

Biocuration:

Helping Researchers Harness the Data Explosion at TAIR and the Plant Metabolic Network



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Overview

- ❑ Biological data **explosion**
- ❑ Biocurators want to help!
- ❑ Biocuration practices and resources at two plant databases
 - The Arabidopsis Information Resource
 - The Plant Metabolic Network
- ❑ Request for *your* help!

Growth of biological data

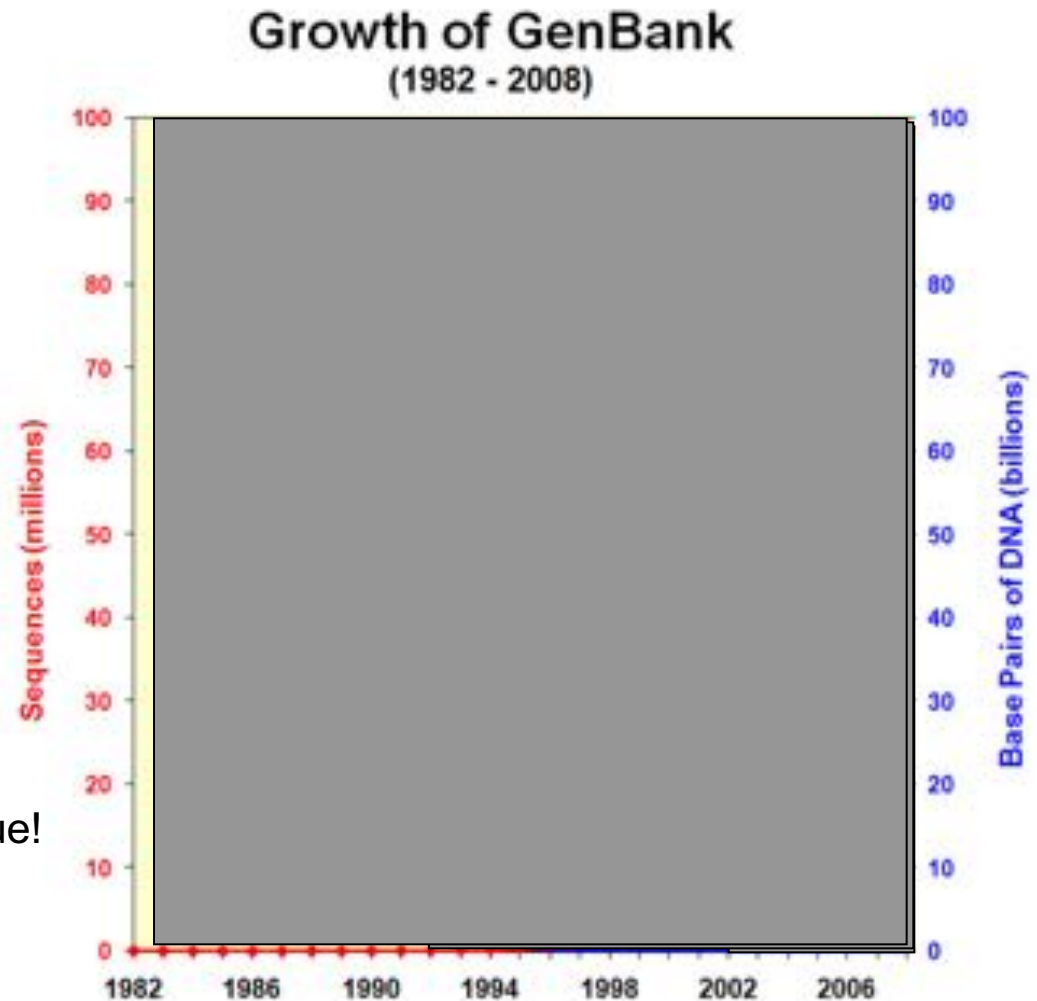
- Over time biological data increases in
 - Quantity
 - Methods improve
 - Costs decrease

Growth of biological data

□ Nucleotide sequences

- Number of sequences in 1982
 - 606
- Number of sequences in 1992:
 - 78,608
- Number of sequences in 2002:
 - 22,318,883
- Number of sequences in 2008:
 - 98,868,465
- And, the acceleration may continue!

Source: National Center for Biotechnology Information (NCBI)



Growth of biological data

- Over time biological data increases in

- Complexity

- Protein data

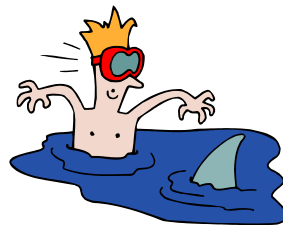
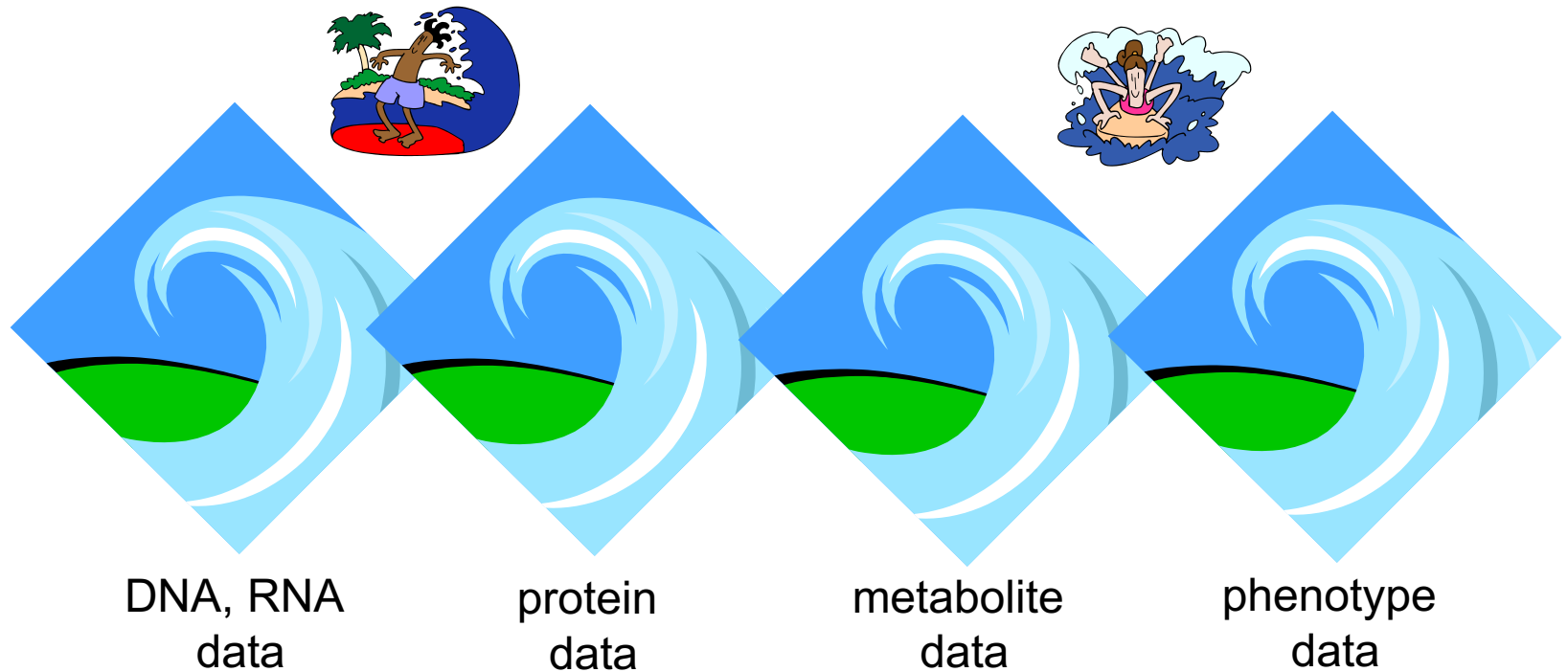
- Primary sequence
 - 3D structure
 - Subcellular localization
 - Rate of degradation
 - Enzymatic activity properties
 - Post-translational modification
 - Phosphorylation
 - Prenylation
 - Methylation
 - Ubiquitination

Is it static or dynamic?

What stimuli cause it to change?

By how much?

The waves of data keep mounting!





Exploring the sea of biological data

- Primary data source
 - Articles published in peer-reviewed journals
 - Over 18 million available through PubMed by 2008!
 - NOT a comprehensive set; many journals are missing

- Answering scientific questions
 - Specific focus:
 - Find **every single piece of information ever discovered** about my favorite gene – XYZ1 – to figure out exactly what it does
 - Broad search:
 - Compare the protein sequence of **every single transcription factor ever discovered** in a prokaryote or a eukaryote to study the evolution of nuclear-localization signals
 - ***How do you collect these data from ALL of the relevant research articles?***
 - Data repositories . . . staffed by **biocurators** . . . try to help!
 - Computer scientists and bioinformaticians contribute to these efforts as well!

Global data repositories / databases

- Centralized data hubs
 - Many data types
 - Many species

- Asia
 - Several in Japan, e.g. RIKEN, China is adding new ones
- Europe
 - European Bioinformatics Institute (EBI)
- USA
 - National Center for Biotechnology Information (NCBI)

Global data re

NCBI Resources How To

NCBI
National Center for
Biotechnology Information

Search

Resources

- NCBI Home
- All Resources (A-Z)
- Chemicals & Bioassays
- Data & Software
- DNA & RNA
- Domains & Structures
- Genes & Expression
- Genetics & Medicine
- Genomes & Maps
- Homology
- Literature
- Proteins
- Sequence Analysis
- Taxonomy
- Training & Tutorials
- Variation

Resources How To

3 A B C D E F G H I J L M N O P R S T U V

Featured items are in bold.

3 3D Domains Database

A Amino Acid Explorer
ASN.1 Format Summary
Assembly Archive

B **BankIt**
Barcode Submission
Batch Entrez
BioAssay Services
BioSystems
BLAST (Basic Local Alignment Search Tool)
BLAST (Stand-alone)
BLAST Link (BLink)
BLAST Microbial Genomes
BLAST Tutorials and Guides
Bookshelf

C Cancer Chromosomes
CDTree
CDTree Installation Page
Clone Registry
Cn3D
Cn3D Installation Page
COBALT
Coffee Break
Concise Microbial Protein BLAST
Consensus CDS (CCDS)
Conserved Domain Architecture Retrieval Tool (CDART)
Conserved Domain Database (CDD)
Conserved Domain Search Service (CD Search)

Specialized data repositories / databases

- ❑ Model organism databases (MODs)
 - Mouse Genome Informatics (MGI)
 - Flybase (*Drosophila*)
 - Saccharomyces Genome Database (SGD) (yeast)
 - **The Arabidopsis Information Resource (TAIR)**
- ❑ Topical databases
 - Worldwide Protein Data Bank (3D structures)
 - miRbase (microRNAs)
 - **Plant Metabolic Network (PMN) (metabolic / biochemical pathways)**

Roles of biocurators at data repositories

- Organize and process raw data
 - Assign unique stable identifiers for nucleotide sequences submitted by researchers
- Review and improve data to generate **curated** data sets
 - Manually correct errors in raw nucleotide sequences to make RefSeq gene structures
- Develop tools for accessing data
 - Provide a protein interaction viewer
- Train users
 - Present at conferences and universities
- ***Try to help researchers harness the data explosion!***
 - ***TAIR***
 - ***Plant Metabolic Network***

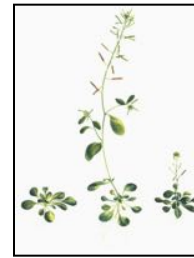
Introduction to TAIR

- TAIR = **T**he **A**rabidopsis **I**nformation **R**esource
- Why Arabidopsis?
- What does TAIR do?
- What can you do with TAIR?



Arabidopsis

Introduction to Arabidopsis

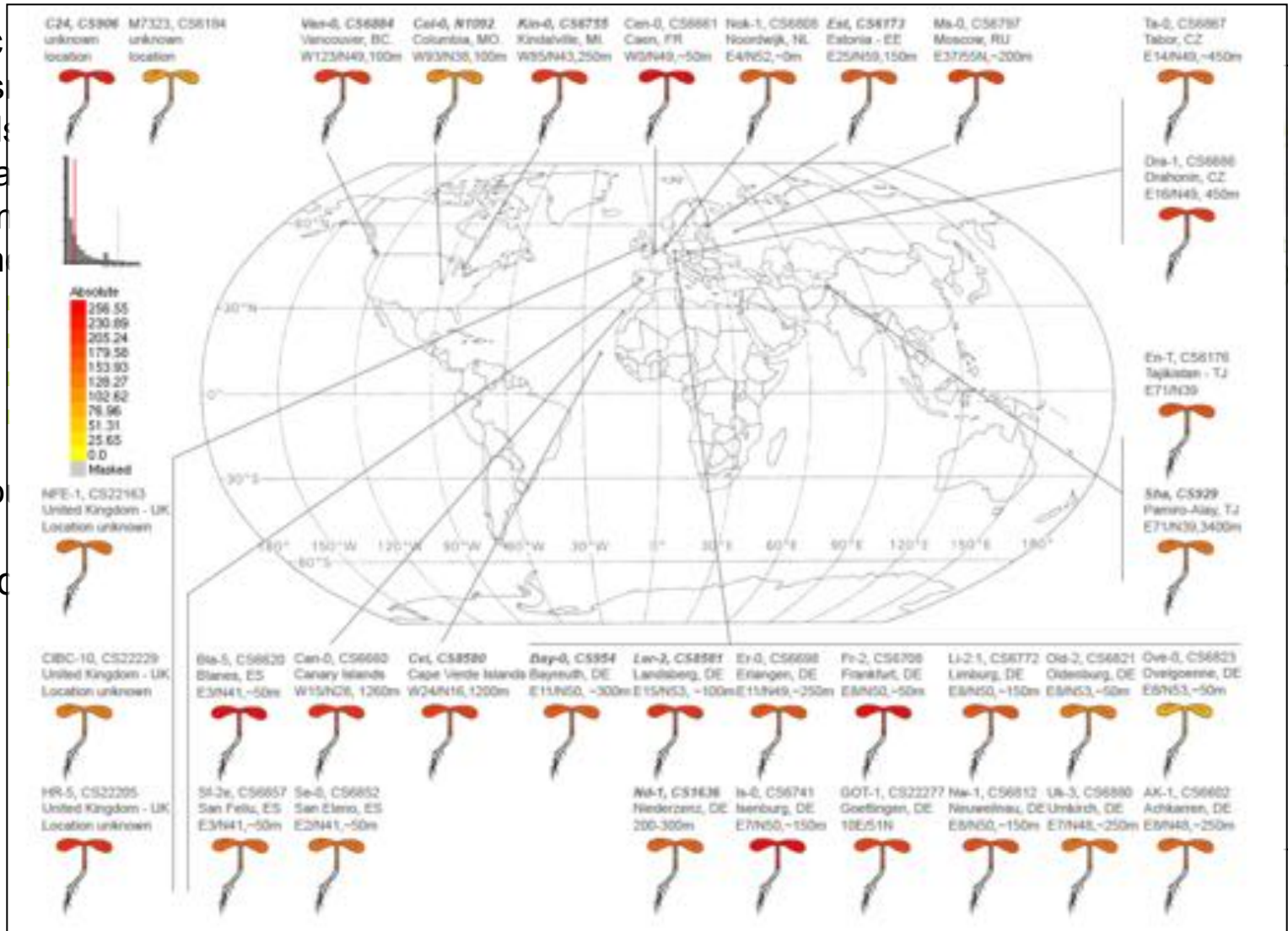


Basic

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Why d



rella

Arabidopsis offers some advantages

- “Good” genome
 - very small: 125 Mb - ~27,000 genes
 - diploid
 - 5 haploid chromosomes
 - fewer/smaller regions of repetitive DNA than many plants
- Quite easily transformable with *Agrobacterium*
 - NO tissue culture required
- Inertia!
 - A group of scientists lobbied for Arabidopsis
 - The genome was sequenced (2000)
 - **MANY resources have been developed**



Arabidopsis research can be applied to “real plants”

- Over-expression of the *hardy* gene from Arabidopsis can improve water use efficiency in rice (*Karaba 2007*)
- A high throughput screen performed using castor bean cDNAs expressed in Arabidopsis found three cDNAs that increase hydroxy fatty acid levels in seeds (*Lu 2006*)
- These experiments and many more benefit from the work of curators trying to help harness the Arabidopsis data explosion . . .
 - ~2400 articles discussing Arabidopsis in PubMed per year!

What

Curator

TAIR

TAIR

TAIR

Full

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The Arabidopsis Information Resource

The Arabidopsis Information Resource (TAIR) maintains a database of genetic and molecular biology data for the model higher plant *Arabidopsis thaliana*. Data available from TAIR includes the complete genome sequence along with gene structure, gene product information, metabolites, gene expression, DNA and seed stocks, genome maps, genetic and physical markers, publications, and information about the Arabidopsis research community. Gene product function data is updated every two weeks from the latest published research literature and community data submissions. Gene structures are updated 1-2 times per year using computational and manual methods as well as community submissions of new and updated genes. TAIR also provides extensive linkouts from our data pages to other Arabidopsis resources.

The Arabidopsis Biological Resource Center at The Ohio State University collects, reproduces, preserves and distributes seed and DNA resources of *Arabidopsis thaliana* and related species. Stock information and ordering for the ABRC are fully integrated into TAIR.

 TAIR is located at the Carnegie Institution for Science Department of Plant Biology and funded by the National Science Foundation. 

Syntenic Viewer at TAIR



A syntenic viewer (gbrowse_syn, a GMCO project) is now available at TAIR. This tool allows the user to compare syntenic regions between *A. thaliana* and *A. lyrata*. Additional plant genomes will be added in the future.

Breaking News

Syntenic Viewer At TAIR [July 7, 2009]
A syntenic viewer, comparing syntenic regions between *A. thaliana* and *A. lyrata*, is now available at TAIR. More genomes will be added soon.

TAIR9 Genome Release [June 15, 2009]
The TAIR9 genome release is now available at TAIR and NCBI, with 262 new loci, updates to 1254 gene structures and 739 new splice variants.

Tips for searching DNA stocks including vectors and amfDNA clones [May 20, 2009]
Are you searching for clones and vectors available from ABRC? Here are some tips for finding them using the TAIR searches and ABRC catalog pages. Specific instructions for finding amfDNA clones and multifunctional vectors are included.

New clones/constructs at ABRC [March 25, 2009]
Ascomycin clones from W. Frommer, J. Schroeder and S. Asmar, auxin related constructs from A. Theologis, and expression clones from S. P. Dinesh Kumar.

ArAtCyc 5.0 and PlantCyc 2.0 release [March 18, 2009]
97 new and updated.

at directors

Structural curation at TAIR

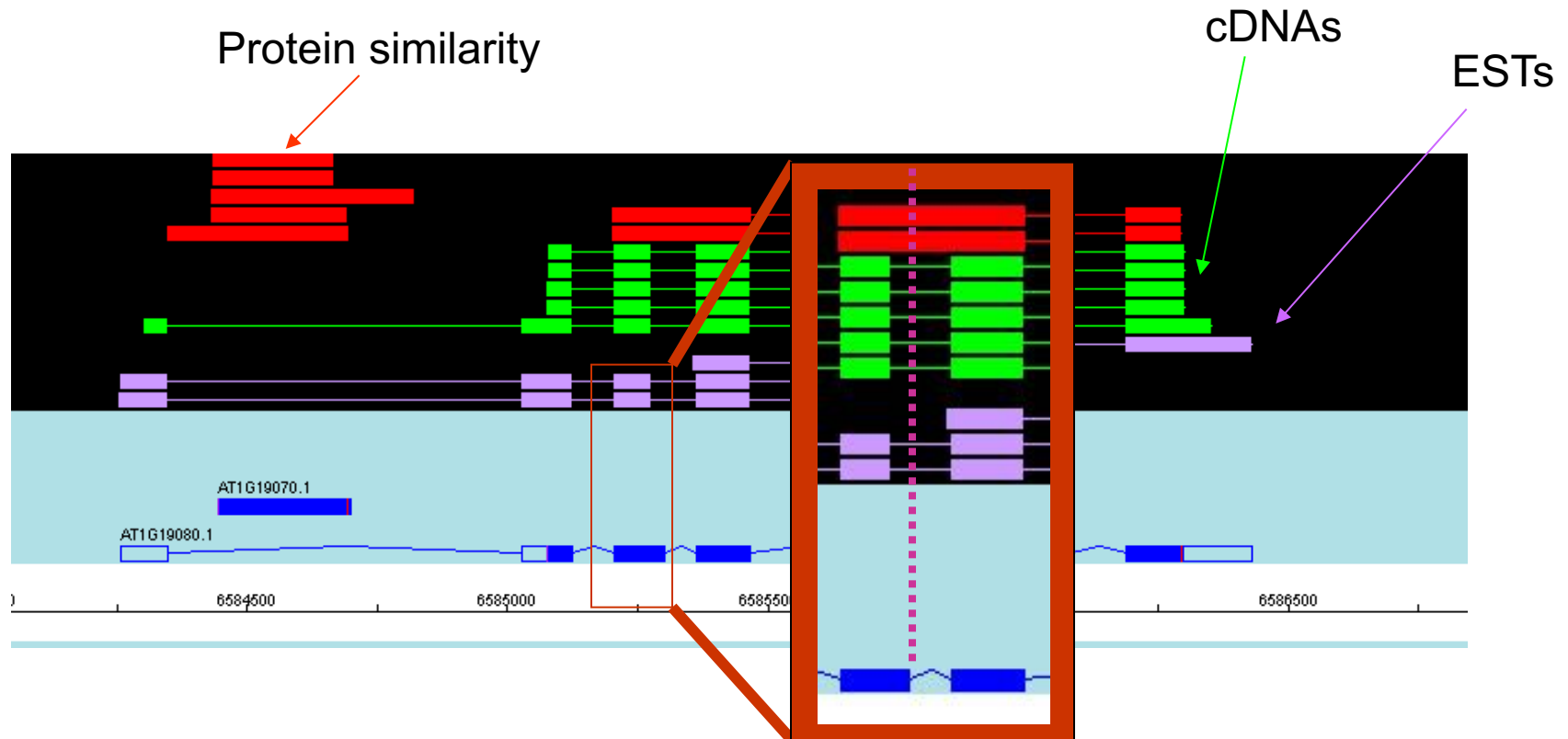
- Structural curators try to answer the question:
What are ALL of the genes in Arabidopsis?
 - Use many types of data
 - ESTs
 - full-length cDNAs
 - peptides
 - orthology
 - RNASeq data**
 - Determine gene coordinates and features
 - Establish intron, exon, and UTR boundaries
 - Add alternative splice variants
 - Classify genes
 - protein coding
 - miRNA
 - pseudogene

Structural curation at TAIR

- Even though the genome was sequenced in 2000 . . .
- . . . the work goes on!
 - TAIR9 – released June 2009
 - 282 new loci and 739 new splice variants
 - TAIR10 – on its way
 - 126 novel genes
 - 1182 updated genes
 - 5885 new splice variants added (18% of all loci)

Structural curation at TAIR

- Apollo is a program to assist with structural curation



Functional TAIR

□ Functional curators

- *What does e*
- *When and w*
- We hope tha

□ Functional curators

- Allow cross
- TAIR curate

achene
berry
capsule
caryopsis
circumcissile
capsule
cypsela
drupe
follicle
grain
kernel
legume
loculicidal capsule
lomentum
nut
pod
pome
poricidal capsule
schizocarp
septicidal capsule
septifragal capsule
silique

questions:

oidopsis do?

form research in other plants

and vocabularies

and agree upon common terms

The seed-bearing structure
formed from the ovary



FRUIT

Plant Ontology:

Structure:

PO:0009001

Functional curation at TAIR

Catalysis of the reaction:

IAA + UDP-D-glucose = indole-3-acetyl-beta-1-D-glucose + UDP

IAA-Glu synthetase activity
IAA-glucose synthase activity
IAGlu synthase activity
indol-3-ylacetylglucose synthase activity
UDP-glucose:(indol-3-yl)acetate beta-D-glucosyltransferase activity
UDP-glucose:indol-3-ylacetate glucosyl-transferase activity
UDP-glucose:indol-3-ylacetate glucosyltransferase activity
UDPG-indol-3-ylacetyl glucosyl transferase activity
UDPglucose:indole-3-acetate beta-D-glucosyltransferase activity
uridine diphosphoglucose-indoleacetate glucosyltransferase activity



indole-3-acetate beta-glucosyltransferase activity

Gene Ontology:
Molecular function:
GO:0047215

Functional curation at TAIR

Category	Relationship Type	Keyword
GO Biological Process	involved in	nitrate transport
GO Cellular Component	expressed in	plant-type vacuole membrane
GO Molecular Function	has	nitrate transmembrane transporter activity
Growth and Developmental	expressed during	petal differentiation and expansion stage, 4 anthesis, 4 leaf senescence stage, C globular stage, D bilateral stage, E expanded cotyledon stage, F

AT5G14570	has	nitrate transmembrane transporter activity	molecular function	inferred from direct assay: protein expression in heterologous system: none: Chopin, et al. (2007)	The Arabidopsis Information Resource 2007-09-13
AT5G14570	has	nitrate transmembrane transporter activity	molecular function	inferred from sequence or structural similarity: Sequence similarity: homologue almost closely related to: none: Vital, John (2003-09-04)	The Arabidopsis Information Resource 2003-03-29
AT5G14570	expressed in	seed	plant structure	inferred from expression pattern: expression of a reporter gene: none: Chopin, et al. (2007)	The Arabidopsis Information Resource 2007-09-13

Gene

Functional curation at TAIR

Associated Loci

Metagen ethylmethane sulfonate

Inheritance recessive

Allele Type loss-of-function

Description

Contains a substitution (G to A) at the 1226th base pair of AT5G63890 (5th exon) resulting in an amino acid change of glycine to aspartic acid at the 217th amino acid of the GALT protein (TAIR accession:4012145350, RefSeq ID:13170834)

Associated Polymorphisms

Substitution Species Variant

Germplasm Home Breeds

Showing 1 of 1 entries

Phenotype

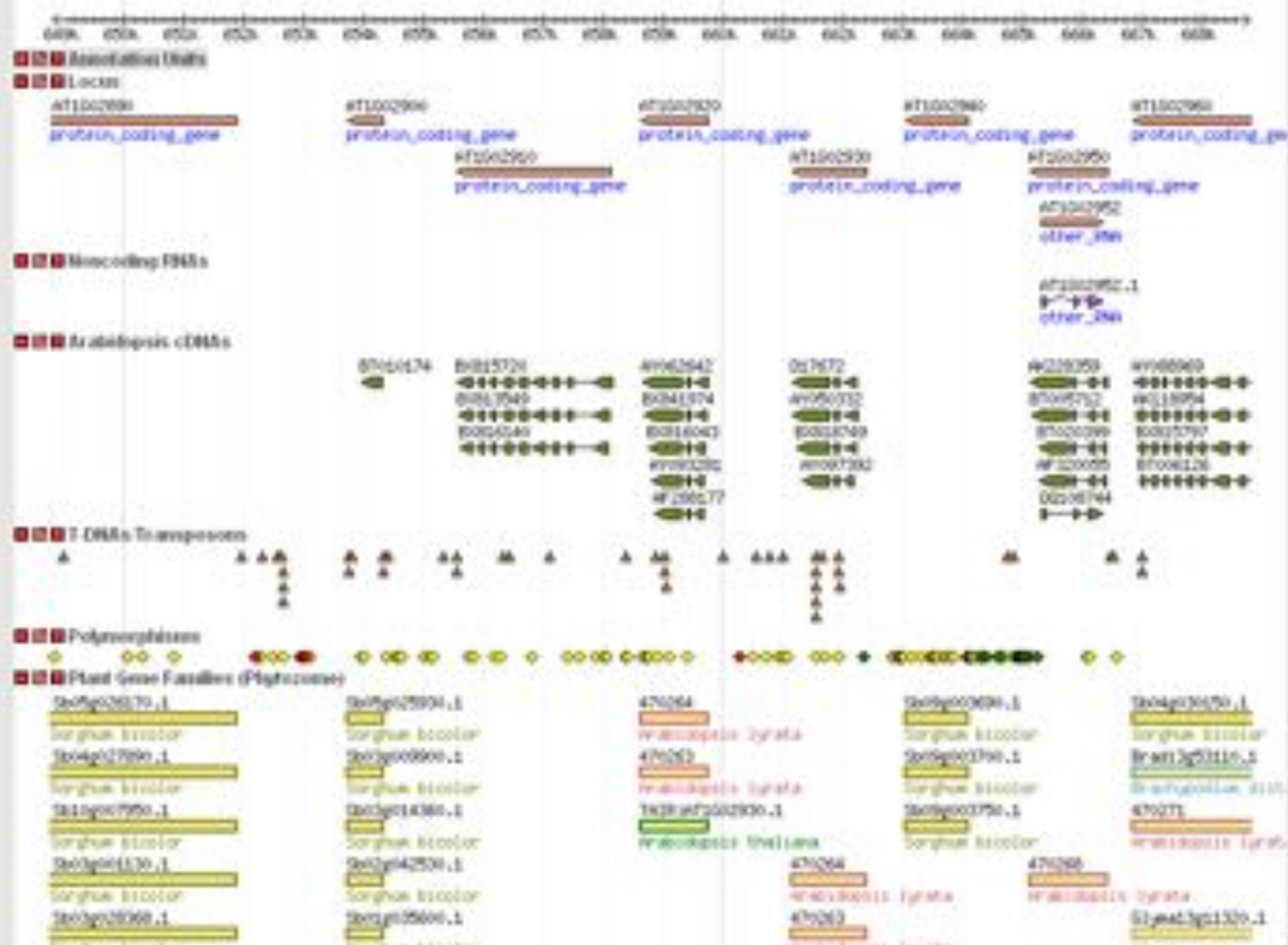
- Loss-of-function seedlings are more viable after 12 d of drought, remain viable 8 d longer, shorter and rounder structure of the leaves, chloroplasts and up-regulated and down-regulated genes in wild type; increased cellulose and sugar derivatives accumulation; no difference in stomatal index on the abaxial

alx8

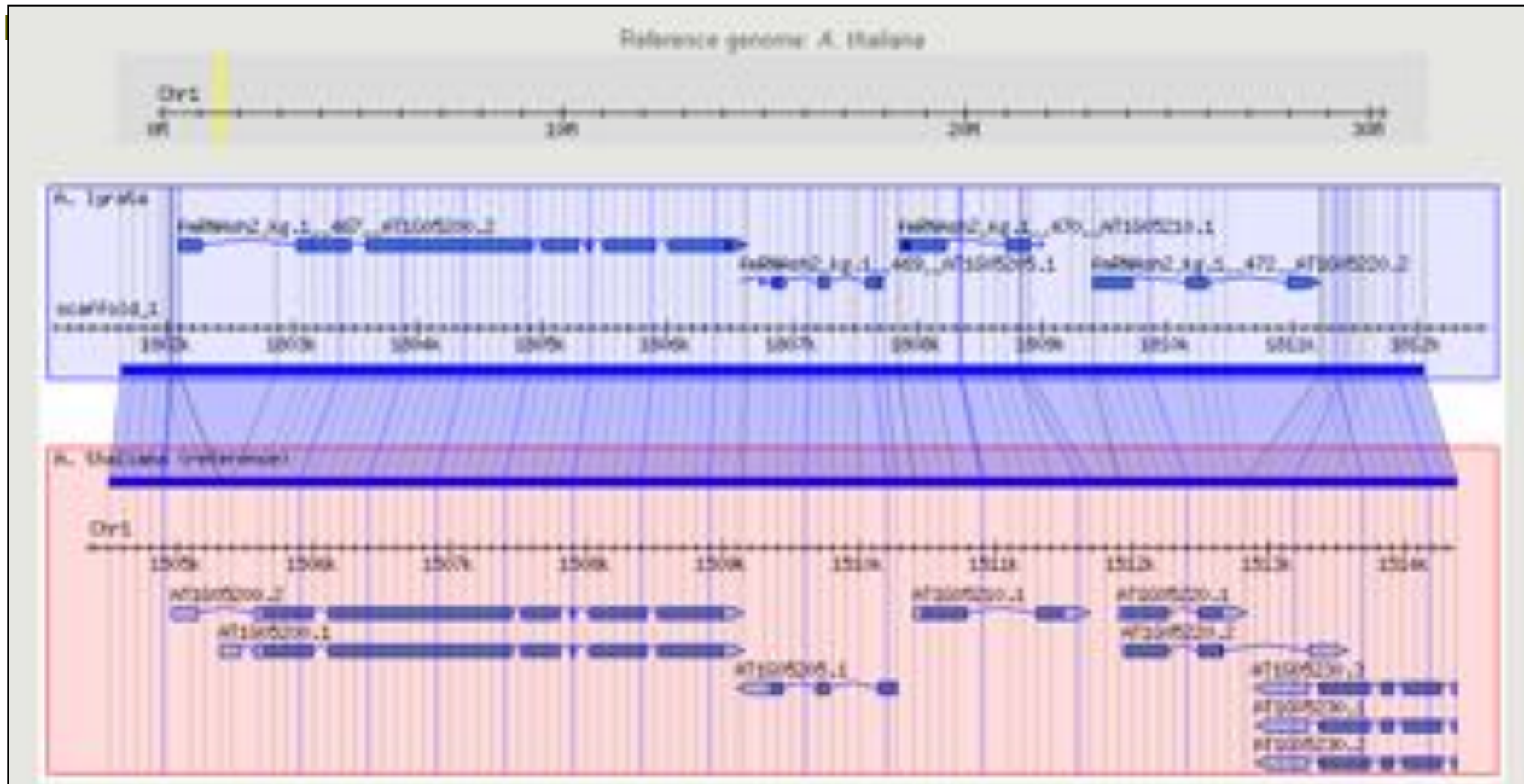
48 day old plant.
Image provided by Barry Pogson

Providing access to external tools and data

[illegible]



Providing Tools at TAIR



Providing Tools at TAIR

1 Search New Gene: From DB: in species: Locate Gene:

3 Multi-Edge Only

- ☒ Biochemical assays
 - ☒ Affinity technology
 - ☒ Enzymatic study
 - ☒ Chromatography technology
 - ☒ Comigration in gel electroph.
 - ☒ Crosslinking study
 - ☒ Cosedimentation
 - ☒ Other biochemical methods
- ☒ Biophysical assays
 - ☒ Fluorescence technology
 - ☒ Surface plasmon resonance
 - ☒ Other biophysical techniques

☒ Node Attribute ☒ Edge Fit

4

Node systematic ID : AT4G10180
Node Common Synonym : DET1
Other Names: ATDET1,
DE-ETIOLATED 1, FUS2, FUSCA 2

Description : Encodes a nuclear-localized protein that acts as a repressor of photomorphogenesis and may be involved

2

step 1: gene(s): HYS (nodes 10, edges 41);

GO Browser Info Met

Edge Table Birds Eye View

10 20 30 40 50 60 70 80 90 100 Gesamte
Barke

[illegible]

Providing

□ Tech team

■ Create

■ Create



TAIR Gene Search

[Help]

Genes may be searched by name, keywords, features, and/or location. In TAIR, a Gene Model is defined as any description of a gene product from a variety of sources including computational prediction, mRNA sequencing, or genetic characterization. A locus is defined as the genomic sequence corresponding to a transcribed unit (e.g. AT2G03360) in the genome. In TAIR, many gene models can exist for a given locus, therefore a search for a gene may result in multiple hits for the same gene name.

Search by Name or Phenotype ⓘ

Gene name

starts with

(Leaving this field blank will return all entries)

Include obsolete genes ☐

Search by Associated Keyword ⓘ

Keyword Term ⓘ

starts with

GO ID (exact match only)

Keyword Type

Any
GO Molecular Function
GO Biological Process
GO Cellular Component

Evidence ⓘ

Any
inferred from direct assay
inferred from electronic annotation
inferred from expression pattern

Restrict by Features ⓘ

Gene Model Type ⓘ

Any
pri tmd
transposable element gene
protein coding

Advanced

gene structure predicted
has associated literature
is sequenced
is not sequenced

Time Restriction ⓘ

☐ only search last 2 months

Other Resources at TAIR

TAIR Community Detail [Help]

Name	Eleanore Wurtzel	
TAIR Accession	Person:1501423405	
Organisms	Rice, Maize, Bacteria, Tomato, Wheat, Arabidopsis	
Primary Job Title	Professor	
Research Interest	regulation of carotenoid/provitamin A biosynthesis in cereal crops; evolution of biosynthetic pathways	
Keywords	carotenoids, molecular biology, provitamin A, genomics, genetics, genes, Rice, Maize, Bacteria, Tomato, Wheat, Arabidopsis	
Address	Dept. Of Biological Sciences Lehman College, The City University of New York 250 Bedford Park Blvd. West Bronx, NY 10468 USA	
E-mail	wurtzel@lehman.cuny.edu	
Websites	http://maize.lehman.cuny.edu	
Office Phone	718-960-8843	
Lab Phone	718-960-4994	
Mobile Phone	516-381-5013	
Affiliations	Organization	Job Title
	*Eleanore Wurtzel Laboratory	
	* denotes Primary Investigator for this organization	
Record last updated	10/31/2006	



TAIR: The Arabidopsis Information Resource

Wall

Info

Photos

Discussions

Events

Share: Post Question Photo Link Video

Suggest to Friends

Subscribe to TAIR: The Arabidopsis Information Resource

Subscribe via SMS

I am an arabidopsis fact and TAIR is my home

23 Friends Like This

6 of 23 Friends

See All



Raymond Chetty



Debbie Alexander



Dor Kelley



Peifen Zhang



Justin Walley



Vanessa Kriup Swing

423 People Like This



Gil Dean



Julia Young



Adwafur Rahman Kolid



Brigitte Kroll



Sahan Liu



John Hugh Snyder



TAIR: The Arabidopsis Information Resource Thank you Dow AgroSciences, GMI, and Syngenta for being our first official TAIR sponsors!
<http://bit.ly/bbv1A>

August 10 at 12:49pm via TweetDeck · Comment · Like

2 people like this.

Write a comment...



TAIR: The Arabidopsis Information Resource eva and the tair advisory board: <http://bit.ly/ajUL90> how many can you name?

August 12 at 2:52pm via TweetDeck · Comment · Like



TAIR: The Arabidopsis Information Resource



TAIR Staff

August 12 at 2:51pm · Comment · Like · Share

6 people like this.



Katika Ilie Looking good!!
June 29 at 9:03pm · Like · Flag

Write a comment...



TAIR: The Arabidopsis Information Resource at the TAIR advisory board meeting today discussing our future priorities.

August 11 at 11:15am via TweetDeck · Comment · Like

	Detail Level: City	Visits ↓	Pages/Visit	Avg. Time on Site	% New Visits	Bounce Rate
1.	Brooklyn	675	6.30	00:07:13	10.22%	25.19%
2.	New York	527	5.75	00:08:14	17.08%	33.02%
3.	Cold Spring Harbor	240	6.48	00:10:01	14.94%	19.09%
4.	Syracuse	199	7.01	00:08:48	14.00%	29.00%
5.	Upton	99	4.66	00:05:43	24.44%	21.11%
6.	Stony Brook	71	2.26	00:01:07	4.23%	16.90%
7.	Bronx	62	7.39	00:07:22	29.03%	41.94%
8.	Corland	38	2.53	00:02:30	0.00%	84.21%
9.	Huntington Station	36	9.69	00:22:34	0.00%	19.44%
10.	Briarcliff Manor	18	1.22	00:00:04	0.00%	94.44%



134,797

This state sent 2,056 visits via 74 cities

How can TAIR contribute to your work?

- If you work on Arabidopsis . . .
 - Find specific information about individual genes and proteins
 - Access large Arabidopsis-specific data sets

- If you work on another species . . .
 - Take your gene / protein of interest and find all the data TAIR contains for its ortholog
 - Look up your favorite:
 - biological process
 - molecular function
 - subcellular compartment
 - organ or tissue
 - developmental stage
 - mutant phenotype

 - Identify many related genes in TAIR and then find orthologs in your species

- But . . . if you want more on plant metabolism

Welcome to the PMN!

- PMN = The **P**lant **M**etabolic **N**etwork
 - Created in 2008
 - Funded by the National Science Foundation



- What is

- What do

- How do

- How can

- How can you help the PMN to grow?



Sue Rhee
(PI)

Peifen Zhang
(Director)

What is the PMN?

□

□

The screenshot shows the PMN website homepage. At the top left is the PMN logo, which consists of the letters 'PMN' in a large, green, serif font, with a small green leaf and a yellow flower-like shape to the right. Below the logo is the text 'Plant Metabolic Network'. To the right of the logo is a faint, light blue background image of a plant's root system. Below the logo and background image is a search bar with a green border. The search bar contains the text 'PlantCyc' and a small blue 'v' icon, followed by a 'search' button. Below the search bar is a dark blue navigation bar with white text links: 'About PMN', 'Databases', 'Downloads', 'Tools', 'Useful Sites', 'Submit Data', 'Help', and 'Feedback'. Below the navigation bar are two main content areas. The left area is titled 'Introduction' and contains text about the PMN project, its goals, and its funding. The right area is titled 'News' and contains information about PMN Summer Road Shows, including dates and locations for the American Society of Plant Biologists and the 20th International Conference on Arabidopsis Research.

PMN
Plant Metabolic Network

Search: PlantCyc

[About PMN](#) [Databases](#) [Downloads](#) [Tools](#) [Useful Sites](#) [Submit Data](#) [Help](#) [Feedback](#)

Introduction

The [Plant Metabolic Network \(PMN\)](#) is a collaborative project among databases and biochemists with a common goal to build a broad network of plant metabolic pathway databases. A central feature of the PMN is [PlantCyc](#), a comprehensive plant biochemical pathway database, containing curated information from the literature and computational analyses about the genes, enzymes, compounds, reactions, and pathways involved in primary and secondary metabolism.

PMN is funded by the [National Science Foundation](#) (Grant # 0640750), governed by an [Editorial Board](#) composed of internationally renowned scientists, and executed at the [Carnegie Institution for Science](#), Department of Plant Biology.

News

PMN Summer Road Shows!

Come see PMN [presentations](#) and posters at conferences and universities this summer.

- [American Society of Plant Biologists](#)
July 18 - 22, Honolulu, HI
- [20th International Conference on Arabidopsis Research](#)
June 30 - July 4, Edinburgh, Scotland

- Facilitate research that benefits society

Connecting the PMN to important research efforts

- **More nutritious foods**

- vitamin A biosynthesis, folate biosynthesis . . .

- **Medicines**

- morphine biosynthesis, taxol biosynthesis . . .

- **More pest-resistant plants**

- maackiain biosynthesis, capsidiol biosynthesis . . .

- **Higher photosynthetic capacity and yield in crops**

- chlorophyll biosynthesis, Calvin cycle . . .

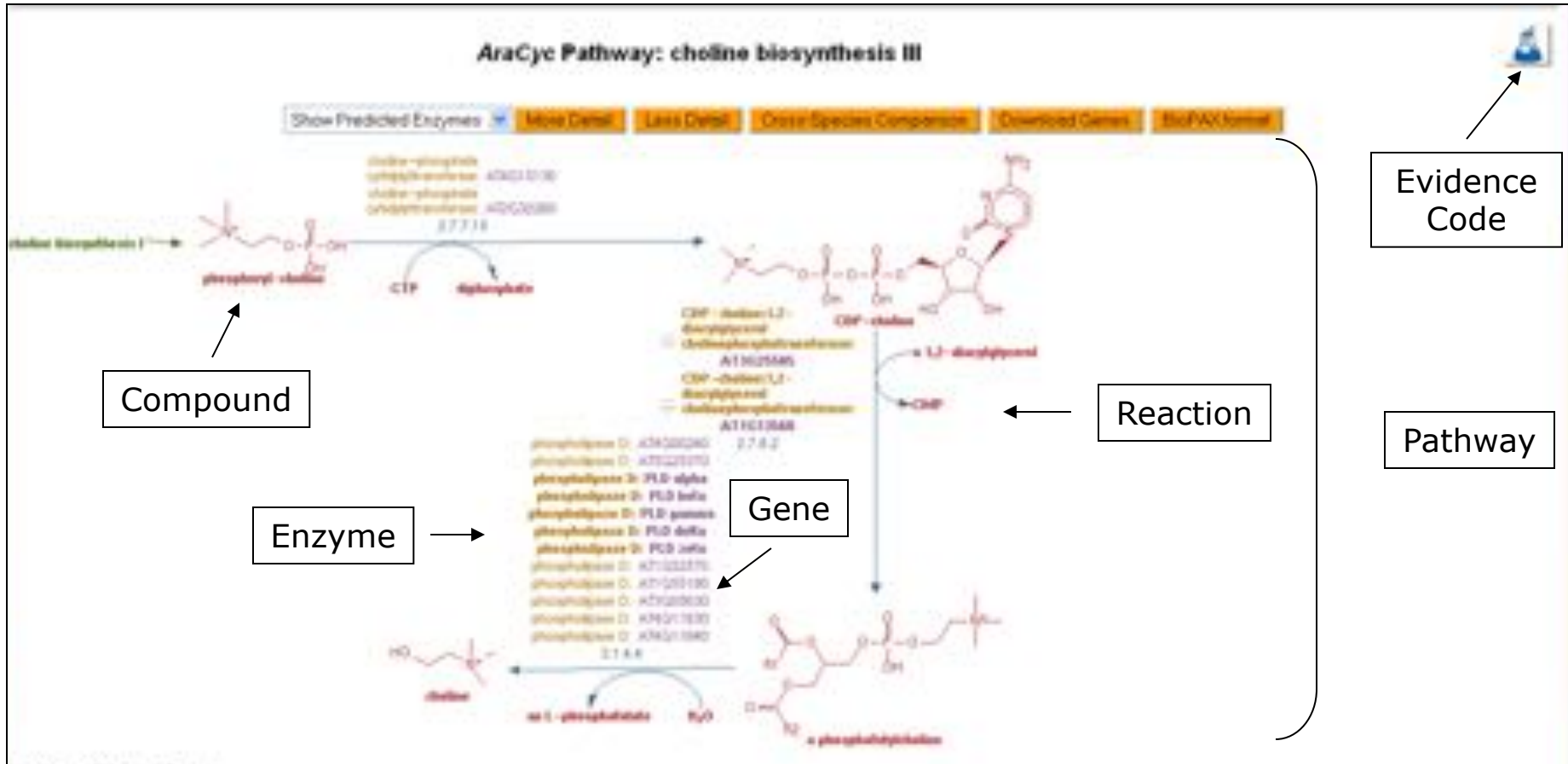
- **Better biofuel feedstocks**

- cellulose biosynthesis, lignin biosynthesis . . .

- **Many additional applications relevant to rational metabolic engineering**

- ethylene biosynthesis, resveratrol biosynthesis . . .

What data are in the PMN?



Pathway Tools software provided by collaborators at SRI International

PMN databases

- Current PMN databases: **PlantCyc, AraCyc, PoplarCyc**
 - Coming soon: databases for wine grape, maize, cassava, Selaginella, and more . . .
- Other plant databases accessible from the PMN:

PGDB	Plant	Source	Status
RiceCyc **	Rice	Gramene	some curation
SorghumCyc	Sorghum	Gramene	no curation
MedicCyc **	Medicago	Noble Foundation	some curation
LycoCyc **	Tomato	Sol Genomics Network	some curation
PotatoCyc	Potato	Sol Genomics Network	no curation
CapCyc	Pepper	Sol Genomics Network	no curation
NicotianaCyc	Tobacco	Sol Genomics Network	no curation
PetuniaCyc	Petunia	Sol Genomics Network	no curation
CoffeaCyc	Coffee	Sol Genomics Network	no curation

** Significant numbers of genes from these databases have been integrated into PlantCyc

PMN database content statistics

	PlantCyc 4.0
Pathways	685
Enzymes	11058
Reactions	2929
Compounds	2966
Organisms	343

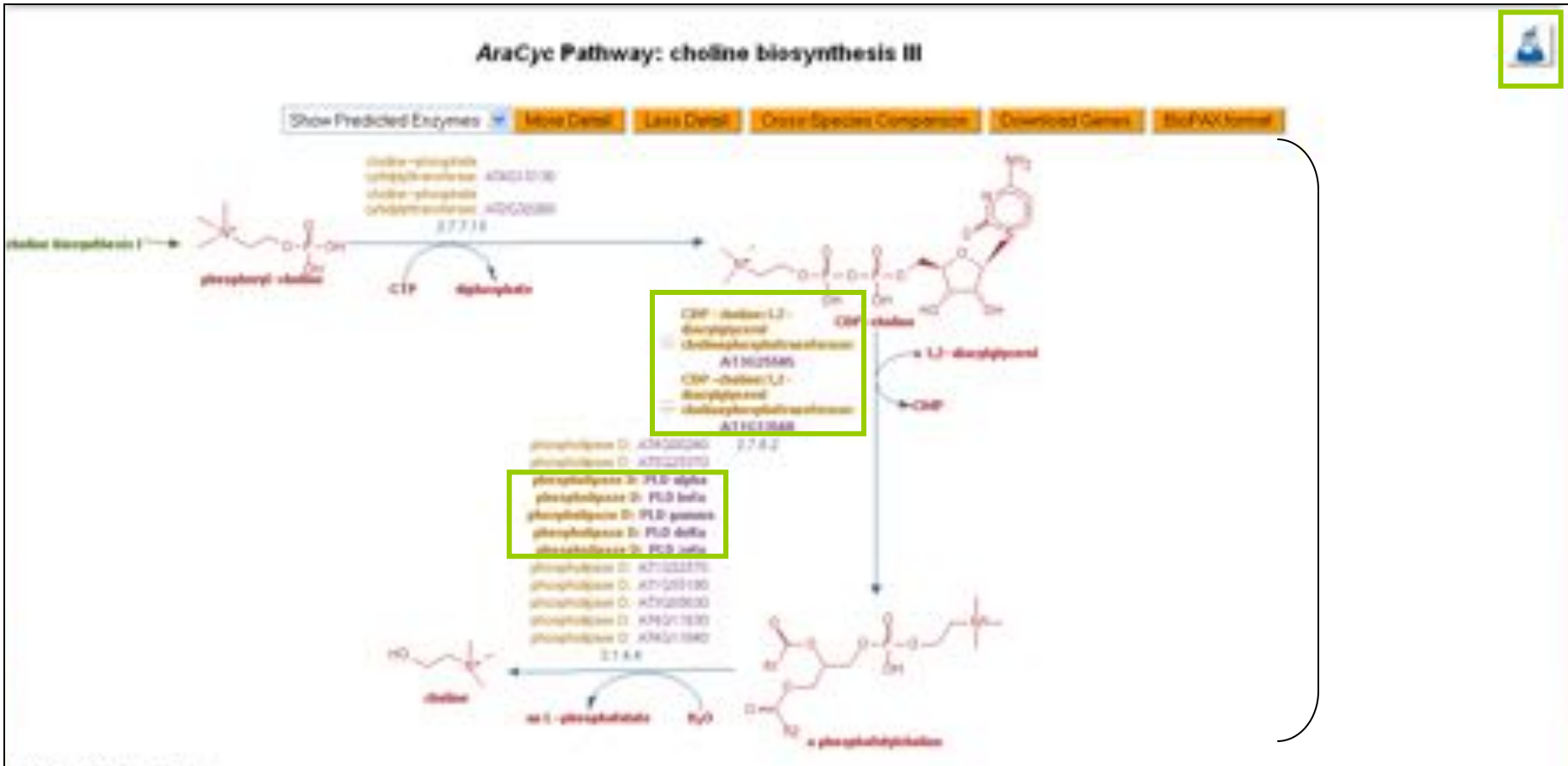
Species (A - E) [back to top](#)

- *Abies grandis* (yellow fir, white fir, silver fir, lowland fir, grand fir)
- *Acer pseudoplatanus*
- *Adonis aestivalis*
- *Adonis annua* (pheasants-eye, blood-drops)
- *Ajuga reptans*
- *Albizia julibrissin*
- *Alectonia*
- *Alliaceae* (onion family)
- *Allium cepa*
- *Allium sativum*
- *Allium tuberosum*
- *Aloe arborescens*
- *Amanita muscaria*
- *Ammi majus*
- *Amorpha fruticosa*
- *Anabaena variabilis*
- *Anchusa officinalis*
- *Anisodus acutangulus*
- *Antirrhinum majus* (snapdragon)
- *Apium graveolens*
- *Aquilegia vulgaris*
- *Arabisopsis lyrata* (brake rockcress)
- *Arabisopsis thaliana*, Col (thale cress, mouse-ear cress)
- *Arabisopsis thaliana*, Ler (thale cress, mouse-ear cress)
- *Arachis hypogaea*
- *Artemisia annua* (sweet Annie)
- *Asclepias syriaca*
- *Atropa belladonna* (deadly nightshade)
- *Avena sativa* (cultivated oat)
- *Berberis beaniana*
- *Berberis stolonifera*
- *Berberis Wilsonae*

How does experimentally verified data enter the PMN?

- Biocurators perform manual curation
 - Use journal articles to enter information
 - Receive helpful messages from researchers
 - Request specific data from experts
 - Invite editorial board members to review metabolic domains

Pathway information



Pathway information

Superclasses: [Biosynthesis](#) -> [Fatty Acids and Lipids](#) -> [Choline Biosynthesis](#)

Summary:

General information: Choline is a fundamental metabolite in plants because of its contribution to the synthesis of the membrane phospholipid phosphatidylcholine, which accounts for 40 to 60% of lipids in non-plant plant membranes [[Mun02](#)]. Choline is also a precursor for the formation of glycine betaine ([glycine betaine Biosynthesis II \(plants\)](#)) in certain plants such as spinach, where this osmoprotectant is accumulated and confers also tolerance to salinity, drought, and other environmental stresses. In addition choline has been recognized as an essential nutrient for humans [[McNeil01](#)].

The choline biosynthetic pathway enables plants to decouple choline synthesis from lipid metabolism (Kennedy pathway - [N-methylglyceral Biosynthesis](#)) and provides them with the metabolic flexibility to adapt to environmental conditions where large and variable amounts of choline are beneficial for survival [[Porten01](#)].

Pathway information: The first step in choline biosynthesis is the direct decarboxylation of serine to ethanolamine [[Porten01](#)], which is catalyzed by a serine decarboxylase unique to plants [[Porten02](#)]. Ethanolamine is widely recognized as the entrance compound to choline biosynthesis.

The pathway variant displayed (nucleotide pathway) represents the biosynthetic route as found in diverse plant families. The synthesis of choline from ethanolamine may take place at three parallel pathways, where three consecutive N-methylation steps are carried out either on free-bases [[Pudhomme02](#)], phospho-bases [[Nucco02](#)], phosphatidyl-bases [[McNeil01](#)] or a mixture of the latter [[Duke00a](#)] [[Duke00b](#)] [[Mull01](#)].

The synthesis of intermediates on both the phospho-base and phosphatidyl-base level includes the nucleotide pathway via CDP-phosphoaminoalcohol and the methylation pathway. However, it has been pointed out that the synthesis of phosphatidylethanolamine and phosphatidylcholine is characterized by a high degree of interaction and fluctuation on the various levels of arising intermediates. Consequently, it has been assumed that the reactions embedded in the nucleotide and methylation pathways may be two characteristics of one overall phosphoaminoalcohol pathway for the formation of phosphatidylcholine in plants [[Kinner03](#)].

The release of choline from the different pathway levels is also species-specific. Phosphocholine can either be directly dephosphorylated to release choline as observed in spinach [[Summer03](#)] or incorporated into phosphatidylcholine with the subsequent release of choline, as in tobacco [[McNeil02](#)]. The latter reaction has been shown to be specifically catalyzed by phospholipase D ([phospholipases](#)) in cacti bean [[Wang04](#)]. Although a well-defined physiological role of phospholipase D (PLD) still await further research, progress has been made to assign some members of the heterogeneous family of PLDs to distinct cellular functions [[Vick00](#)]. The remaining enzymes involved in this pathway, phosphoaminoalcohol cytidyltransferase and CDP-aminoalcohol phosphotransferase, cover a broader spectrum of substrates. This may be beneficial to process the heterogeneous mixture of possible substrates but it also indicates that the pathway flux is probably controlled more upstream [[Kinner03](#)].

Superpathways: [superpathway of choline biosynthesis](#)

Variants: [choline biosynthesis II](#), [choline biosynthesis I](#)

Unification Link: [MetaCyc:PWY-3681](#)

Legend for Pathway Diagram

If an enzyme name is shown in **bold**, there is experimental evidence for this enzymatic activity.

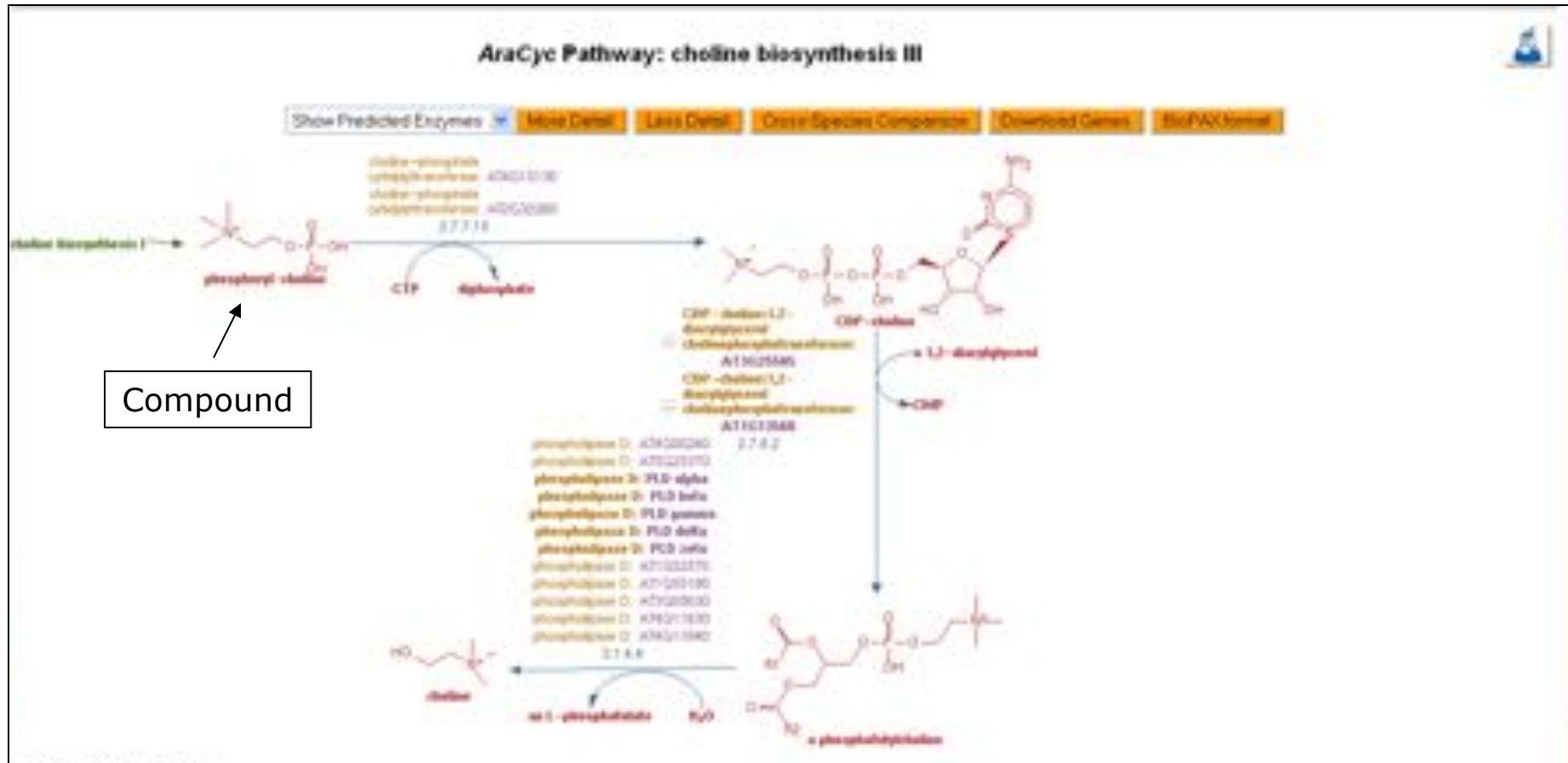
References

[Duke00a](#), Duke AH, Mudd SH (2000) "Estimates of phosphatidylcholine synthesis in *Lotus*, *soybean*, and *corn*." *Plant Physiol* 128(3): 1330-1340.

[Duke00b](#), Duke AH, Mudd SH (2000) "Phosphatidylcholine synthesis: Differing patterns in *soybean* and *corn*." *Plant Physiol* 128(3): 854-861.

[Hill01](#), Hill WD, Rhodes D, Hanson AD (2001) "Redox-state evidence implicating phosphoryl and phosphatidyl bases as intermediates in betaine synthesis by water-stressed *barley leaves*." *Plant Physiol* 128(2): 601-614-602.

Compound information



Compound: CDP-choline

Com

Synonyms: citicoline , citicholine , citilins , cytocholine , cytidine 5'-diphosphocholine , cytidine diphosphate choline

Synonyms

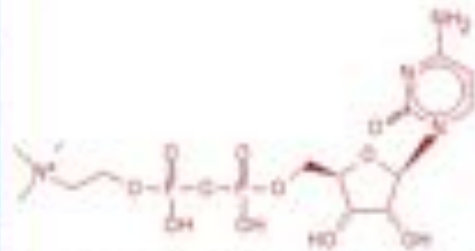
Superclasses: [a nucleic acid component](#) -> [a base derivative](#)
[a nucleic acid component](#) -> [a pyrimidine-related compound](#)

Classification(s)

Empirical Formula: $C_{14}H_{27}N_4O_{13}P_2$

Molecular Weight: 489.34 daltons

Molecular Weight / Formula



Smiles: C(=O)OP(=O)(=O)OP(=O)(=O)OCC(N+)(C)CC(C)C(=O)C(=O)C(=O)N(C)C(=O)N(C)C

Unification Links: CAS:987-78-0

Gibbs Energy of Formation (kcal/mol, estimated): -116.7

In Pathway Reactions as a Reactant:

[phospholipid biosynthesis:](#)

[a 1,2-dicacylglycerol](#) + CDP-choline = [a phosphatidylcholine](#) + CMP

Appears as Reactant

[choline biosynthesis III:](#)

[a 1,2-dicacylglycerol](#) + CDP-choline = [a phosphatidylcholine](#) + CMP

In Pathway Reactions as a Product:

[phospholipid biosynthesis:](#)

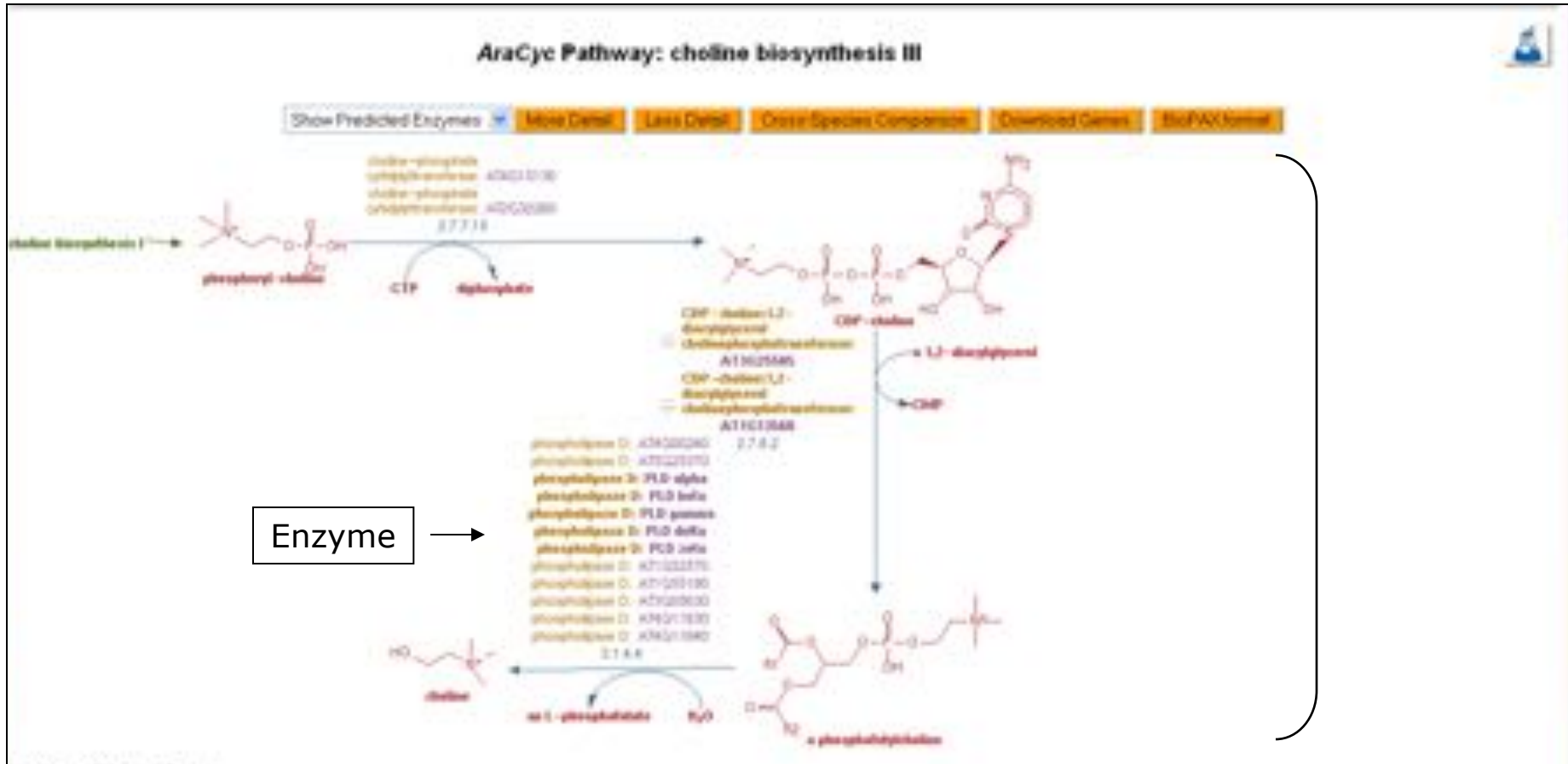
[phosphorylcholine](#) + CTP = CDP-choline + diphosphate

Appears as Product

[choline biosynthesis III:](#)

[phosphorylcholine](#) + CTP = CDP-choline + diphosphate

Enzyme information



Enzyme information

Arabidopsis Enzyme: phosphatidyltransferase

Enzymatic reaction of: cholinephosphotransferase (phosphatidyltransferase)

[1,3-diacylglycerol](#) + CDP-choline \rightleftharpoons a phosphatidylcholine + CMP

Reaction

The reaction direction shown, that is, $A + B \rightleftharpoons C + D$ versus $C + D \rightleftharpoons A + B$, is in accordance with the Enzyme Commission system.

Reversibility of this reaction is unspecified.

In Pathways: [choline biosynthesis II](#), [phospholipid biosynthesis](#)

Pathway(s)

Summary:

Two cDNA's from *Arabidopsis thaliana* (AtAAPT1, AtAAPT2) have been isolated from an *Arabidopsis* cDNA library using the AAPT cDNA from soybean as a heterologous hybridization probe. Both cDNA's encode aminoalcoholphosphotransferases involved in the nucleotide pathway of the biosynthesis of phosphatidylethanolamine (PE) and phosphatidylcholine (PC) as demonstrated by gene expression in yeast mutants lacking these enzyme activities. It has been demonstrated that both AtAAPT1 and AtAAPT2 convert CDP-ethanolamine and CDP-choline into the corresponding phosphatidylethanolamines although with slight differences regarding the substrate preference. AtAAPT2 showed a higher preference for CDP-choline over CDP-ethanolamine in comparison to AtAAPT1 and was also inhibited to a lesser degree by Ca^{2+} and Cytidine monophosphate (CMP) than AtAAPT1 [[Goode99](#)]. Both enzymes (AtAAPT1, AtAAPT2) were able to catalyze the reverse reaction supporting the proposal that diacylglycerol, involved as substrate in both PC and triacylglycerol biosynthesis [[triacylglycerol biosynthesis](#)], is in equilibrium with PC and maintains this equilibrium via the reversibility of the cholinephosphotransferase reaction [[Slack85](#)]. AtAAPT1 and AtAAPT2 seem to represent the only aminoalcoholphosphotransferases in *Arabidopsis* as concluded from the Southern blotting patterns. Both polypeptides contain seven membrane spanning regions as shown by their hydropathy files. This finding is in agreement with other AAPT's isolated so far and confirms their localization in membranes [[Goode99](#)].

Inhibitors (Allosteric): [CMP](#) [[Goode99](#)]

Inhibitors (Unknown): [Ca²⁺](#) [[Goode99](#)]

Primary Physiological Regulators of Enzyme Activity: [CMP](#)

Inhibitors, Kinetic Parameters, etc.

Summary

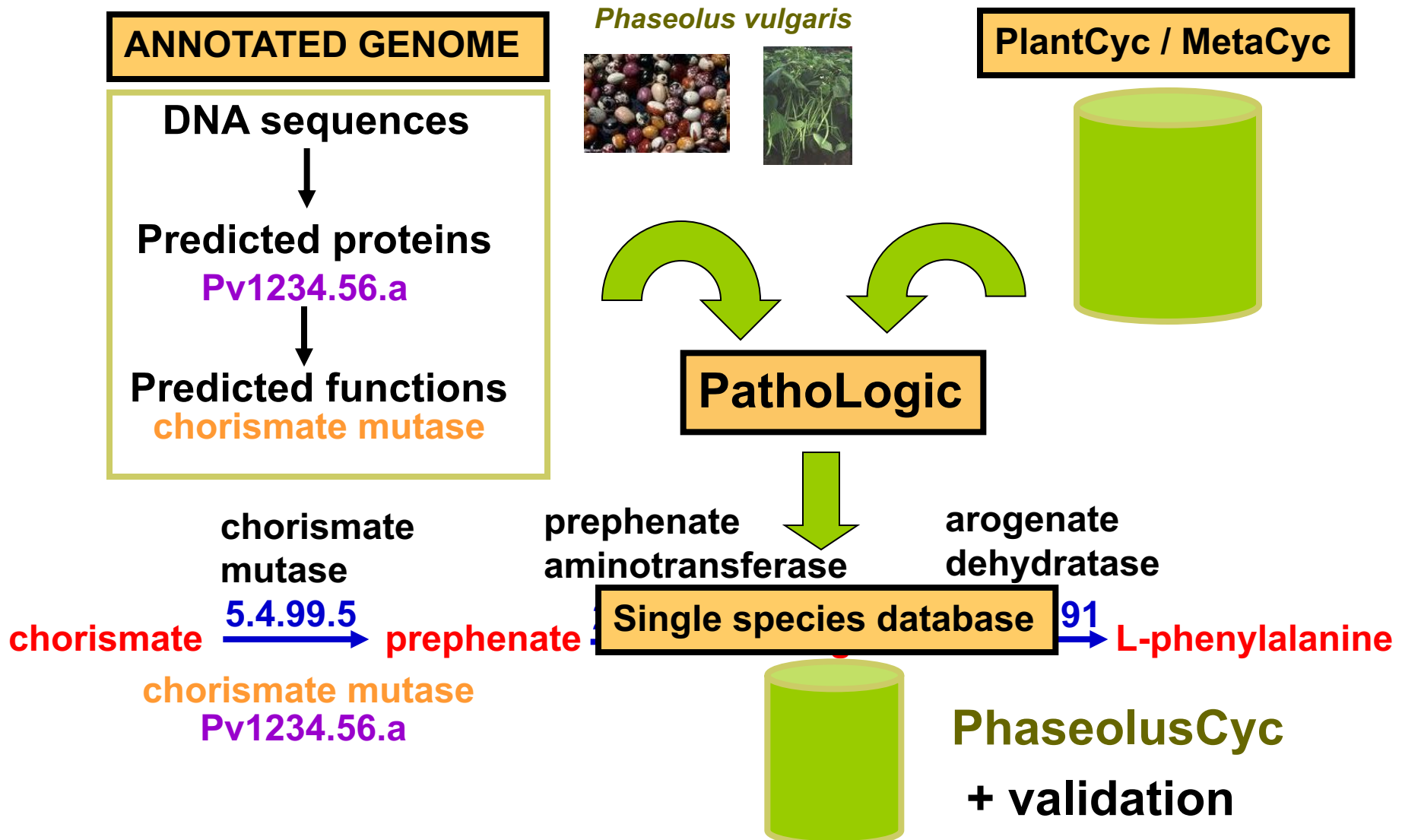
References

[Goode99](#), Goode JH, Dewey SE, (1999) "Characterization of aminoalcoholphosphotransferases from *Arabidopsis thaliana* and soybean." *Plant Physiol. Biochem.* (1999), 37(5), 445-457.

[Slack85](#), Slack CR, Broughan PG, Brown JA, Gardner SE, (1985) "Some properties of cholinephosphotransferase from developing safflower cotyledons." *Biochim. Biophys. Acta* (1985), 833, 439-446.

References

How does computationally predicted data enter the PMN?





How can researchers use the PMN?

- Learn background information about particular metabolic pathways
 - Utilize simple and advanced search tools
 - Quick search bar



- Specific search menus

How can researchers use the PMN?



Search Database *PlantCyc* [change](#)

About PMN

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Databases Overview

Compounds

PlantCyc Compound Search

▼ Search for compound by name or ID

Enter a compound name, or a database identifier from this database or from an external database such as ChEBI, LIGAND, PubChem or CAS. This database may not contain mappings to all of these other databases. Partial names will generate a substring search on compound names only (not on database identifiers).
Examples: "tryptophan", "C00036"

▶ Search/Filter by ontology (inactive)

▶ Search/Filter by molecular weight (inactive)

▶ Search/Filter by chemical formula (partial or full) (inactive)

▶ Search by InChI string (inactive)

How can researchers use the PMN?

- Compare metabolism across species

Pathway Class: Biosynthesis - Amines and Polyamines Biosynthesis		AraCys col	P. trichocarpa
glycine betaine biosynthesis III (plants)		X	X
Organism	Evidence Glyph	Enzymes and Genes for glycine betaine biosynthesis III (plants)	
AraCys.col		EC# 1.14.15.7	None
		RxN-6268	Spontaneous
		EC# 1.2.1.8	3-chloroalanyl aldehyde dehydrogenase: AT1G74920 3-chloroalanyl aldehyde dehydrogenase: AT3G48170
P. trichocarpa		EC# 1.14.15.7	None
		RxN-6268	Spontaneous
		EC# 1.2.1.8	betaine-aldehyde dehydrogenase: JGI-666405 betaine-aldehyde dehydrogenase: JGI-661953

☐ Arabidopsis thaliana col
 ☐ PlantCyc
 ☐ Populus trichocarpa



How will the PMN grow in the future?

- ❑ Help from the research community!!!
- ❑ You are the experts with great knowledge to share!



Building better databases together

- ❑ To submit data, report an error, or volunteer to help validate . . .
 - Send an e-mail: **curator@plantcyc.org**
 - Use data submission “tools”



- **Meet with me this afternoon**
 - ❑ . . . or later this week
 - ❑ . . . or later this year

Building better databases together

- **Details are very, very welcome!!**
 - Reactions:
 - All co-factors, co-substrates, etc.
 - EC suggestions – partial or full
 - Compounds
 - Structure – visual representation / compound file (e.g. mol file)
 - Synonyms
 - Unique IDs (e.g. ChEBI, CAS, KEGG)
 - Enzymes
 - Unique IDs (e.g. At2g46480, UniProt, Genbank)
 - Specific reactions catalyzed

Community gratitude

□ We

The screenshot shows the PMN website interface. The top navigation bar includes links for Search, Tools, Downloads, Useful Sites, Submit Data, Help, and Feedback. The left sidebar contains a list of links: About PMN, Project Overview, Documentation, Presentations, Statistics, Release Notes, Contact Info, News, PMN Staff, Collaborators, Contributors (highlighted with a yellow box), and Editorial Board. The main content area is titled 'PMN Contributors' and contains the following text:

PMN contributors from around the world have added to or helped to improve the content of AraCyc, PlantCyc, and the other PlantCyc-derived databases that are part of the PMN.

In addition to the active contributions from the PMN editorial board and PMN collaborators, the following individuals have contributed significantly in improving the content of PlantCyc, AraCyc, and the other PlantCyc-derived databases that are part of the PMN.

- Some contributors have [contacted us](#) with suggestions and revisions.
- Some have generously responded to appeals for help from the curators.
- Some have attended curatorial jamborees.

Contributors for PMN release... June 2010

- Lodger Beechhus - Technische Universität Braunschweig, Germany
- David Doon - Durham University, UK
- Natalia Dudareva - Purdue University, USA
- Andrzej Guranowski - The University of Life Sciences, Poland
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- Jungmi Hur - University of California, Los Angeles, USA
- Peter Lundquist - Cornell University, USA
- Rüdiger Sadre - RWTH Aachen University, Germany
- Bernd Schneider - Max Planck Institute for Chemical Ecology Braunschweig Campus, Germany
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- [Gene Family Data](#)
- [Online Submission for Authors and Others](#)
- [Gene Structure Annotations/Modifications](#)
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- [Metabolic Pathway Data](#)
- [Phenotypes](#)
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- [2010 Functional Genomics Gene List](#)

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Data Submission Overview

We encourage users of TAIR to share their data.

Please see below Data Types Accepted by TAIR by TAIR).

DATA TYPES ACCEPTED BY TAIR

Please take a look at the specific guidelines asked to download an Excel worksheet to fill in suggestions, please contact us at: [curating@tair.org](#)

ARDC Stock Donation - Guidelines - Seed Don

For seed stocks, cones, vectors, libraries, etc. of interest to the community.

External Links - Guidelines - Submission Form

We provide links from TAIR detail pages to our

Gene Class Symbol Registration - Guidelines -

Reserve a gene symbol prior to publication to ensure uniqueness and consistency in gene nomenclature.

Gene Family Data - Guidelines - Submission Form

For inclusion on the TAIR Arabidopsis Gene Family page.

Gene Function Submission Forms for Authors and Others - Online Form - Large Dataset Submission Form

[Submit Overview](#)[ARDC Stock Donation](#)[External Links](#)[Gene Class Symbol](#)[Registration](#)[Gene Family Data](#)[Online Submission for Authors and Others](#)[Gene Structure](#)[Annotations/Modifications](#)[Locus Identifier Request](#)[Marker and Polymorphism Data](#)[Metabolic Pathway Data](#)[Phenotypes](#)[Protocols](#)[2010 Functional Genomics](#)[Gene List](#)[Gene List](#)

Have just a few genes?

Fill out our online submission form. * means a field is required.

TAIR curators will review your submission and will get in touch with any issues or questions using the e-mail from your user profile.

Molecular Function Annotations

Annotating locus from article

Molecular Function

Examples:

ion transmembrane transporter activity

protein kinase activity

phytoene synthase activity

Start typing, then choose from list or add a new term.

Then click outside the field to accept the new term.

Method

Example:

Enzyme assays

Choose a method or enter a new one.

histidine

Choose method or enter a new one

Delete

Term	Id	Synonyms
CAU codon-amino acid adaptor activity	GO:0033425	histidine tRNA
CAC codon-amino acid adaptor activity	GO:0033426	histidine tRNA
protein histidine kinase binding	GO:0043424	histidine kinase binding
protein histidine kinase activity	GO:0004673	histidine kinase activity
polar-amino acid-transporting ATPase activity	GO:0015426	histidine permease activity
histidine-tRNA ligase activity	GO:0004821	histidine transase activity
L-histidine 2-oxoglutarate aminotransferase activity	GO:0008110	histidine transaminase activity
L-histidine transmembrane transporter activity	GO:0006290	L-histidine transporter activity
histidine decarboxylase activity	GO:0004398	
histidine ammonia-lyase activity	GO:0004397	
L-phenylalanine-pyruvate aminotransferase activity	GO:0047312	histidine aminotransferase activity
histidine N-acetyltransferase activity	GO:0047981	histidine acetyltransferase activity

Biological networking . . .

- Please use our data
- Please use our tools
- Please help us to improve our databases!
- Please contact us if we can be of any help!



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Eva Huala (*PI-TAIR*)
Peifen Zhang (*Director-PMN*)

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- Donghui Li (*curator*)
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- A. S. Karthikeyan (*curator*)
- Marga Garcia (*curator*)
- Leonore Reiser

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- Ron Caspi (SRI)
- Suzanne Paley (SRI)
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- Anuradha Pujar (SGN)
- Gramene and MedicCyc

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- Larry Ploetz (*Sys. Administrator*)
- Anjo Chi
- Raymond Chetty
- Cynthia Lee
- Shanker Singh
- Chris Wilks

PMN project post-doc

- Lee Chae



Biological networking . . .

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Out-takes

- The following slides are relevant but were removed from the presentation due to time constraints



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Arabidopsis has good model organism traits

- ❑ Fast life cycle (6 weeks)
- ❑ Thousands of plants fit in a small space
- ❑ Fairly easy to grow
- ❑ Thousands of seeds produced by each plant
- ❑ Self-fertile (in-breeding)
- ❑ Many different subspecies/ecotypes
- ❑ Serves as a good model for crop plants
- ❑ But why Arabidopsis instead of other plants?



Arabidopsis data explosion

- TONS of data are generated about Arabidopsis
 - Over 2400 “Arabidopsis” articles published each year are indexed in PubMed
 - Tens of thousands of mutants have been generated
 - Hundreds of microarray experiments have been performed
 - Proteomics and metabolomics studies are becoming popular
 - “1001” Arabidopsis genomes are being sequenced
 - Large-scale phenotypic studies are scheduled to start soon
- TAIR tries to bring data together to benefit scientists and society

Pr

□ T

Enter a peptide sequence or pattern (examples):

SDExCA

Choose a Sequence Database (click and

All public Arthropod sequences can be found within

TAFIS Proteins (protein)

[\[START PATTERN SEARCH\]](#) or [reset form](#)

PLEASE WAIT FOR EACH REQUEST TO COMPLETE

More Options :

Maximum hits: 75000 [more](#)

RNA, strand: both strands [more](#)

Mismatch: 2 [more](#)

Mismatch Type: ☐ Insertions ☐ Deletions ☐ Substitutions

Minimum Hits per Sequence: 1 [more](#)

Maximum Hits per Sequence: 100 [more](#)

Supported Pattern Syntax and Examples:

Search Type	Character	
Peptide Searches	$\{F\}LW\{R\}Q\{C\}P$ TSHEQWR	Exact match
	$\{J\}$ more	Any letter of J
	$\{U\}$ more	Any letter of U
	$\{B\}$ more	O or N
	$\{Z\}$ more	E or Q
	$\{X\}$ more	Any amino acid

Hits found	66
Sequences with hits	66
Sequences searched	13810
Bytes searched	13434913
Pattern	SDExCA
Database searched	TAFIS Proteins (protein)
Download all matches as a textfile	download

Next Results >

Hit #	Sequence name	# of hits	Hit pattern	Matching Positions		Hit sequence
				start	end	
1	KT1010050.1	1	SDExCA	337	343	sequence
2	KT1011020.1	1	SDExCA	242	248	sequence
3	KT1018710.1	1	SDExCA	499	505	sequence
4	KT1020020.1	1	SDExCA	195	201	sequence
5	KT1020020.2	1	SDExCA	447	453	sequence
6	KT1021700.1	1	SDExCA	209	214	sequence
7	KT1021700.2	1	SDExCA	209	214	sequence
8	KT1022630.1	1	SDExCA	50	56	sequence
9	KT1028330.1	1	SDExCA	545	551	sequence
10	KT1027950.1	1	SDExCA	611	617	sequence

What data are in the PMN?

- ❑ Plants provide crucial benefits to the ecosystem and humanity
- ❑ A better understanding of plant metabolism may contribute to:
 - More nutritious foods
 - New medicines
 - More pest-resistant plants
 - Higher photosynthetic capacity and yield in crops
 - Better biofuel feedstocks
 - Improved industrial inputs (e.g. oils, fibers, etc.)
 - Enhanced ability to do rational metabolic engineering
 - . . . many more applications
- ❑ How can the PMN help?

What metabolites are in the PMN?

- “Primary” metabolites (“essential”)
 - sugars
 - glucose, fructose, . . .
 - amino acids
 - tryptophan, glutamine, . . .
 - lipids
 - waxes, phosphatidylcholine , . . .
 - vitamins
 - A, E, K, C, thiamine, niacin, . . .
 - hormones
 - auxin, brassinosteroids, ethylene . . .

What metabolites are in the PMN?

- “Secondary” metabolites (important, but not “essential”)
 - terpenoids
 - orzyalexin, menthol, . . .
 - organosulfur compounds
 - glucosinolates, camalexin . . .
 - isoflavonoids
 - glyceollin, daidzein. . .
 - alkaloids
 - caffeine, capsaicin, . . .
 - polyketides
 - aloesone, . . .
 - many more . . .

How do computational predictions enter the PMN?

- **New sets of DNA sequences -> predicted proteome**
 - Genomes are sequenced
 - Large RNAseq or EST data sets are created
- **Predicted proteome -> set of predicted enzyme functions**
 - Performed using computer algorithms
 - The PMN is working to develop better algorithms to increase the accuracy of the predictions
- **Set of predicted enzyme functions -> set of predicted metabolic pathways**
 - The PathoLogic program uses a reference database to predict the metabolic pathways for the enzyme sets
- **Set of predicted metabolic pathways -> set of “validated” metabolic pathways**
 - Curators remove incorrect information and add additional data