an enzyme called

acetylcholinesterase (AChE)

- · AChE is found at the nerve synapse
- · FACTAFESIGNER COBLEGESTOPS ARE IN PUTSE TO SET OF STREET TO SET OF SET OF STREET TO SET O

# Examples



#### Organochlorines

- Endosulfan
  - Gamma HCH
- Gamma BHC

#### Organophosphates



- -Diazinon
- -Fenitrothion
- -Dichlorvos
- -Dimethoate
- Malathion



Carbamates - Aldicarb -Carbofuran



- Pyrethroids -Tefluthrin
- Deltamethrin
- Lambda
- -cyhalothrin
- Permethrin
- Cypermethrin

#### Neonicotenoids

- -imidacloprid
- -nitempyram
- acetamiprid
- thiamethoxam

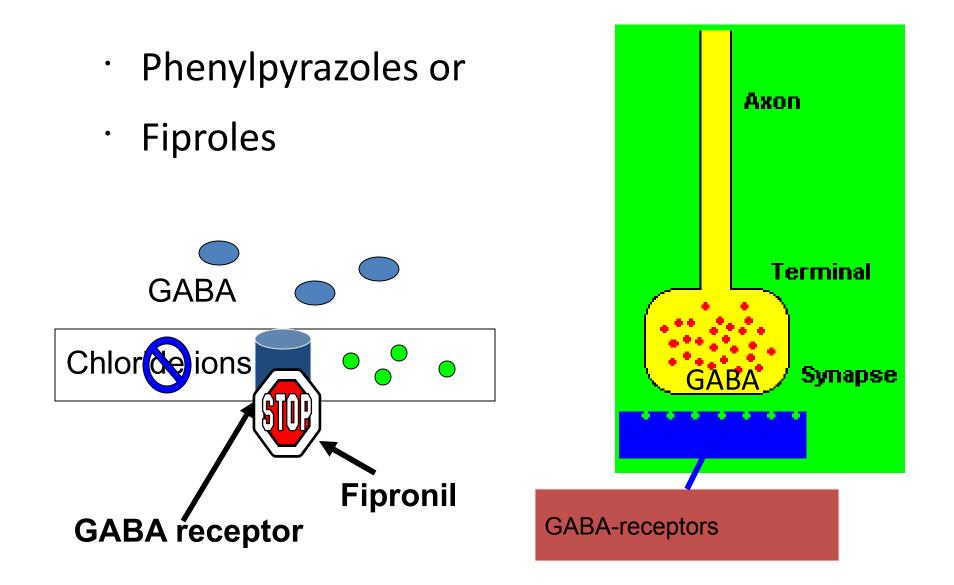
# Neonicotinoids

- Imidacloprid (subgroup: pyridylmethylamine)
  - Advantage pet treatment
  - Merit -- turf ornamentals
    - · Water soluble

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- Systemic in plants
- Premise termites
  - · Affects insects first by paralyzing mouthparts
- Bayer Advantage OTC
- Thiamethoxam (subgroup: nitroguanidine)
  - Used in crops

#### **GABA Receptor in Central Nervous System**



## MOA: GABA Antagonist

- Phenylpyrazole (fipronil), cyclodienes
- · Antagonists of GABA activated chloride channels
  - GABA Antagonists: chemicals that bind to but do not activate GAMMA-AMINOBUTYRIC ACID receptors, thereby blocking the actions of endogenous GAMMA-AMINOBUTYRIC ACID or GAMMA-AMINOBUTYRIC ACID agonists.
- GABA can't bind to receptor

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- Doesn't activate channel (doesn't open)
- Interferes with (prevents) the passage of chlorine ions through the channel.
- Concerns with cross resistance with other GABAergic insecticides

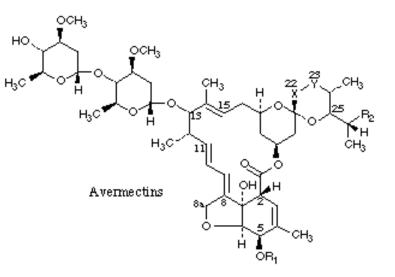
### MOA: GABA Agonist

- · Avermectins
- Bind to GABA (Gamma-Amino-Butyric Acid gated chloride channel
- · Opens channel
- · Chloride influx
- · Cannot reach threshold for action potential
- Causes membrane to hyperpolarize, making it less excitatory, decreasing nerve transmission
- · Result: flaccid paralysis and death

#### Avermectins

- · Abamectin
  - Avert -- cockroaches
  - Advance -- ants

Produced from soil microbials. Stomach poison for ants and cockroaches slow acting low toxicity to mammals



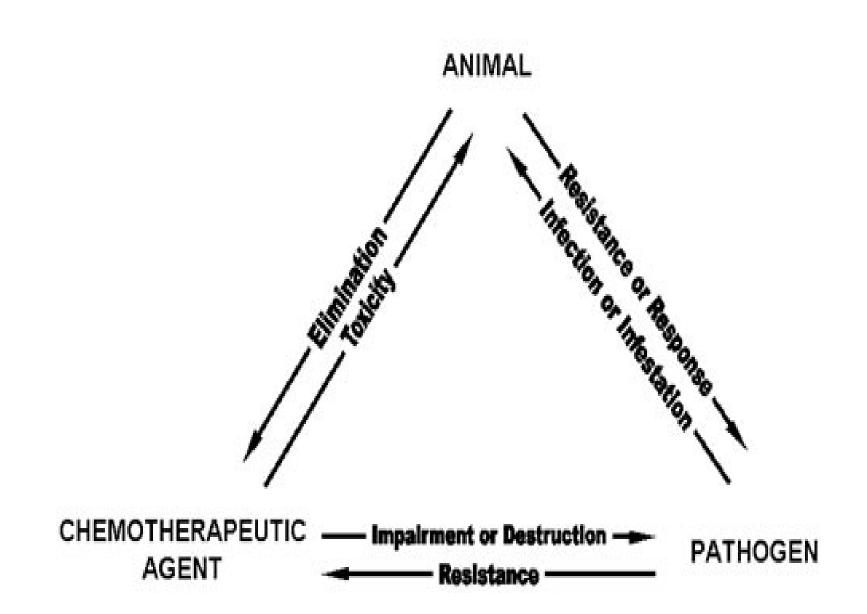
#### Avert Kill

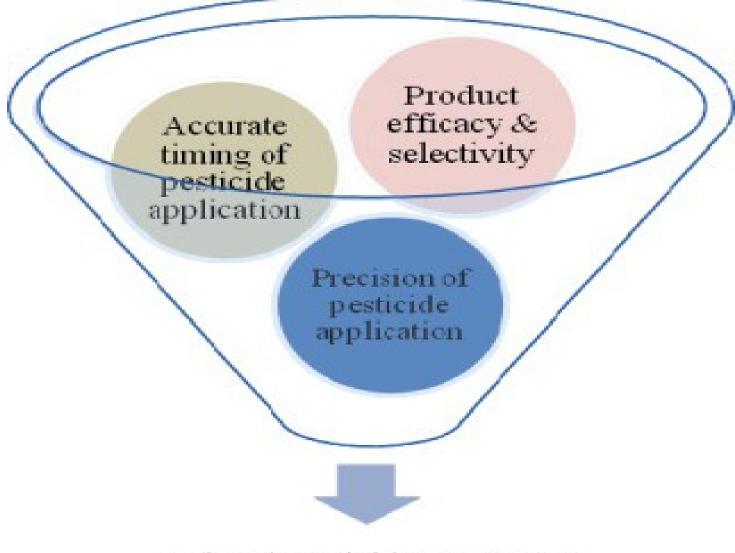
- · 22 hours to nerve effects
- · 40 hours to kill









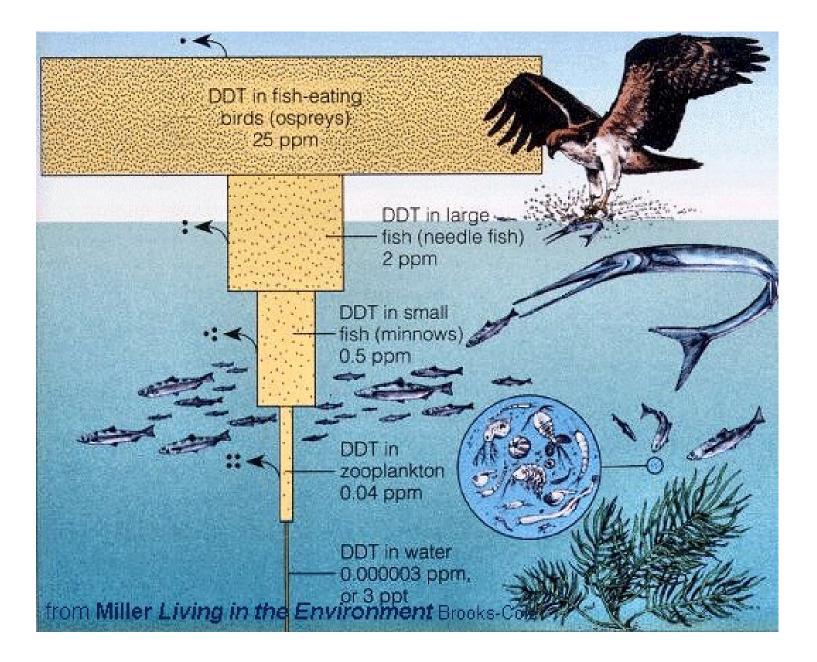


Rational Pesticide Use (RPU)

### Environmental hazard

Environmental hazard of insecticides is generally evaluated as a function of persistence often compared to effectiveness

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#### Public concern about Insecticide use

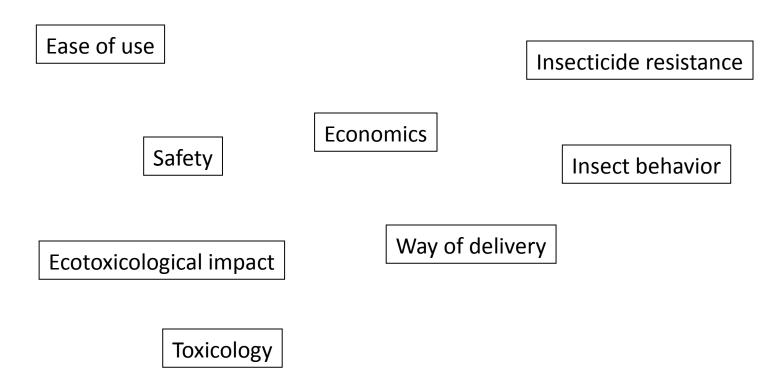
~100% of us have some insecticide residue in our bodies



# World Health Organization (WHO) Classification of Pesticides

	by Hazard (mg/kg body weight)			
Class	Oral		Dermal	
	Solids	Liquids	Solids	Liquids
la Extremely hazardous	5 or less	20 or less	10 or less	40 or less
Ib Highly hazardous	5 - 50	20 - 200	10 - 100	40 - 400
II Moderately hazardous	50 - 500	200 - 2000	100 - 1000	400 - 4000
III Slightly hazardous	Over 500	Over 2000	Over 1000	Over 4000

# For the Correct use of insecticides, we need to consider:



#### Thank you for your attention