SOURCES OF CHOLINERGIC BLOCKERS

- Veterinary Medicines
 - · Atropine, glycopyrrolate
 - · Hyoscine, propantheline



- Plants

- · Belladonna, Datura,
- Solanaceae (nightshades, potatoes)
- Mushrooms (Amanita panterina)



SOURCES OF MUSCARINIC AGONISTS

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

SOURCES OF CHOLINESTERASE INHIBITORS

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

SOURCES OF NICOTINIC AGONISTS

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



ANS - ORGANOPHOSPHORUS Toxicity



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







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

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

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

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

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

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

- Muscarinic Signs: SLUDDE
- Nicotinic Signs: Muscle Tremors
- CNS: Anxiety, hyperactivity, clonictonic seizures

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

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

- TREATMENT:
 - DECONTAMINATE

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

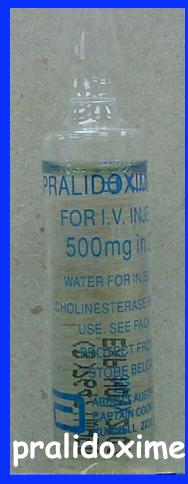


- TREATMENT:
 - 2-PAM (Protopam chloride or aka pralidoxime chloride)
 - SUPPORTIVE CARE
 - · Fluid therapy? Oxygen?
 - · Seizure control-diazepam (Valium)



ORGANOPHOSPHORUS Toxicity Intermediate Syndrome

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



SOURCES:

∞ Carbaryl - insecticides

Slug and Snail bait-MESUROL
 ∞LD₅₀ = 25 mg/kg



MECHANISM OF ACTION

SAME AS OPs except reversible binding to acetylcholinesterase

- · CLINICAL SIGNS:
 - Similar to Organophosphorus compounds

- 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)

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

∞ POTENTIATION:

Phenothiazine tranquilizers (e.g. ACP)

30 day wait after exposed to OPs

Blocks acetylcholinesterase



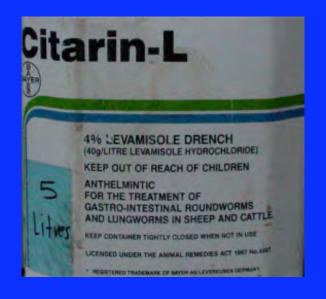
POTENTIATION:

Levamisole:

-nicotinic stimulation

Aminoglycosides:

-blocks acetylcholinesterase





- Muscarinic: SLUDDE + 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