

Myotoxic Plants

Poisonous Plant Class ADVS 586

Bryan Stegelmeier April 3, 2008

PPRL 2008

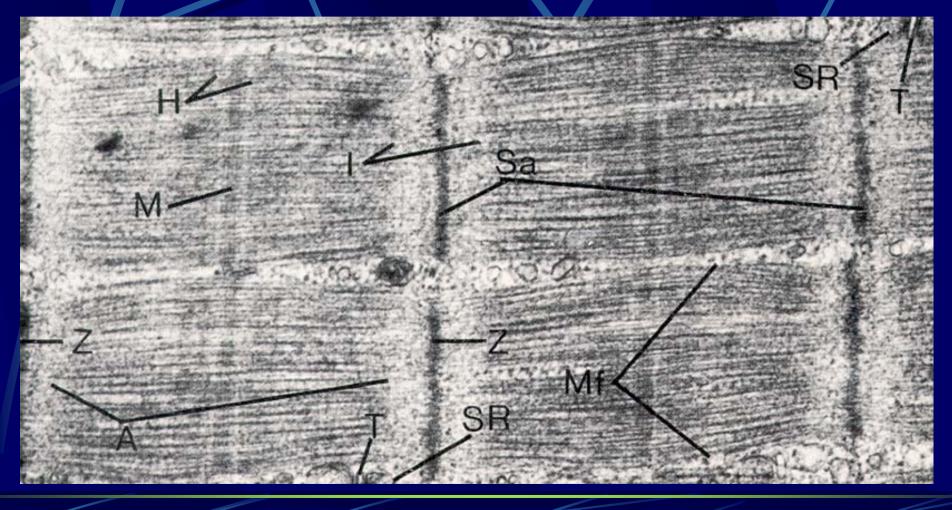
Striated Muscle

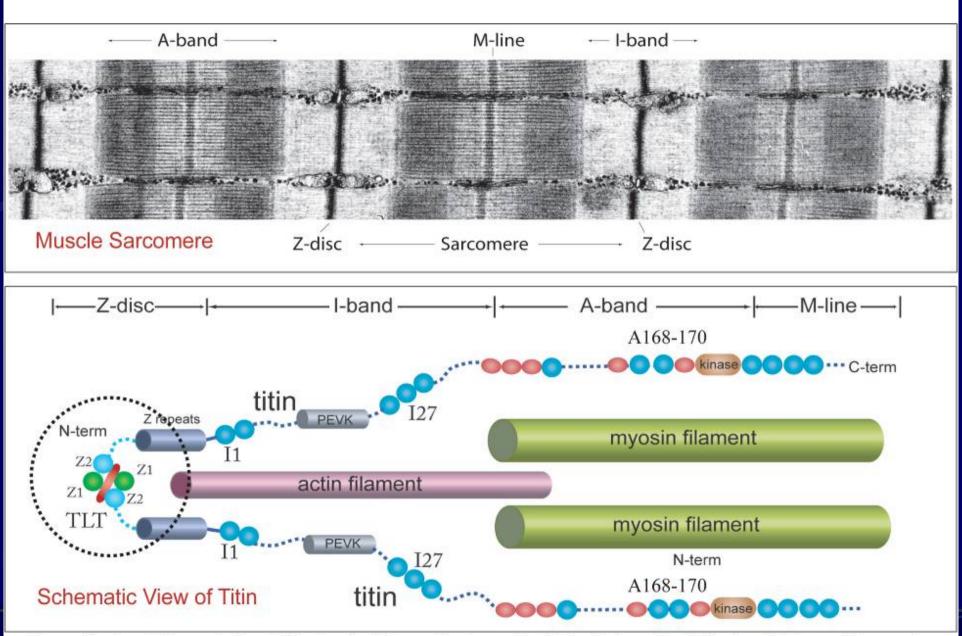
Types of muscleMitochondriaSarcolemna



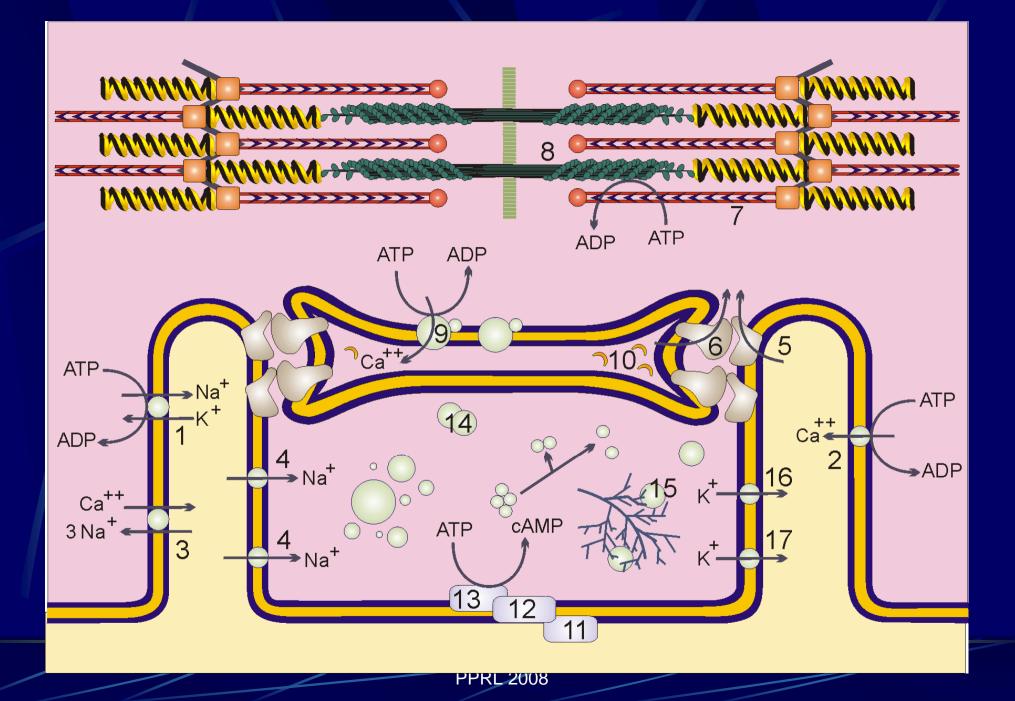
PPRL 2008

Ultrastructure





Theoretical and Computational Biophysics Group, Beckman Institute, University of Illinois at Urbana-Champaign



Myotoxic Plants

- Muscle structure and physiology
- Clinical and histologic lesions

Myotoxic Plants

- Thermopsis montana
- Eupathorium rugosum
- Haplopappus or Aplopappus spp.
- Cassia occidentalis O. obtusifolia
- Kwarwinskia humboldtiana
- Gossypium spp.
- Lathyrus spp.
- Vicia villosa
- Solanum spp. (enzootic calcification)
- Cardioglycoside Containing Plants
 - Digitalis purpurea
 - Nerium oleander
 - Convallaria majalis and C. montana
 - Apocynum spp.
 - Adonis aestivalis
 - Rhododendron spp.
 - Kalmia spp.
- Pieris japonica and P. floribunda
- Other potential myotoxic plants



Clinical Signs

Anorexia, depression, droopy ears

- Reluctant to stand or move
- Swollen hard muscle
- Walk with slow, labored gait
- Weakness, trembling, ataxia
- Recumbency, coma, death



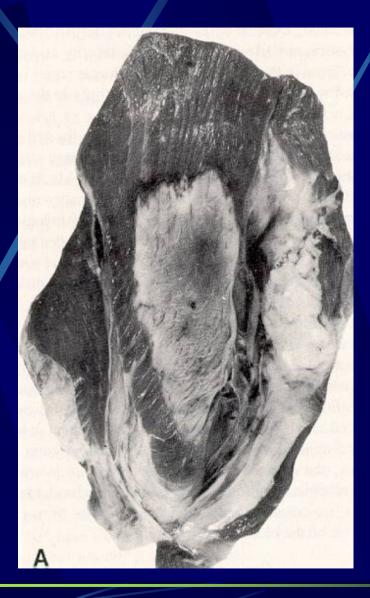
Biochemical Changes

AST
CPK
K
Myoglobinuria
Secondary changes

Gross Lesion

- Hard swollen muscles
 Pale streaking in muscle
 Secondary changes

 Disuse atrophy
 - Concretive beart for
 - Congestive heart failure
 - Nephrosis
 - Hepatic lipidosis



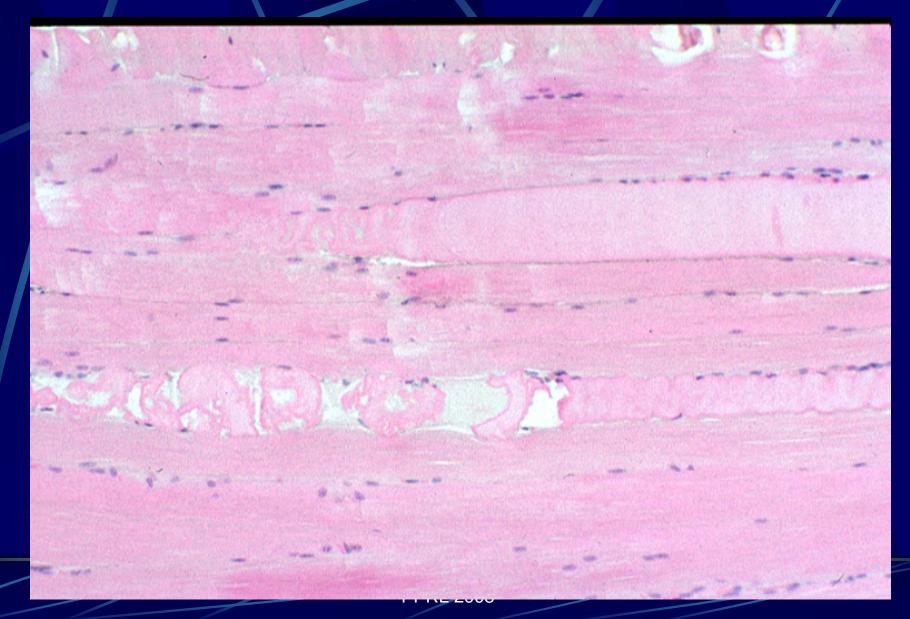
PPRL 2008

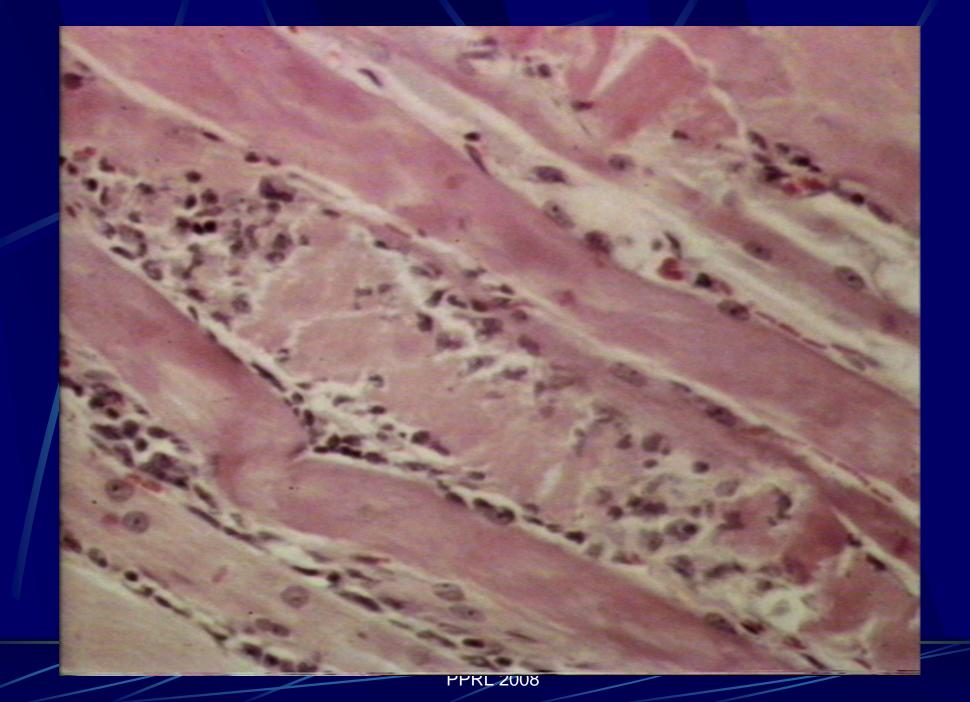


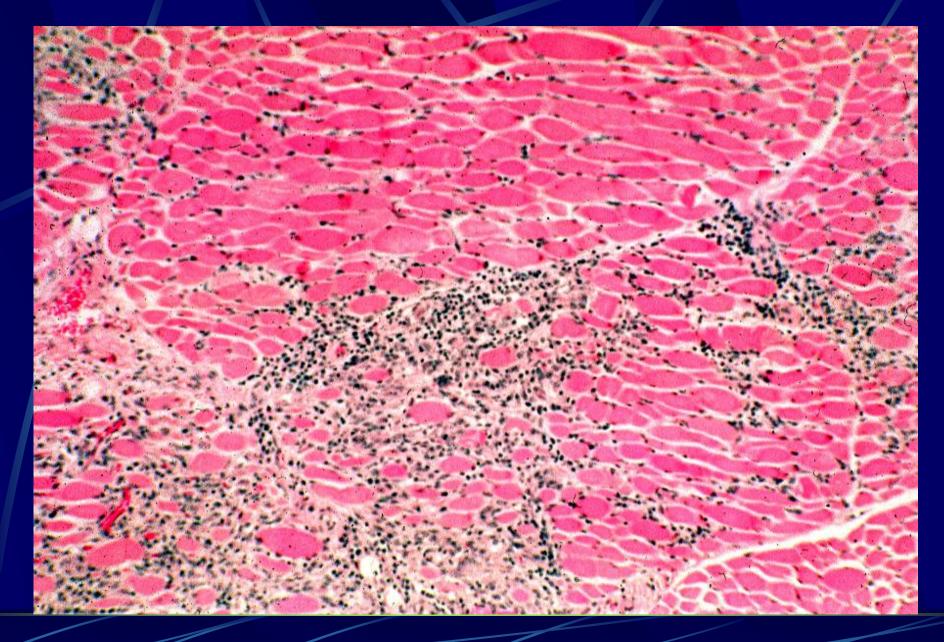


PPRL 2008

Histologic Lesions







PPRL 2008

Thermopsis montana of *T. rhombifolia*- Golden Banner, Mountain Thermopsis, False Lupine, Yellow Pea

- A perennial pea like plant with a rhizomatous root system and erect, branching stems that reach a height of 30 to 46 cm.
- Alternate with three leaflets (lupine has 5+).
 - Bright yellow flowers in dense racemes from the leaf axils
- Densely haired, erect seed pods that are straight (*T.* montana) or curved (*T.* rhombifolia).





MO ID OR WA NE UT CO

- Quinolizidine alkaloids:
 - n-methylcytisine
 - cytisine
 - 5,6 dehydrolupamine
 - thermopsine
 - Anagyrine

1 g/kg BW for 2 to 4 days



PPRL 2008

Signs and Lesions



- Depression, weakness, trembling, recumbency and death
- Edema, arched back, swollen eyelids
- Increased serum enzymes
- Muscle degeneration and necrosis

Eupathorium rugosum- snakeroot, white snakeroot, richweed



 Epidemics of weakness, nausea, prostration, death
 Milk sickness, 'the slows' (Mrs. Lincoln, General McClellan) 'trembles' in cattle

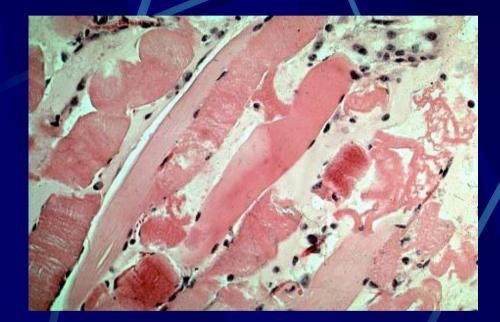
100 years to connect disease with plant

1927 Trematol

- Trematol (mixture of tremetone, dehydroytremetone, dihydroytremetone and hydroxytremetone)
- Cyrochrome P450 activated and quickly detoxified
- Green, dry and frosted plant are toxic
- Lipid soluble results in relay of secondary toxicity
- Stiffness, depression, ataxia, sternal recumbency, anorexia, tremors, coma, death
- Horses develop CHF



Disease in Livestock



0.5-1.5% BW disease in 7-11 days
6 month old hay toxic
Lactating cows protected

- Histology
 - Myonecrosis
 - Hepatic lipidosis
 - Hemorrhages and congestion
 - Gastroenteritis

Jimmy Weed, Rayless Goldenrod, Burrow Weed Isocoma pluraflora (Isocoma wrightii), (Haplopappu

 An erect, sparsely branched, woody perennia growing to 1 meter high
 Sticky leaves are linear and

- Sticky leaves are linear an alternate
 - Yellow numerous flowers form small, terminal flat topped heads of 7 to 15 flowers



Alkaline soils of drier rangeland, river valleys, drainage areas, and irrigation canals TX, NM, AZ, and CO Horses, cattle, sheep and goats 1.5% bw toxic in cattle







Senna or Cassia spp.- coffee weed or coffee sena

- Troublesome weeds southeastern United States, Hawaii, Mexico. Opportunist annuals that grow in waste areas, roadsides, fence lines. Common as weeds of corn and soybean fields.
- Green and dry plants are toxic
- Poison horses, cattle, sheep and goats.



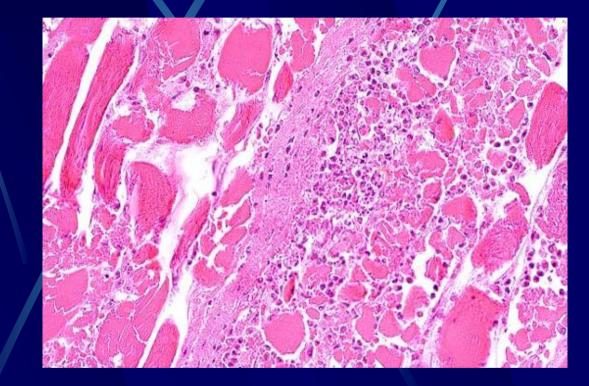


Cassia obtusifolia



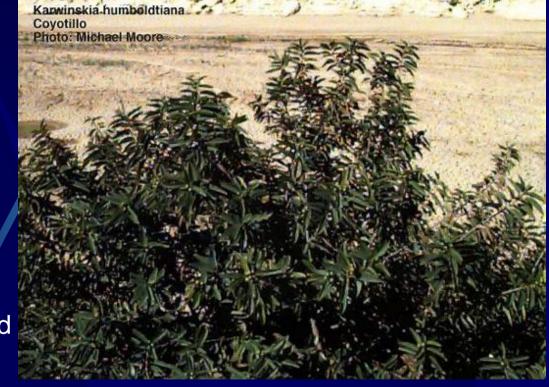
- Woody, erect, lightly branched annual, 2-3 m tall
- Alternate pinnate leaves with 4-5 pairs of leaflets spaced on common stalk
 - Flowers are yellow in loose clusters on leaf axils
- Curved seed pods (20 cm) are thick, dark brown and slightly flattened with with pale longitudinal stripes and brown seeds

- Most poisoning in cattle occurs in Nov. and Dec. after frosts. Calves are more susceptible
- Horses may have liver disease sooner than the myonecrosis.
- Toxin is unknown but speculated to be substituted quinones- some evidence it uncouples oxidative phosphorylation.
- 0.4-12% BW toxic
- Skeletal and cardiac toxicity
- Recovery depends on the severity. Rarely does an animal recover once it has become recumbent.



Karwinskia humboldtiana

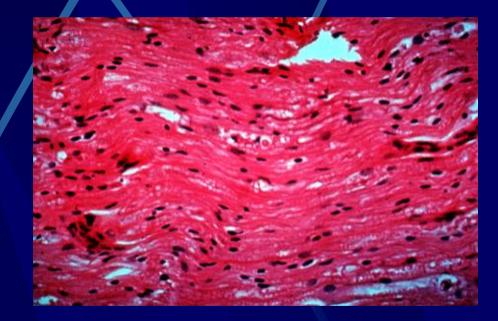
- Coyotillo, buckthorn, tanglefoot, tullidora
- Woody shrub or small tree
- TX, Mexico and SW States
- Gravely hills, canyons, and along arroyos
- Polyneuropathy with acending paralysis
- Anthracenones (T496, T514, T516, T544) usually called tullidinol and possibly other neurotoxins
- Interfere with neuronal synthesis and axonal transport
- Large, long axons most severely affected



Karwinskia humboldtiana

Cattle most sensitive but poisoning reported in goats, sheep, hogs, fowl, horses and man

- Signs
- Lesions: Demyelinating neuropathy, lymphadenopathy, epicardial hemorrhage, skeletal and myocardial degeneration and necrosis, nephrosis and lipidosis
- Axoplasmal dysruption, wallerian degeneration, myelin degeneration



Gossypol

- Gossypium spp. (cotton plants)
- Polyphenolic binaphthalene found in the seed
- Monogastrics and young ruminants most susceptible
 - Lesions
 - CSM for several weeks
 - Inappetence, weight loss, weakness, ascietes, hydrothroax, CHF, skeletal and cardiac muscle degeneration and necrosis, regeneration



Lathyrus spp.

- Europe, Africa, Russia and India
- People eat Lathyrus seeds
- L. hirsutus, L. incanus, L. pusillus, L. sylvestris, L. odoratus used in US
 - Horses may be more susceptible
 - Beta-(gamma-L-glutamyl)aminopropionitrile
 - Metabolized to aminoproprionitrile that is thought to inhibit collagen cross linking (inhibits lysyl oxidase)
- Results in osteolathyrism and angiolathyrism- spinal cord and nerve degeneration, vasuclar aneurysms



 Cattle- stilted gait, weak, shift weight often
 Horses- severe weakness, laryngeal hemiplegia (roaring disease), lameness, sudden death



Vicia villosa

Hairy vetch
 OK and midwest
 Myotoxin plus hepatotoxin, and neurotoxin
 Granulomatous inflammation in heart, skeletal muscle, adrenal glands, kidney, thryroid, brain and lungs (hypersensitivity ?)



Cestrum diurnum

- 1,25-dihydroxycholecalciferol
- Increases Ca absorption from GI, increases Ca mobilization from bone, decreases renal Ca excretion
- Hypercalcemia and hyperphosphatemia- >60 product=soft tissue mineralization
- Cardiac, pulmonary, renal, and gastrointestinal mineralization
- Dystrophic calcification



US Solanum spp.





S. verbascifolium, S. torvum, Nierembergia veitchii, Cestrum diurnum (jessamine, wild jasmine, day cestrum, king of the day, Chinese inkberry) FL

Enzootic Calcification

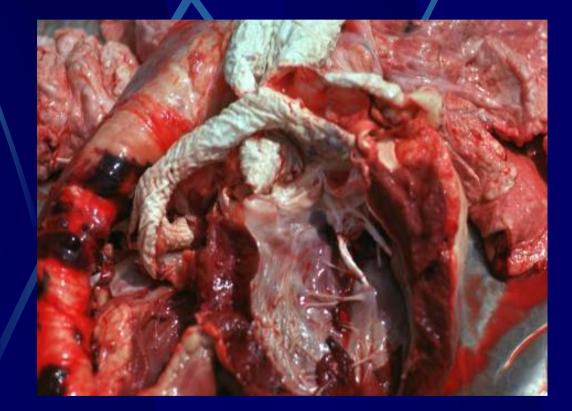
Signs/Lesions:

- Chronic weight loss despite normal appetite
- Stiffness » lameness » recumbency
- Pain in the ligaments and tendons
- Heart murmurs » failure
- Calcification of tendons, ligaments, and elastic arteries » calcinosis of aorta, pulmonary arteries, heart valves, and endocardium



Prognosis

Recovery is rare if poisoning is chronic
 Less severely poisoned animals will probably recover if they are denied further access to the plant and are given a balanced ration.



Cardiac Glycoside Containing Plants

- Digitalis (model compound)
- 100-200 mg/kg lethal
- 8% use results in toxicity
- Blocks Na/K ATPase causing increased intracellular Na and lowering the membrane potential
- Resulting increased Ca causes a positive ionotropic effect
- High doses interfere with the cardiac conduction system especially the SA and AV nodes

Asystole

Lesions (Cardiac Glycosides)

Arrhythmias (tachycardia), cold extremities, dilated pupils, blue mucous membranes, sweating, colic, anorexia, vomiting, diarrhea, bradycardia, heart block, asystole, and death.

Minimal myocardial hemorrhage, myofiber vacuolation with minimal inflammation.

Digitalis purpurea

Foxglove
 Biennial herb from Europe, common on west coast
 Digtoxin, digoxin, gitoxin
 Toxic green or when dry



Nerium oleander

- Ornamental throughout North America
- Evergreen shrub

Nerioside, oleandroside, oleandrin, digitoxigenin, neriin, folinerin, oleandromycin, rosagenin, and odoroside that are similar to digitoxin

- Toxic green and dry
- Most poisonings from clippings



Convallaria majalis and C. montana

- Lilly of the valley
- Ornamental throughout North America
- C. montana native to eastern US
- Convallarin, convallarmarin, convallatoxin (cardiac glycosides)
- All parts, green and dry are toxic
- Signs persist for 3 weeks including dermatitis and gastrointeritis



Apocynum spp.

 Dogbane, Indiana hemp
 Perennial erect plant of North America
 Green and dry plant are toxic

Root used therapeutically



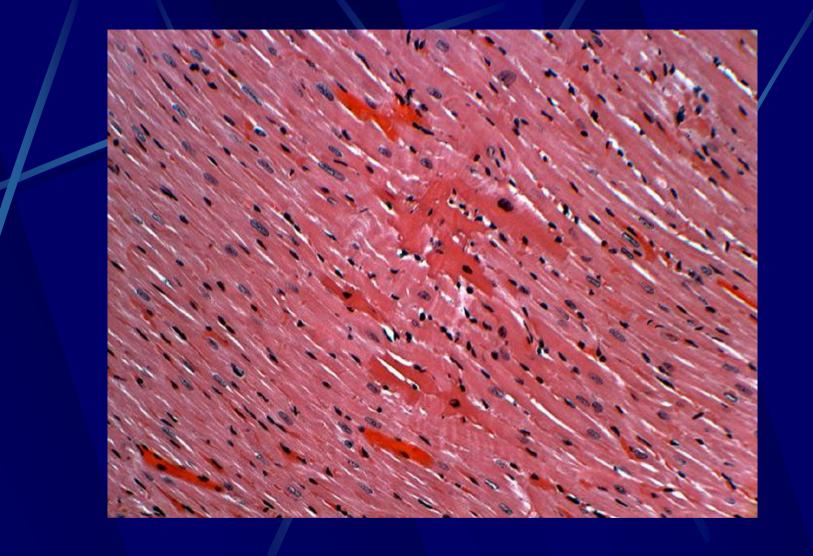
Adonis aestivalis (Pheasant's Eye)

Less toxic or more toxic?









Rhododendron spp.

Rhododendron

- Deciduous shrub found throughout North America
- Andromedotoxin, grayanotoxin (Alters Na channels)
- Cattle, sheep, goats, rarely horses and people have been poisoned
- All parts both green and dry are toxic
- Gastroenteritis, colic, salivation, epiphora, anorexia, depression, nausea, vomiting, defecation, weakness, incoordination, paralysis, absent pupillary reflexes, coma, nephrosis, liver degeneration, aspiration pneumonia



Kalmia spp.

Laurels Evergreen shrub grayanotoxin

Kalmia microphylla Western Mountain Laurel photo by David Parke

Pieris japonica and P. foribunda

Japanese PierisWoody shrubGrayanotoxin



Other potentially myotoxic plants

- Macadamia nuts- transient muscular weakness in dogs
- Hops (Humulus lupulus)- malignant hyperthermia syndrome in dogs
- Ixiolaena brevicompta- Austrialian plant causing tiring syndrome in sheep
- Helichrysum argyrophaerum- South Africa
- Geigeria ornative- South Africa
- Cytisus scoparius- Scotch broom, leguminous shrub



Disease of neglect

Centauria spp.

Centauria repens or Acroptilon repens (Russian knapweed)







- Creeping perennial with black horizontal roots
- Erect, rather stiff, and branched plant up to 1 meter high
- Stems are covered with soft gray hair or nap
- Lower leaves are linear, alternate with toothed margins
- Lavender-white thistle-like flowers have papery spineless bracts
- The grayish seeds are 1-2 mm with bristles at one

Centauria solstitialis (yellowstar thistle, Barnaby's thistle)

- Annual herbaceous weed, branching from the base up to 30 cm tall
- Winged ascending branches with cottony hair covered, basal, lobed leaves
- Yellow disc flowers tipped with characteristic stiff yellow spines 1 to 2 cm) long



- Aspartic and glutamic acids
- Sesquiterpene lactones, solstitialin A 13-acetate and cynaropicrin
- Dopaminergic neurotoxin,
 2,3 dihydro-3, 5 dihydroxy-6methyl-4 (H) pyran-4-1



Weeks to months of exposure
 Green yellow star thistle equal to 86 to 200 percent of their body weight before clinical signs develop



Chewing Disease

- Dysfunction of facial, mouth, and throat muscles (chewing disease)
- Facial paralysis that causes "smiling", tongue lolling, protruding tongue, and head tossing
- Depression, loss of interest in food, dehydration and malnutrition, difficult breathing, incoordination, muscle tremors



Negropallidal encephalomalacia

 Necrosis of the substancia nigra and globus pallidus (negropallidal encephalomalacia)
 As there is no treatment and the disease is irreversible, it is best to avoid exposure.

