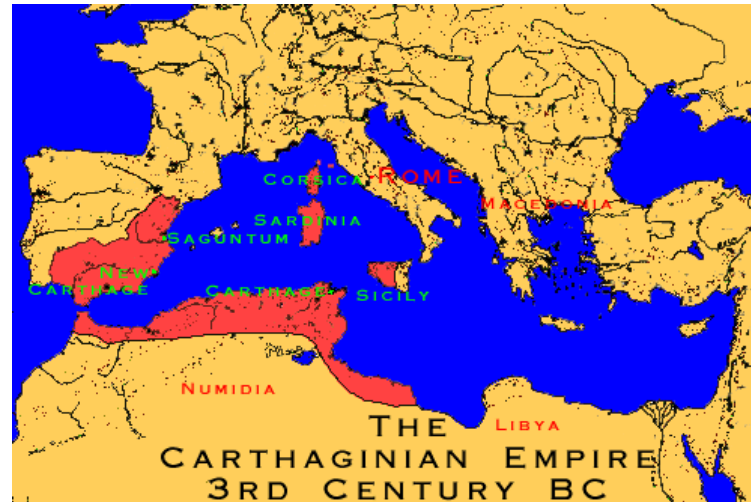


# History of Herbicides and Herbicide Resistance

Scott Steinmaus, PhD  
Biological Sciences

# Carthage

- Romans brought down Carthage in 164 BC using salt to destroy their agriculture
- Sulfuric acid, salts of copper and arsenic, carbon bisulfide
- Very high rates
- Non-selective





# Growth regulator problems

- Dioxin contaminant in making Agent Orange (2,4,5-T)

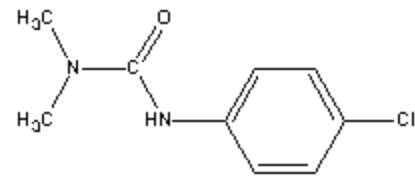


A guerrilla in the Mekong Delta paddles through a mangrove forest defoliated by Agent Orange (1970). Photograph by Le Minh Truong © National Geographic Society

<p><b>2,4-D</b> 2,4-dichlorophenoxyacetic acid</p>	<p><b>MCPA</b> 4-chloro-2-methylphenoxyacetic acid</p>
<p><b>Dichlorprop (2,4,-DP)</b> 2-(2,4-dichlorophenoxy) propionic acid</p>	<p><b>Mecoprop (MCP)</b> α-(4-chloro-2-methylphenoxy) propionic acid</p>
<p><b>Fenoprop (Silvex, 2,4,5-TP)</b> 2-(2,4,5-trichlorophenoxy) propionic acid</p>	<p><b>2,4,5-T</b> 2,4,5-trichlorophenoxyacetic acid</p>
<p><b>2,4-DB</b> (2,4-dichlorophenoxy) butyric acid</p>	
<p>Structures of 2,4-dichlorophenoxyacetic acid (2,4-D) and chemically-related herbicides.</p>	

# Phenyl substituted ureas

- Discovered in 1951
- Examples are Monuron, diuron
- Photosynthetic inhibitor (the most common mode of action of all herbicides)



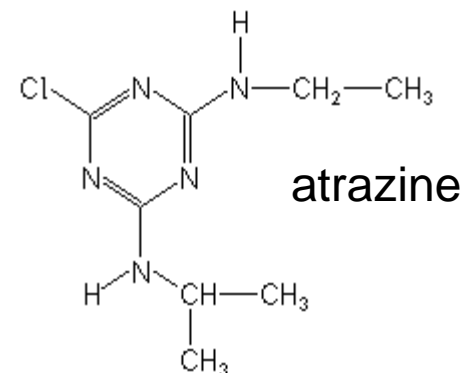
Monuron



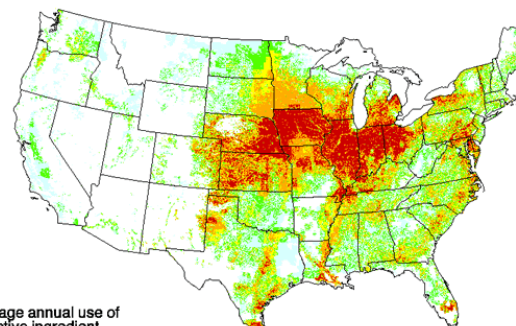
Symptoms of PSII e-transport inhibitors on bean

# Triazines

- Developed in 1955
- Atrazine, simazine, Velpar
- Photosynthetic inhibitor
- Long residual activity



ATRAZINE - herbicide  
1997 estimated annual agricultural use



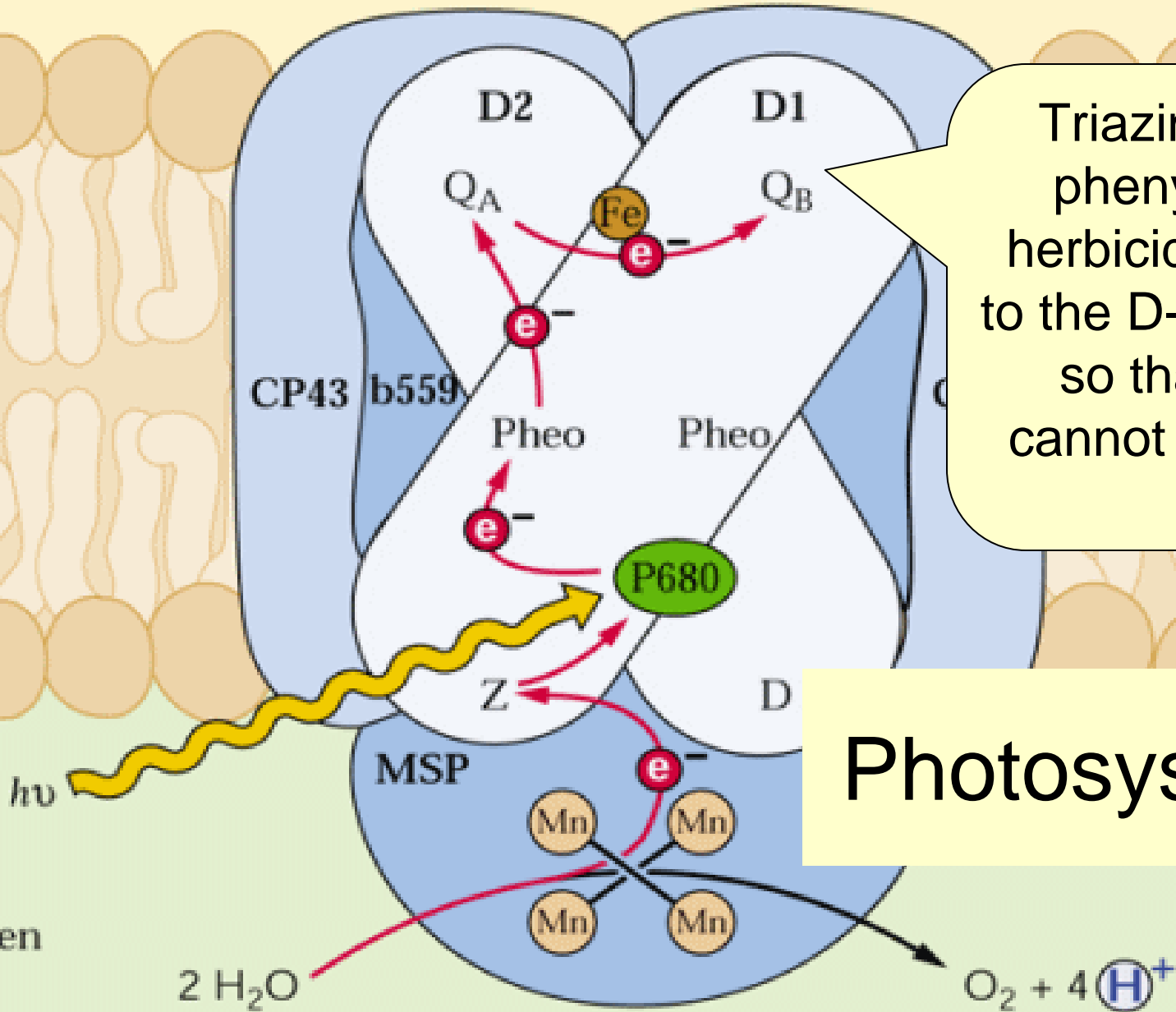
Average annual use of active ingredient (pounds per square mile of agricultural land in county)

- no estimated use
- 0.001 to 0.358
- 0.359 to 2.151
- 2.152 to 9.855
- 9.856 to 32.77
- >= 32.771

Crops	Total pounds applied	Percent national use
corn	62,381,038	84.00
sorghum	5,750,038	3.09
summer fallow	2,539,189	3.42
sugarcane	2,203,421	2.97
sweet corn	340,452	0.46
sod harvested	30,214	0.04
other hay	13,224	0.02
seed crops	5,833	0.01

Atrazine use (kg/km<sup>2</sup>) is high in corn growing regions of midwestern US (USGS 1991).

Stroma

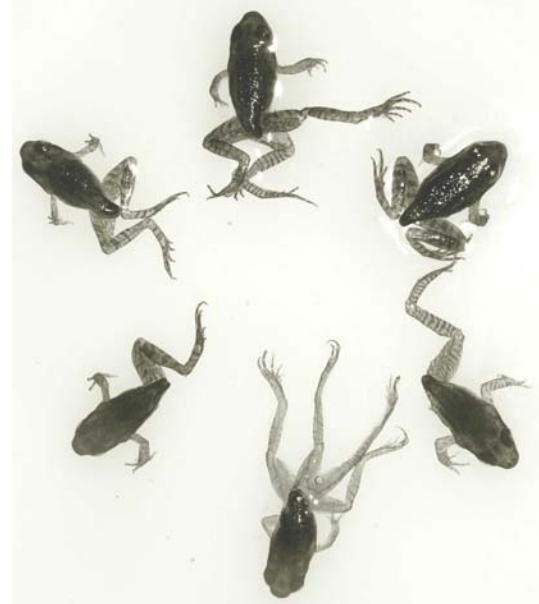
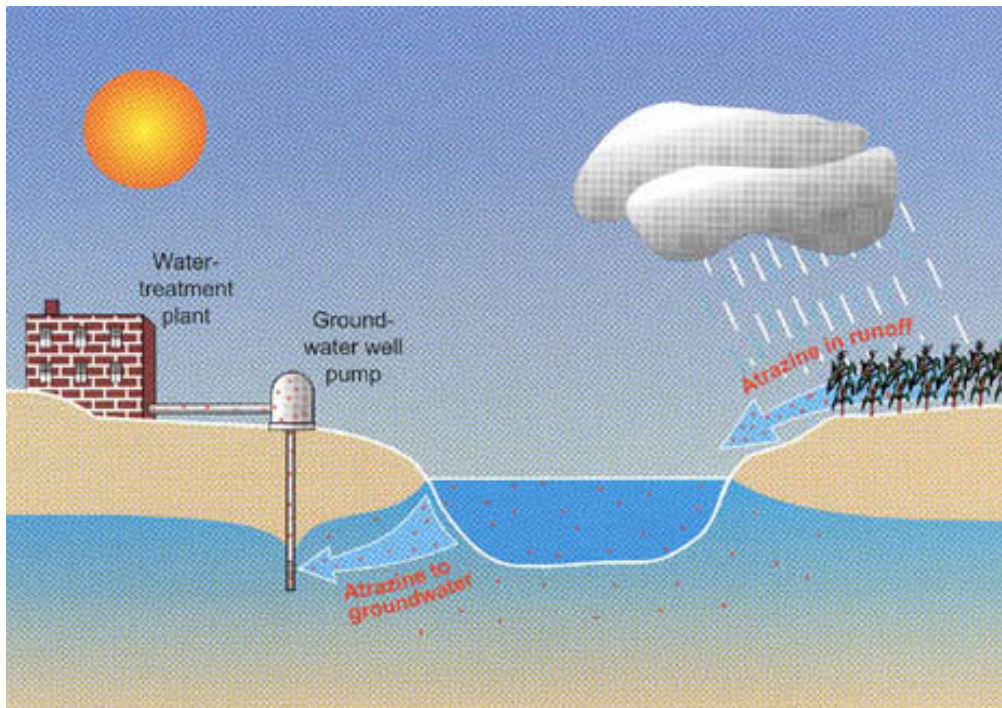


Triazine and phenylurea herbicides bind to the D-1 protein so that  $Q_B$  cannot carry  $e^-$

Photosystem II

# Problems with the triazines

- Aquifer contamination
- Hormone mimics/  
endocrine disruptors





# ACCase Inhibitors: the 'dims' and 'fops'

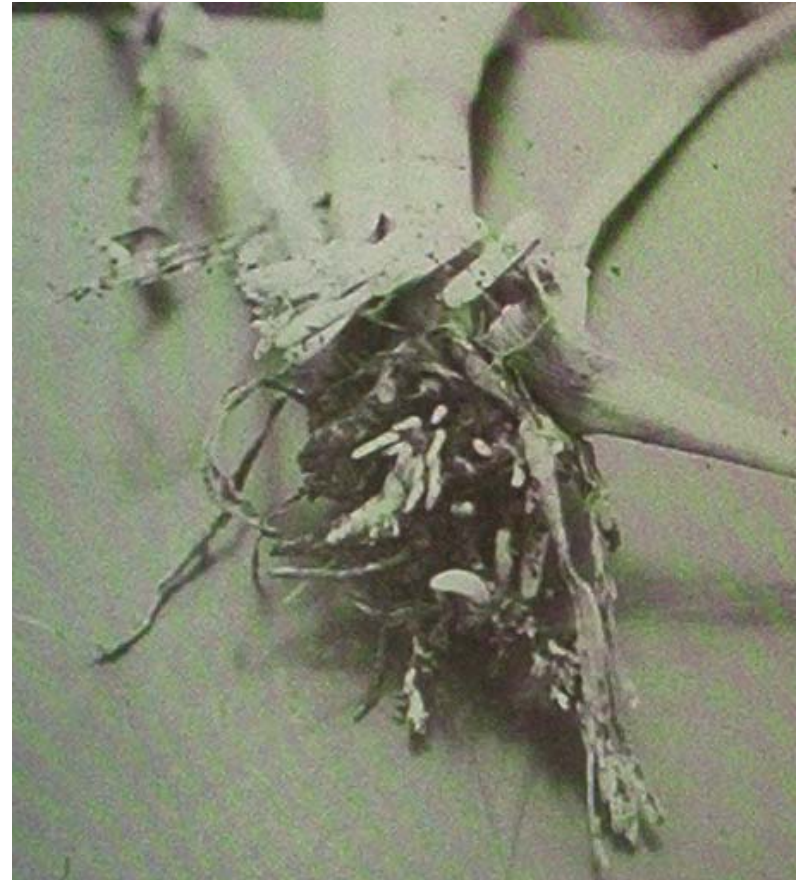
- Grass-specific ACCase inhibitors:  
sethoxydim=  
Poast  
fluazifop=  
Fuslade
- Inhibits acetyl-CoA to fatty acids



ACCase inhibitor on rice

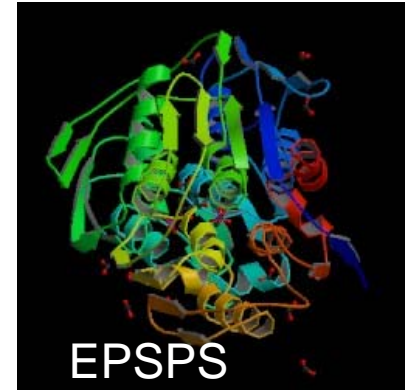
# Dinitroanilines

- Introduced in the 1960's
- Dinitroanilines (trifluralin (Treflan®), oryzalin (Surflan®)) are carbamate herbicides
- Binds to tubulin during the formation of microtubules stops cell division.



# Glyphosate

- Developed in the late 1970's
- Marketed first as Roundup
- Inhibits EPSPS
- Non-selective until...



velvetleaf



Roundup Ready  
soybean

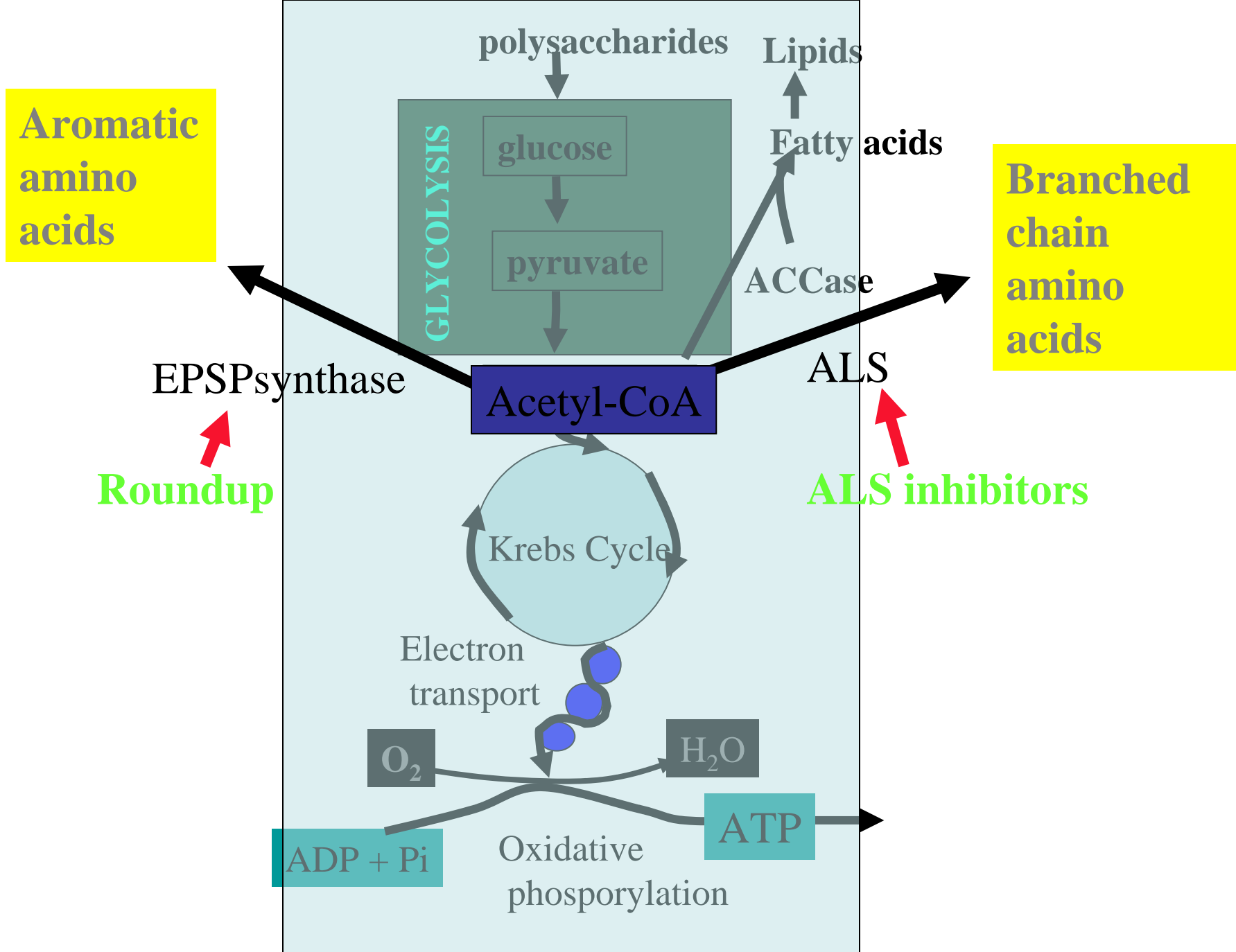
conventional  
soybean

# ALS Inhibitors

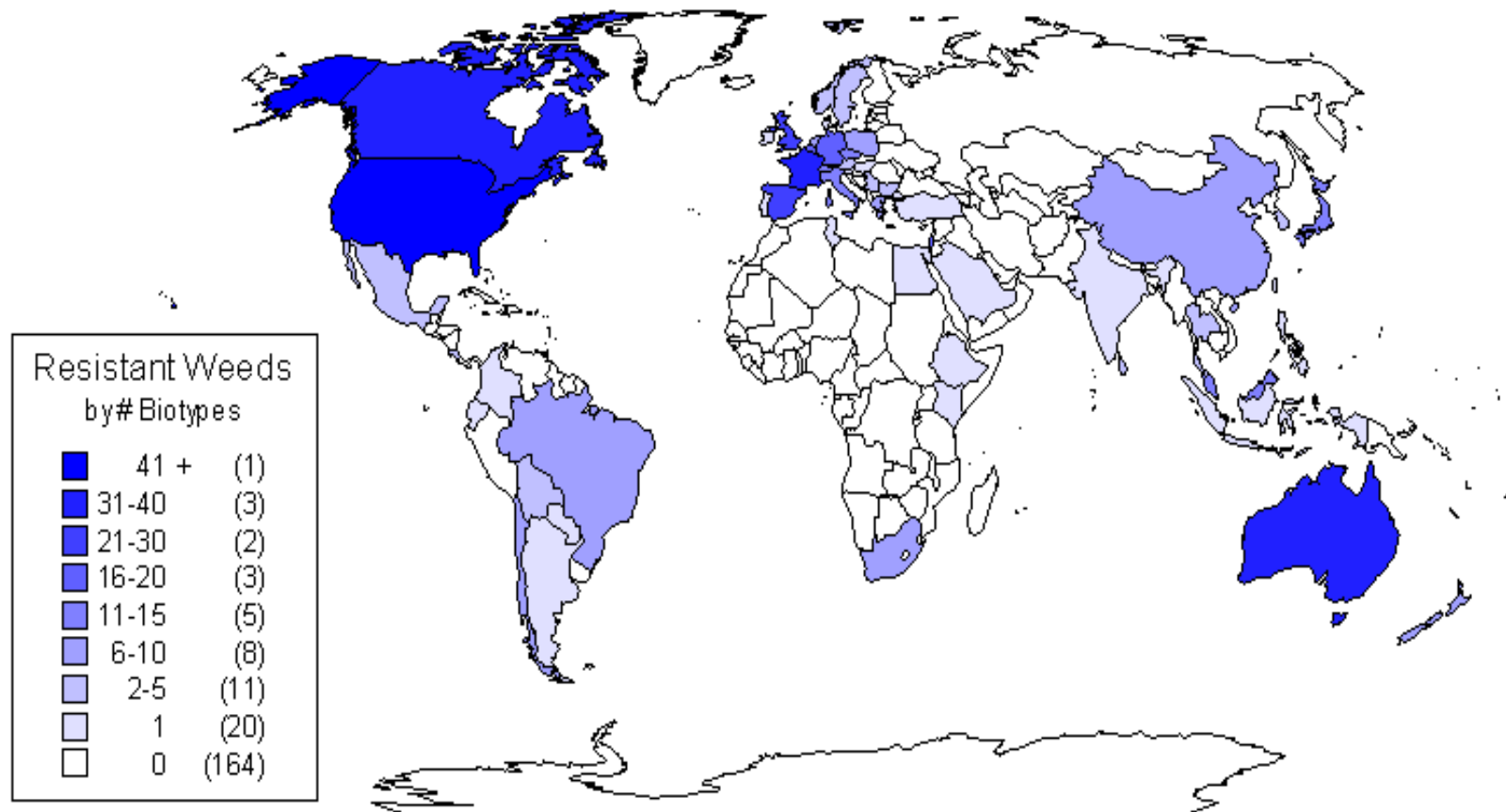
- Developed in the 1980's
- Imidazolinones:  
imazethapyr (Pursuit)
- Sulfonylureas: 'urons'  
bensulfuron (Londax),  
chlorsulfuron (Glean, Telar)
- pyrimidinyloxybenzoate,  
triazolopyrimidines
- Very low rates (g/A)



ALS inhibitor symptoms begins at growing points where branched chain amino acids are needed



# Distribution of Herbicide Resistant Biotypes



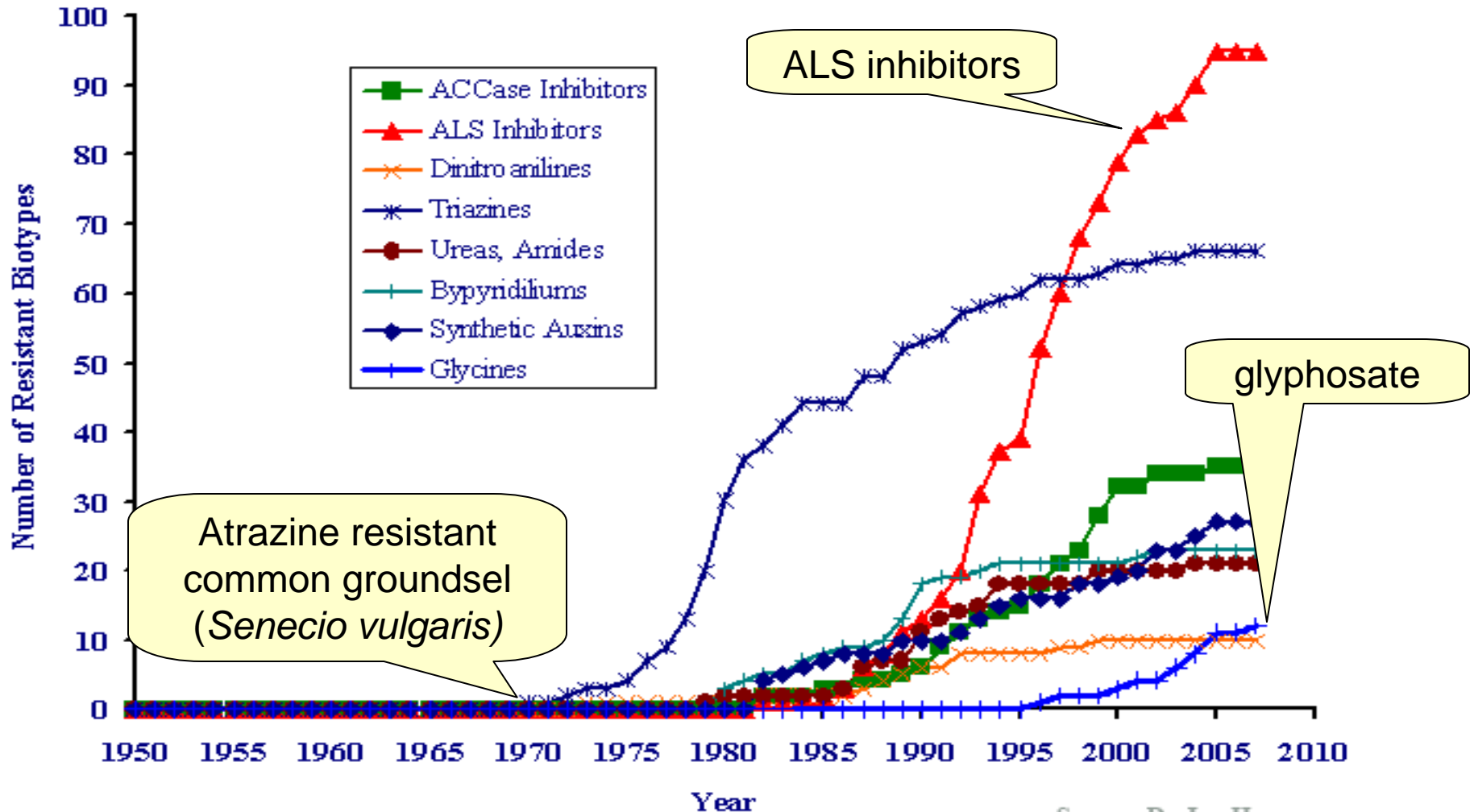
Source: Dr. Ian Heap  
[www.weedscience.com](http://www.weedscience.com)

# Most Important Herbicide-Resistant Species

1. Rigid Ryegrass *Lolium rigidum*
2. Wild Oat *Avena fatua*
3. Redroot Pigweed *Amaranthus retroflexus*
4. Common Lambsquarters *Chenopodium album*
5. Green Foxtail *Setaria viridis*
6. Barnyardgrass *Echinochloa crus-galli*
7. Goosegrass *Eleusine indica*
8. Kochia *Kochia scoparia*
9. Horseweed *Conyza canadensis*
10. Smooth Pigweed *Amaranthus hybridus*



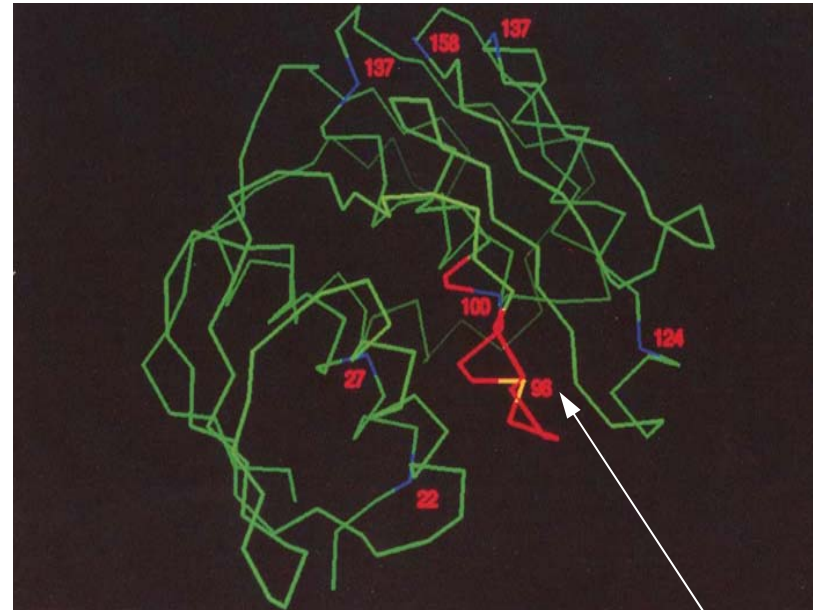
# Development of herbicide resistance



Source: Dr. Ian Heap  
<http://WeedScience.com>

# Glyphosate resistance

- 2 possible mutations in EPSPS can confer resistance
- First reported in 1996 in rigid ryegrass (*Lolium rigidum*) in Australia
- Next in California 1998 in Italian ryegrass (*Lolium multiflorum*)



Substituting Ala for Gly at the residue site 96 Confers glyphosate resistance. The other numbers correspond to active site residues for glyphosate and substrate binding.

# Roundup (6qt./A) at highest labeled rate



ryegrass  
(resistant)

Roundup  
Ready corn

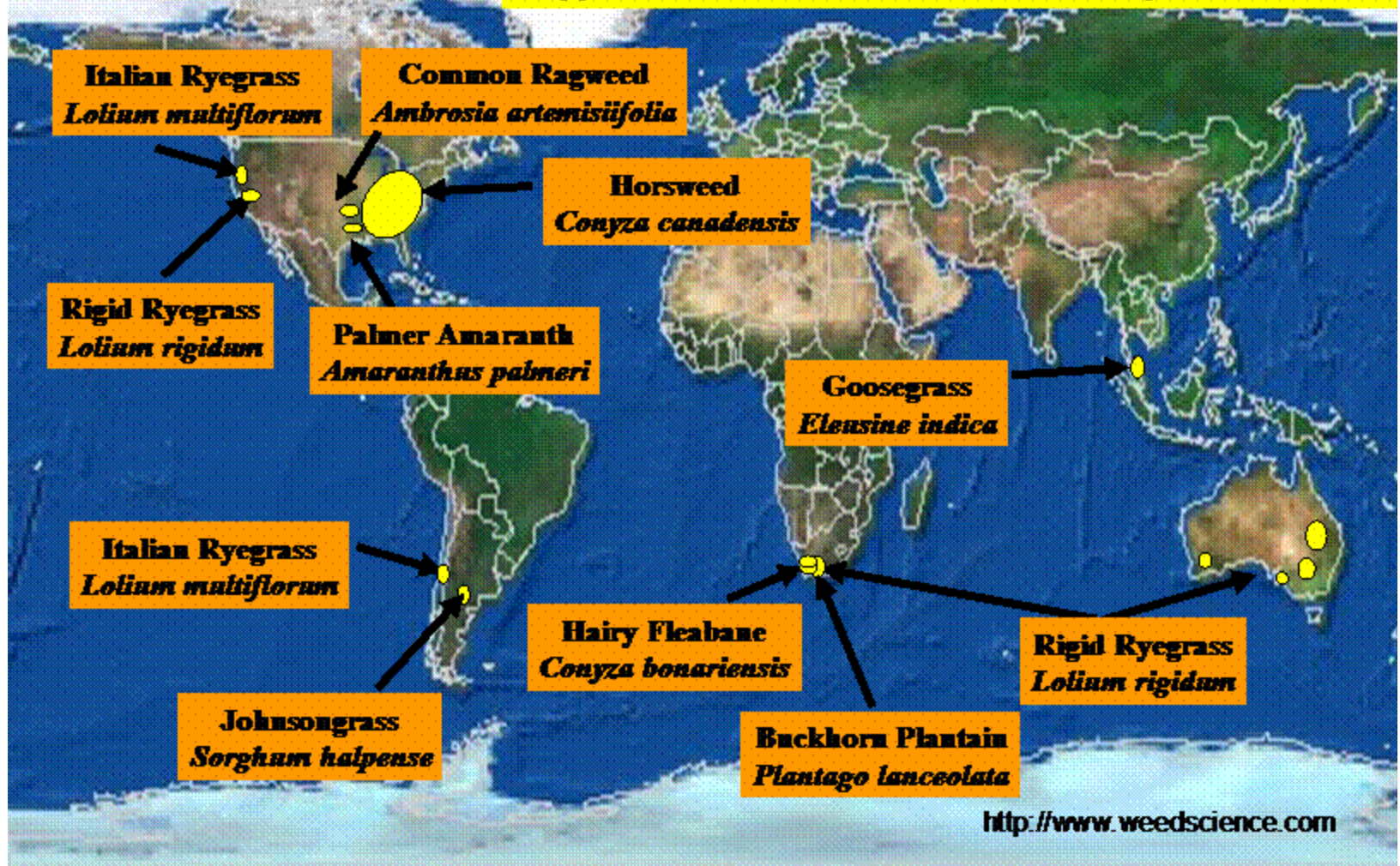
Roundup  
Ready cotton

cotton

ryegrass  
corn

# Glyphosate resistance

Glyphosate Resistant Weeds – September 2006



# ALS Inhibitor resistance

- 10+ possible mutations on ALS that can confer resistance to a plant
- Highest number of cases of resistance (95 biotypes)
- Latest mode of action



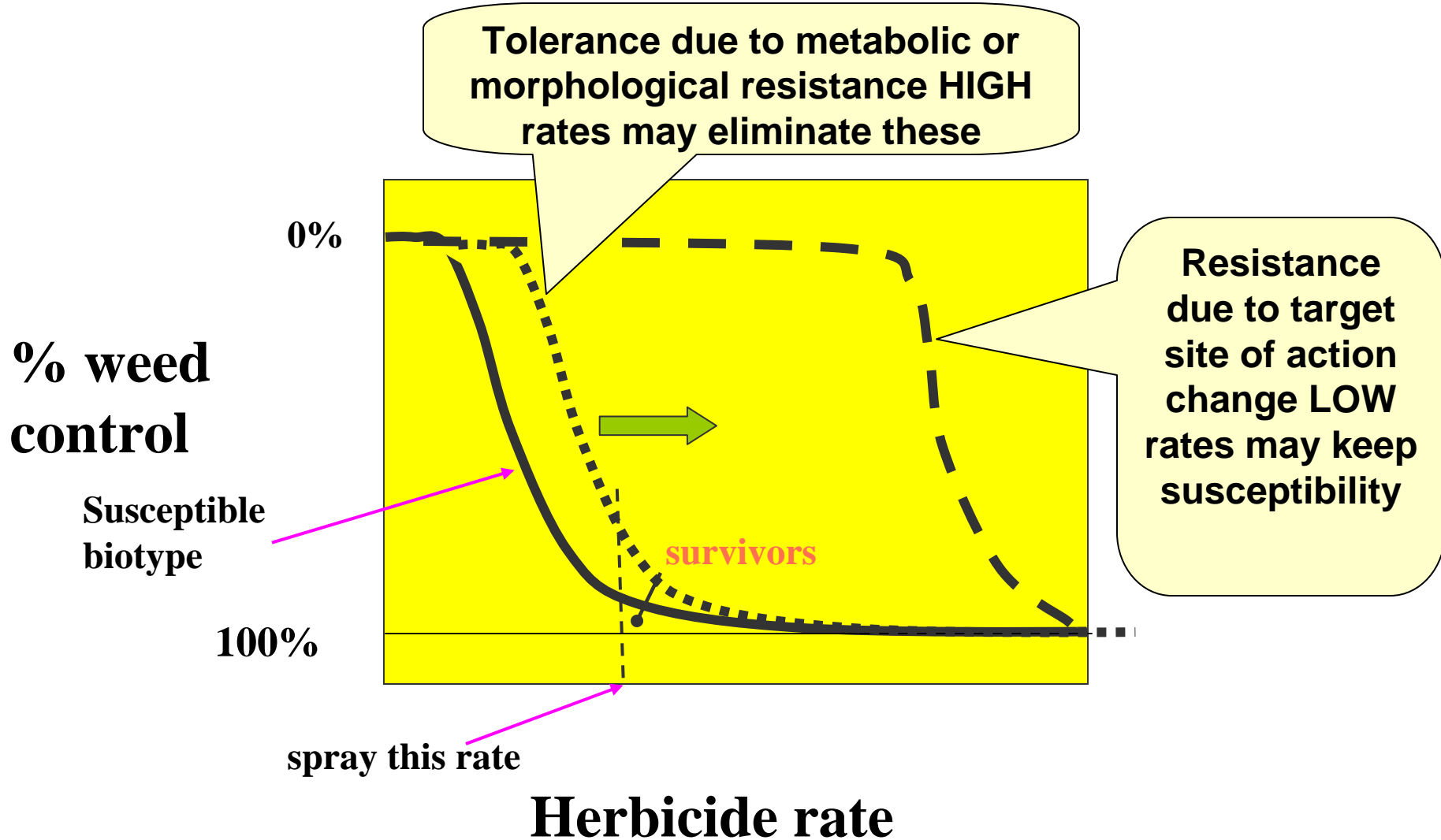
# Rate of evolution depends on:

- Generation time
- Initial frequency of resistant allele
- Threshold at which resistance is recognized
- Strength of selection: intensity and duration

# Herbicide characteristics that influence weed resistance

- herbicides with a single target site
- selection intensity: rate and duration
- herbicides used multiple times during the growing season
- herbicides with long residual activity
- herbicides used for consecutive growing seasons
- Herbicides used alone without other control strategies

# How weeds are resistance determines how to manage them





# Summary

- Herbicides began as non-selective, highly toxic, used at high rates
- They became selective, non-toxic, and used at low rates
- Resistance developed primarily because of reliance on single modes and methods of action
- Resistance is "micro"-evolution: a function of selection pressure