

# ANS - TOXICITIES

## SOURCES OF CHOLINERGIC BLOCKERS

- Veterinary Medicines
  - Atropine, glycopyrrolate
  - Hyoscine, propantheline
- Plants
  - Belladonna, Datura,
  - Solanaceae (nightshades, potatoes)
  - Mushrooms (*Amanita panterina*)



# ANS - TOXICITIES

## SOURCES OF MUSCARINIC AGONISTS

- Veterinary Medicines
  - Pilocarpine, bethanecol
- Plants
  - Fungal toxin in red, white & alsike clovers (toxin-slaframine)
  - Mushrooms (*Amanita muscaria*)



# ANS - TOXICITIES

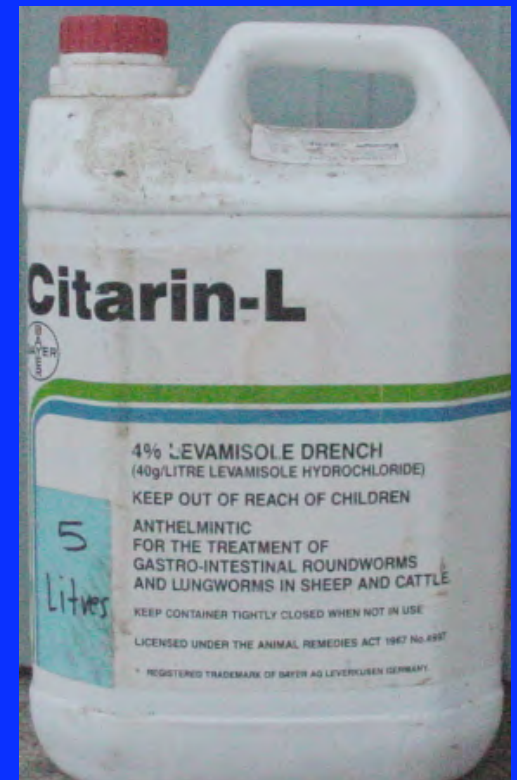
## SOURCES OF CHOLINESTERASE INHIBITORS

- Veterinary Medicines
  - Physostigmine, neostigmine (NMB reversal)
- Plants
  - Blue-green algae [anatoxin-a(s)]
- Cholinesterase Inhibitors
  - Organophosphates
  - carbamates

# ANS - TOXICITIES

## SOURCES OF NICOTINIC AGONISTS

- - Toxicants with Nicotinic Effects
  - Plants
    - Tobacco (*Nicotiana*)
    - Poison Hemlock (*Conium maculatum*)
    - Sophora, Laburnum
  - Chemicals
    - Levamisole (anthelmintic)



# ANS - ORGANOPHOSPHORUS Toxicity

- SOURCES
  - Insecticides
  - Pesticides
  - Dips
  - Pour-ons
  - Flea Collars
  - Sprays
  - Anthelmintics



# ORGANOPHOSPHORUS Toxicity

- TOXICITY
  - HIGHLY VARIABLE 100'S OF OP Compounds
  - Rats
    - LD 50 ranges from <1 mg/kg to > 4 grams/kg
  - Birds and Fish very sensitive

# ORGANOPHOSPHORUS Toxicity

- MECHANISM OF ACTION:
  - Inhibition of acetylcholinesterase at cholinergic receptors  
(IRREVERSIBLE when aged)
  - Inhibition of acetylcholinesterase (RBCs)

# ORGANOPHOSPHORUS Toxicity

- CLINICAL SIGNS (MUSCARINIC):
  - SLUDGE
    - Salivation
    - Lacrimation
    - Urination
    - Defaecation
    - Dyspnoea
    - Emesis



# ORGANOPHOSPHORUS Toxicity

- CLINICAL SIGNS (MUSCARINIC):
  - sweating
  - brady or tachy cardia depending on adrenaline release
  - pinpoint pupils (usually)
  - nasal discharge

# ORGANOPHOSPHORUS Toxicity

- CLINICAL SIGNS (NICOTINIC)
  - Tremors
  - Weakness
  - Paralysis

# ORGANOPHOSPHORUS Toxicity

- CLINICAL SIGNS (CNS):
  - Nervousness
  - Apprehension
  - Ataxia
  - Convulsions
  - Coma
  - Small animal:  $\pm$  seizure, hyperactive, hyperreflexive
  - Large animal: rarely seizure,  $\pm$  hyperactive

# ORGANOPHOSPHORUS Toxicity

- Muscarinic Signs: SLUDGE
- Nicotinic Signs: Muscle Tremors
- CNS: Anxiety, hyperactivity, clonic-tonic seizures

# ORGANOPHOSPHORUS Toxicity

## Intermediate Syndrome

- CATS and DOGS
- anorexia, diarrhoea, weakness,
- muscle tremors,
- abnormal posture and behaviour,
- clonic-tonic seizures

# ORGANOPHOSPHORUS

## Induced Delayed Neuropathy

### OPIDN

#### Mechanism of Action

- Caused by inhibition of neuropathy target esterase (NTE)
- Loss of myelin and axons in the spinal cord

#### Known Substances Causing OPIDN:

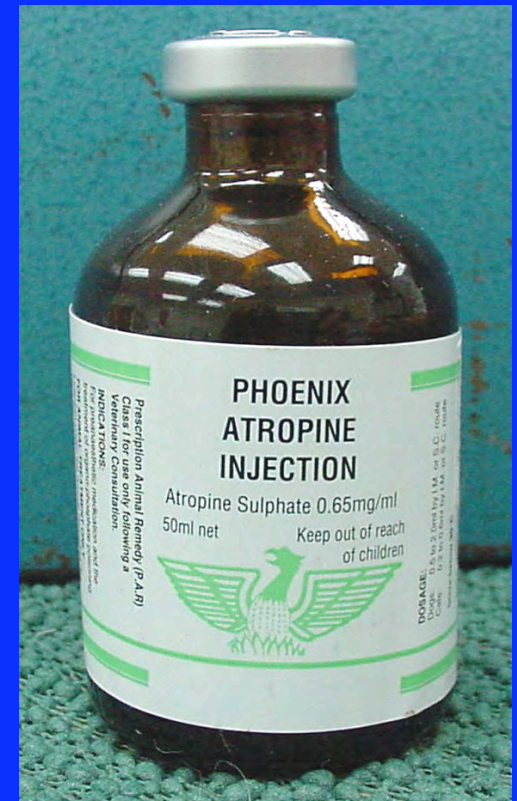
- ∞ leptofos, fenitrothion, trichlorfon, trichloronat and others

# ORGANOPHOSPHORUS Toxicity

- DIAGNOSIS
  - History
  - Garlic odour?
  - decreased acetylcholinesterase activity
  - Test dose of 0.02 mg/kg atropine

# ORGANOPHOSPHORUS Toxicity

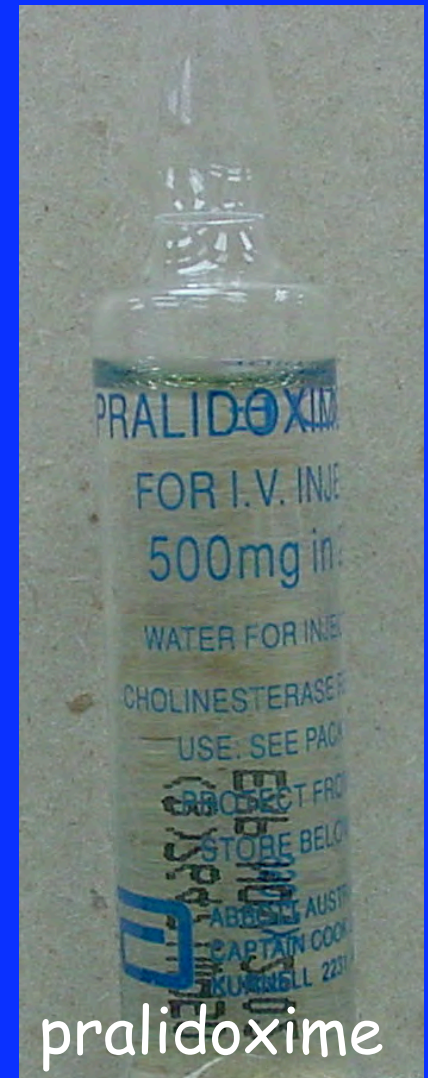
- TREATMENT:
  - DECONTAMINATE  
(dermal vs oral exposure)
  - Atropine sulphate
    - Part given Intravenously,
    - the rest Subcutaneously
      - 0.25-1 mg/kg





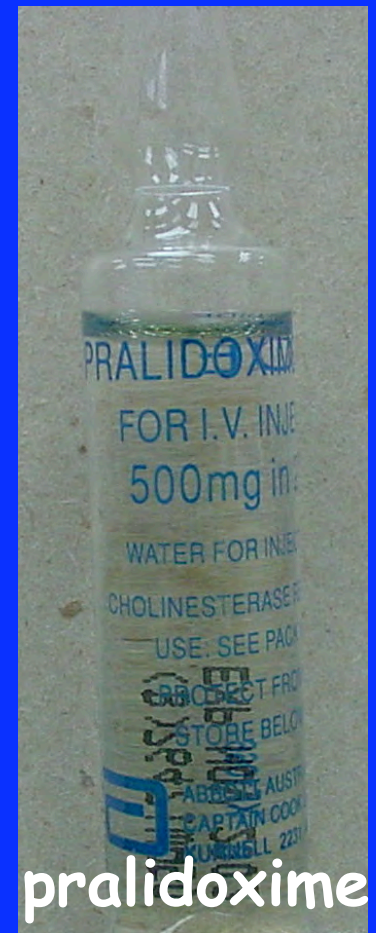
# ORGANOPHOSPHORUS Toxicity

- TREATMENT:
  - 2-PAM (Protopam chloride or aka pralidoxime chloride)
  - SUPPORTIVE CARE
    - Fluid therapy? Oxygen?
    - Seizure control-diazepam (Valium)



# ORGANOPHOSPHORUS Toxicity Intermediate Syndrome

- TREATMENT:
  - 2-PAM (Protopam chloride or  
aka pralidoxime chloride)
  - SUPPORTIVE CARE



pralidoxime

# CARBAMATES

## SOURCES:

- ∞ Carbaryl - insecticides
- ∞ Slug and Snail bait-MESUROL
  - ∞  $LD_{50} = 25 \text{ mg/kg}$



# CARBAMATES

- MECHANISM OF ACTION

SAME AS OPs except reversible binding to acetylcholinesterase

# CARBAMATES

- CLINICAL SIGNS:
  - Similar to Organophosphorus compounds

# CARBAMATES

- TREATMENT
  - Similar to Organophosphorus compounds  
EXCEPT DO NOT USE 2-PAM
  - 2-PAM is not necessary and may be harmful

# ANS Toxicities

## ORGANOPHOSPHATES DDX

- Amitraz
- Pyrethrins
- Cationic Surfactants e.g. benzalkonium chloride
- Garbage intoxication (endotoxins)

# OPs and Carbamates

## KEY POINTS

- ∞ SOURCES-numerous
- ∞ Do NOT accumulate (in fat) \*
- ∞ RAPIDLY EXCRETED
- ∞ Comp Inhib ACETYLCHOLINESTERASE
- ∞ "AGING" OF OP-ENZYME complex



# OPs and Carbamates

## KEY POINTS

∞

POTENTIATION:

Phenothiazine tranquilizers (e.g. ACP)

30 day wait after exposed to OPs

Blocks acetylcholinesterase



# OPs and Carbamates

## KEY POINTS

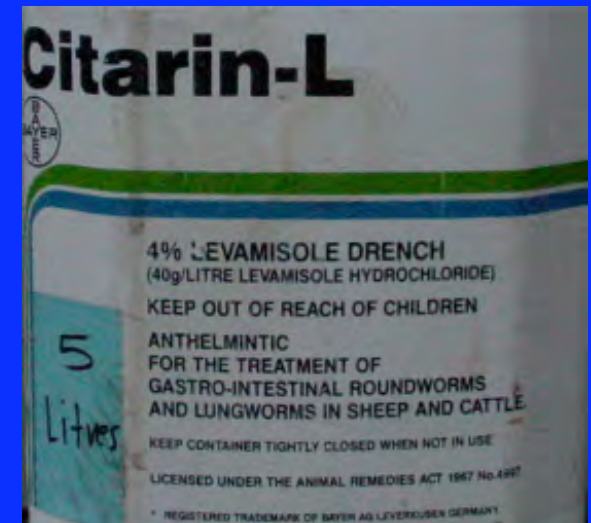
### POTENTIATION:

Levamisole:

- nicotinic stimulation

Aminoglycosides:

- blocks acetylcholinesterase



# OPs and Carbamates

## KEY POINTS

- ∞ Muscarinic: SLUDGE + other clinical signs
- ∞ Nicotinic and CNS effects
- ∞ ATROPINE AND 2 - PAM (oxime)

(2-PAM is not necessary with carbamate poisoning)

# OPIDN

- ∞ Cattle drenched with trichloronat
- ∞ Trichloronat - pasture insecticide
- ∞ OPIDN Onset 2-3 weeks after drenching