

Metabolic profiling unravels complex biological phenomena in the model plant *A. thaliana*

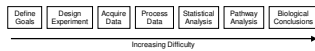
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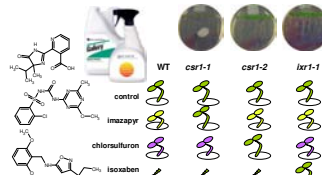
Environmental Stress: Hypoxia



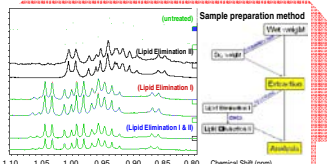
Metabolic Profiling



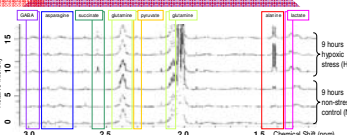
Chemical Stress: Herbicides



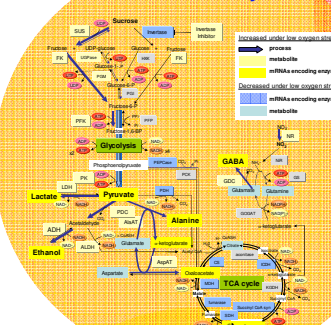
Chemistry



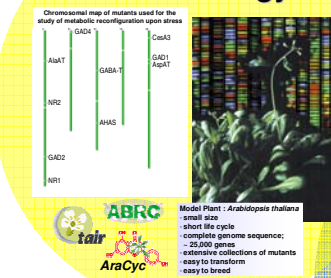
Multivariate Statistics



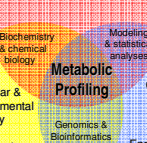
Biochemistry



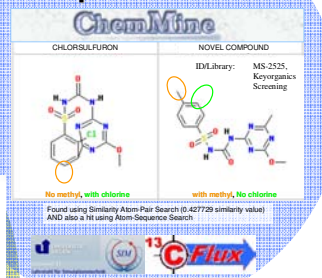
Plant Biology



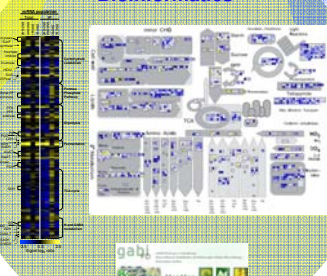
Chemistry



Computer Science



Bioinformatics



References

MapMan: Thimm, et al. *Plant J.* (2004) 37, 914-939.
 TAIR: Garcia-Hernandez, et al. *Funct. Integr. Genomics* (2002) 2, 239-253.
 ChemMine: Girke, et al. *Plant Physiol.* (2005) 138, 573-577.
 Hypoxic Stress: Branco-Price, et al. *Plant J.* (2008) submitted.
 Metabolic Profiling: Kaiser, et al. *NMR Spectroscopy in Pharmaceutical Analysis*, Ed. Holtzgrabe, et al. (2008) in press.

Acknowledgements

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An example of INNOVATIVE Interdisciplinary Science

Chemical Genomics:

Use of chemical compounds (rather than mutations) to study fundamental biology of cells; through the chemical's effects on genes and gene products

Chemical Libraries

ChemBridge DiverSet: 20,000 compounds
 ChemBridge Microformat: 10,000 compounds
 ChemBridge NovaCore: 10,000 compounds
 Microsource Spectrum collection: 2,000 compounds
 Sigma/TimTec MyriaScreen: 10,000 compounds

Value ~ \$100,000

Biomek FX[®] Fluid Handling Robot

*Sterile hood for contamination prevention
 *96 tip pool with gripper for moving lab ware
 *8-pin tip rack with individual tip control
 *50ml and 200ml pin tool for one-step dilution
 *High temperature deck position for agar
 *No incubating to wash for agar cleanup
 *Cryostat hole for high capacity storage

Chemical Genomics Resources, Facilities and Support Staff

Microscopy and Imaging, Dr. David Carter
 5 confocal microscopes, other imaging instruments

Bioinformatics, Dr. Thomas Girke
 64-CPU Linux cluster, 4 quad core Mac Pro, 8 large servers,
 1 Silicon Graphics (SGI)

Proteomics, Dr. Songqin Pan
 3 mass spectrometers

Genomics Facility, Dr. Glenn Hicks
 3 robots, DNA sequencer, 2 arrayer machines, 5 scanners,
 Affymetrix hand and software, 2 qPCR cyclers, FACS cell sorter

Analytical Facility, Dr. Cynthia Larive
 5 mass spectrometers, 2 x-ray diffraction systems,
 5 magnets for NMR, 1 EPR, 2 FT-IR, 2 Raman spectrometers,
 1 fluorescence spectrometer, 1 polarimeter

Example: Chemical Genomics "Screen"

Screen #238
 Surpin & Ralston

10,000 chemical compounds
 218 chemical compounds
 149 auxin-like compounds
 69 unique chemical compounds
 34 compounds
 4 compounds
 1 compound

Surpin, et al. "The power of chemical genomics..." *J. N.A.S.* (2005) 102, 4902-07.
 Branco-Price, et al. "PCP is a Target of Gracillin" *Chem. Biol.* (2007) 14, 1366-76.

ChemGen IGERT Projects

IGERT Student	GP	Project Description
SAMER ELKASHAF	GGB	Screen for inhibitors of plant proteins that respond to viral infection
CHARLES JANG	GGB	Screen for inhibitors of MAPK3 (cellular signal transducer)
JAMES KIM	CMB	Screen for compounds that affect G-protein coupled receptors
COLLEEN KNOTH	PB	Screen for elicitors of pathogen defense responses within a plant
CHRIS MERRYNOW	C	Screen for disruptors of protein trafficking
EDDIE CAO	CS	Virtual screen using compound similarity searching algorithms
JOLINE DIEDRICH	C	Screen for compounds that alter protein phosphorylation status
THERESA DINH	PB	Screen compounds that disrupt transcriptional gene silencing
KAYLA KAISER	C	Screen for inhibitors of tobacco pollen tube germination
AUGUSTA JAMAN	GGB	Screen for inhibitors of ROP-RacGAP interaction (cell signaling)
SEAN BOYLE	GGB	Virtual screen of lead compound DAG B1 analog library
MICHELLE BROWN	GGB	Screen for inhibitors of tobacco pollen tube germination
ANNA CHARISHI	GGB	Virtual screen based on physicochemical properties of compounds
ANDREW DEFRIES	PB	Screen for modulators of protein-protein interactions
MELINDA SALLUS	PB	Screen for elicitors of JA & ethylene dependent defense pathways
MELISSA SMITH	PB	Screen for Lipo-A inhibitor & elicitors (JA dependent defense)

GGB = Genetics, Genomics and Bioinformatics CMB = Cell, Molecular & Developmental Biology PB = Plant Biology C = Chemistry CS = Computer Science GP = Graduate Program