Plant Reactome: Leveraging Neo4j Knowledge Graphs for Pathway Integration, Visualization, Analysis, and Discovery

https://plantreactome.gramene.org



An overview of the Plant Reactome knowledgebase



 Open access and free-of-cost, FAIR data









We manually curate reference pathways in rice

- Gene-orthology-based pathway projections for 129 species
- Open access, free-of-cost, FAIR data
- Videos Tutorials, and Training material





Plant Reactome is a Reaction Network Database

- Explicitly describes biological processes as reactions that transport, transform, and reorganize physical entities.
- Thus, provides molecular representations of many biological events;







Interoperability at the level of data annotation Controlled vocabularies for entities and events

		Referrers Dialog		
[GO_MolecularFunction:261470] (+)-abscisic acid 8'-hydroxylase activity's Referrers:				
Referrer Prop	oerty Name	Referrer		
	Go: Molecular Function	(+)-abscisic acid 8'-hydroxylase activity of (+)-abscisic acid 8'-hydroxylase [cytosol]		
activity		(+)-abscisic acid 8'-hydroxylase activity of AT1G19630 [cytoplasm]		
		(+)-abscisic acid 8'-hydroxylase activity of AT4G19230-MONOMER [cytoplasm]		

Entities: proteins, Gene, transcripts GO: Cellular Component

EntityWithAccessionedSequence Properties			
Property Name	Value		
🖉 cellType			
e compartment	Inucleoplasm		
er endCoordinate	416		
^r hasModifiedResidue			
referenceEntity	UniProt:Q2R435 LAZY1		
e ^r startCoordinate	1		
♣ DB_ID	-104		
👎 _displayName	LAZY1 [nucleoplasm]		
🖉 authored	🕸 Naithani, Sushma, 2022-09-21		
👌 created			
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GO: Biological process Property Na deletedInstanceDB_ID entityFunctionalStatus 🗗 input d output 1 x LAZY1 [nucleoplasm] 🚰 DB_ID -394 displayName Expression of LAZY 1 doRelease authored Naithani. Sushma. 2022-09-21 catalystActivity catalystActivityReference nucleoplasm ^m compartment d created crossReference definition deletedStableIdentifier disease r edited entityOnOtherCell Inferred from Experiment evidenceType 0 figur goBiologicalProcess mRNA transcription nasinteraction InferredFrom IsChimeric false RNA-Seq Analysis of Diverse Ric... **F** literatureReference

Reactions:

We have provision in our data schema to accommodate Details

- Cell type
- Protein modifications
- Disease models and pathogens





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Plail Diviniormatics Resource workshop on 22 June 2024, Plail Diviogy 2024 meeting at nonoiulu nawaii

Exploring Pathways in Pathway Browser

✓ Oryza sativa Pathways Aegilops tauschii Amborella trichopoda Event Hierarchy: Arabidopsis lyrata Arabidopsis thaliana 😑 🍝 Hormone signaling, tr Arachis duranensis Arachis ipaensis 🗄 🏪 IAA biosynthesis I Beta vulgaris 🗉 🕌 IAA biosynthesis II Brachypodium distachyon Brassica napus Brassica oleracea IAA conjugate biosy Brassica rapa Cajanus cajan Fassinosteroid bio Capsicum annuum 🕀 🕌 Trans-zeatin biosyr Chlamydomonas reinhardtii Chondrus crispus 🕀 🕌 Cytokinins 7-N-gluc Cicer arietinum 🗄 🍹 Cytokinins 9-N-gluc Citrus sinensis Coffea canephora 🗄 🏝 Cytokinins-O-gluco Corchorus capsularis Jasmonic acid bios Cucumis sativus Cyanidioschyzon merolae 🗄 🕌 Jasmonic acid sign Daucus carota 🗉 🕌 GA12 biosynthesis Dioscorea rotundata Erythranthe guttata 🕀 🕌 Gibberellin biosynth Eucalyptus grandis 🗉 🕌 Gibberellin biosynth Fragaria vesca Galdieria sulphuraria 🗄 🕌 Gibberellin biosynth Glycine max 🗉 🖫 ABA biosynthesis a Gossypium raimondii Helianthus annuus Abscisic acid home Hordeum vulgare Ethylene biosynthes Jatropha curcas Leersia perrieri Krigolactone biosyl Lupinus angustifolius 🗉 🕌 Strigolactone signa Malus domestica Manihot esculenta Brassinosteroid sign Medicago truncatula 🕀 🕌 Gibberellin signalin Musa acuminata Nicotiana attenuata 🕀 🎽 Salicylic acid signa Orvza australiensis 🕀 🕌 Auxin signalling Oryza barthii

Summation

Secondary metabolism
https://plantreactome.gramene.org

🕀 🕌 Auxin transport



Plant

Reactome

Gramene Pathway

Salicylic acid (SA) is a small phenolic compound that could act as a phytohomone and/or a signaling molecule that effect plant immunity, specifically Systemic Acquire Resistance (SAR). Constitutive

Pathway Projections in Arabidopsis



Displaying Gene Interaction Data on Plant Reactome Pathways

Synthesizing gene-gene interactions information



Users can also upload their own data

Users can send us the data from their favorite species







ne

Gramene Pathway

Analysis Tools: Species Comparisons





Analysis Tools

Users can upload their own omics data and download analysis results



LIMITATIONS for Pathway Biocuration



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16

The annotation status of the Arabidopsis proteome as of 2024



Genetics, Volume 227, Issue 1, May 2024, iyae027, https://doi.org/10.1093/genetics/iyae027

The annotation status of the Rice proteome as of 2021



Research | Open access | Published: 19 April 2022

New Data and New Features of the FunRiceGenes (Functionally Characterized Rice Genes) Database: 2021 Update

<u>Fangfang Huang, Yingru Jiang, Tiantian Chen, Haoran Li, Mengjia Fu, Yazhou Wang, Yufang Xu, Yang Li,</u> <u>Zhengfu Zhou, Lihua Jia, Yidan Ouyang & Wen Yao</u>

<u>*Rice*</u> **15**, Article number: 23 (2022) Cite this article

4125 Accesses | 12 Citations | 2 Altmetric | Metrics

Up to Nov 2021

~ 4100 functionally characterized rice genes

Rice genome Draft genome in 2002 and high-quality 2005

~<u>39,045 protein-coding</u> genes

- ~ 6000 members of different gene families
- 22.3% of the 39,045 annotated protein-coding genes in the rice genome.



Can we use existing knowledge from other species to curate pathways in the Plant Reactome?



Use of gene-orthology relationships to improve functional annotation of a rice gene

Barley EGT2 is a Transcriptional regulator of root growth angle



Gene-orthology relationships to predict potential functions of a rice gene

PNAS RESEARCH ARTICLE PLANT BIOLOGY

antigravitropic mechanism

Root angle is controlled by EGTI in cereal crops employing an

crops represents a key trait for efficient capture of soil resources. Root nined by competing gravitropic versus antigravitropic offset (AGO) mechwe report a root angle regulatory gene termed ENHANCED GRAVI-GCTU that senders a nutrition AGO component who is the set

Servers Marguel¹⁴ (B. Houge Lou^{14,4}), Guorge Graggel², Biocolds Donell, Jacob F. Neme¹⁴, B. Alt N. Noeth¹, I. Charlin Friezand¹⁵, C. Stato March², Jees L. Laffe⁴, R. Neeb Laff⁴, B. Neeb Laff⁴

OPEN ACCESS

egt1 mutant has a steep root growth angle phenotype

hypergravitropic root phenotype

EGT1 appears to function as a component of an antigravitropic offset mechanism that regulates tissue stiffness, which impacts final root growth angle.



Gene-gene interaction network based on evidence from rice and other species



> Proc Natl Acad Sci U S A. 2014 Jul 15;111(28):10377-82. doi: 10.1073/pnas.1401680111. Epub 2014 Jun 30.

Rice actin-binding protein RMD is a key link in the auxin-actin regulatory loop that controls cell growth

Gang Li ¹, Wanqi Liang ¹, Xiaoqing Zhang ¹, Haiyun Ren ², Jianping Hu ³, Malcolm J Bennett ⁴, Dabing Zhang ⁶ Affiliations + expand

PMID: 24982173 PMCID: PMC4104909 DOI: 10.1073/pnas.1401680111

Abstract

The plant hormone auxin plays a central role in plant growth and development. Auxin transport and signaling depend on actin organization. Despite its functional importance, the mechanistic link between actin filaments (F-actin) and auxin intracellular signaling remains unclear. Here, we report that the actin-organizing protein Rice Morphology Determinant (RMD), a type II formin from rice (Oryza sativa), provides a key link. Mutants lacking RMD display abnormal cell growth and altered configuration of F-actin array direction. The rmd mutants also exhibit an inhibition of auxinmediated cell elongation, decreased polar auxin transport, altered auxin distribution gradients in root tips, and suppression of plasma membrane localization of auxin transporters O. sativa PIN-FORMED to (OsPIN1b) and OsPIN2 in root cells. We demonstrate that RMD is required for endocytosis; acosytosis, and auxin-mediated OsPIN2 recycling to the plasma membrane. Moreover, RMD expression is directly regulated by heterodimerized O. sativa auxin response factor 23 (OsARF23) and OsARF24, providing evidence that auxin modulates the orientation of F-actin arrays through RMD. In support of this regulatory loop, osarf23 and lines with reduced expression



Re-use and Re-analysis of Transcriptome Data



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23

Biocuration of rice transcription factor network involved in stress response



Open Access Article

Biocuration of a Transcription Factors Network Involved in Submergence Tolerance during Seed Germination and Coleoptile Elongation in Rice (*Oryza sativa*)

by ∰ Sushma Naithani ^{1,*} ⊠ ⁽⁰⁾, ⊗ Bijayalaxmi Mohanty ² ⊠ ⁽⁰⁾, ⊗ Justin Elser ¹ ⊠ ⁽⁰⁾, ⊗ Peter D'Eustachio ³ ⊠ ⁽⁰⁾ and ⊗ Pankaj Jaiswal ¹ ⊠ ⁽⁰⁾

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- * Author to whom correspondence should be addressed.

Plants 2023, 12(11), 2146; https://doi.org/10.3390/plants12112146









Needed: Increased Support for Biocuration





27

Our Undergraduate Biocurators : Plant Gene and Pathway Curation



Eli Waddell 2013-2014



Christina Partipilo 2014-2016



Ami (Amitavo) Ganguli URSA-Fellow, 2021 winter-spring



Sarah El-Husseini URSA-Fellow, 2021 winter-spring Honors Thesis (2021-2022)



Shayla Rao (spring 2020)

Quentin Beers URSA-Fellow, 2021 winter-spring



Callan Stowell (winter-spring 2020) URSA-fellow



Olivia Worley URSA-Fellow, 2021 winter-spring



Maddie Goebel URSA-Fellow, 2021 winter-spring

chna

Gwendolyn Michna ring URSA-Fellow, 2021 winter-spring



Kennedy Sparks URSA-Fellow, 2022 winter-spring

Tutorials and Videos on Gramene YouTube Channel tinyurl.com/grchannel



Elizabeth Hooks URSA-Fellow, winter-spring

2022, continued until Fall

2023

We train undergraduate students in critical review of scientific literature and gene biocuration



Plant Bioinformatics Resource Workshop on 22 June 2024, Plant Biology 2024 meeting at Honolulu Hawaii

Plant

URSA undergraduate Research Fellows of 2024 working on Biocuration of genes and pathways



Siena Buchanan second-year biology major

Isabella Ramirez second-year microbiology major

Carolyn Crofford Bioresource Research and Sustainability major



30

Biocuration of a gene family using transcriptome data to infer tissue specific expression and association with stress response



Daemon A. Dikeman (2019-2020) (Honors Thesis, undergraduate) Joined Ph.D. at Univ. of Maine > PeerJ. 2021 Mar 15;9:e11052. doi: 10.7717/peerj.11052. eCollection 2021.

Beyond gene ontology (GO): using biocuration approach to improve the gene nomenclature and functional annotation of rice S-domain kinase subfamily

Sushma Naithani ¹, Daemon Dikeman ¹, Priyanka Garg ¹, Noor Al-Bader ¹, Pankaj Jaiswal ¹

Affiliations + expand PMID: 33777532 PMCID: PMC7971086 DOI: 10.7717/peerj.11052 Free PMC article

Abstract

The S-domain subfamily of receptor-like kinases (SDRLKs) in plants is poorly characterized. Most members of this subfamily are currently assigned gene function based on the S-locus Receptor Kinase from Brassica that acts as the female determinant of self-incompatibility (SI). However, Brassica like SI mechanisms does not exist in most plants. Thus, automated Gene Ontology (GO) pipelines are not sufficient for functional annotation of SDRLK subfamily members and lead to erroneous association with the GO biological process of SI. Here, we show that manual bio-curation can help to correct and improve the gene annotations and association with relevant biological processes. Using publicly available genomic and transcriptome datasets, we conducted a detailed



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EMBL-EBI

NYU Langone Medical Center MEDICAL CENTER •Peter D'Eustachio (Biocuration)



Oregon State

Undergraduate Research, Scholarship and the Arts engage program, OSU



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European Bioinformatics Institute

- Irene Papatheodorou (ATLAS)
- IntAct



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