

Target Gene Notebook

User Guide
1.2

Target Gene Notebook allows the curation of persistent, editable, and distributable databases of genomic, biological, chemical, and other data that pertain to the local neighborhood around a gene of interest.

The current version draws heavily on the many harmonized data sources available through Ensembl¹.

1. Daniel R. Zerbino, Premanand Achuthan, Wasiu Akanni, M. Ridwan Amode, Daniel Barrell, Jyothish Bhai, Konstantinos Billis, Carla Cummins, Astrid Gall, Carlos García Giro'n, Laurent Gil, Leo Gordon, Leanne Haggerty, Erin Haskell, Thibaut Hourlier, Osagie G. Izuogu, Sophie H. Janacek, Thomas Juettemann, Jimmy Kiang To, Matthew R. Laird, Ilias Lavidas, Zhicheng Liu, Jane E. Loveland, Thomas Maurel, William McLaren, Benjamin Moore, Jonathan Mudge, Daniel N. Murphy, Victoria Newman, Michael Nuhn, Denye Ogeh, Chuang Kee Ong, Anne Parker, Mateus Patricio, Harpreet Singh Riat, Helen Schuilenburg, Dan Sheppard, Helen Sparrow, Kieron Taylor, Anja Thormann, Alessandro Vullo, Brandon Walts, Amonida Zadissa, Adam Frankish, Sarah E. Hunt, Myrto Kostadima, Nicholas Langridge, Fergal J. Martin, Matthieu Muffato, Emily Perry, Magali Ruffier, Dan M. Staines, Stephen J. Trevanion, Bronwen L. Aken, Fiona Cunningham, Andrew Yates, Paul Flicek
Ensembl 2018.
PubMed PMID: 29155950.

Target Gene Notebook was originally supported and created at Eisai Inc.'s Andover Innovative Medicine Institute in Andover, MA.

We encourage further development of this resource.

Target Gene Notebook team

Eisai (design, implementation and testing):

Andrew Kirby

Janna Hutz

Jamey Wierzbowski

Mary Pat Reeve

Massachusetts General Hospital (design input):

Mark Daly

Hailiang Huang

Requirements

Server:
Java >= 1.7

Client:
Google Chrome
JavaScript enabled

Target Gene Notebook uses these frameworks/libraries:

Bootstrap 3.3.7 (MIT license)

FileSaver.js 1.3.2 (MIT license)

Huebee 2.0.0 (MIT license)

jQuery 2.2.4 (MIT license)

jQuery UI 1.12.1 (MIT license)

Lightbox2 2.9.0 (MIT license)

Select2 4.0.3 (MIT license)

tablesorter 2.28.14 (MIT license)

Spark micro framework (sparkjava.com, Apache 2 license)

Dependencies:

commons-io 2.4

commons-lang3 3.4

gson 2.7

jsoup 1.8.1

slf4j-simple 1.7.21

sqlite-jdbc 3.7.2

GUI Overview

Target Gene Notebook 1.1:1.1 User Guide About

Marker	r2	Coding
rs12111485	0.922	
rs1990760		Y
rs10555400	0.728	
rs13023390	0.674	
rs7587426	0.616	

View Mode * Credible Sets | LD Summary (0.6 r2 threshold, 1KGp3-EUR) Hide non-coding genes

Summary (IFIH1 TSS Ensembl 93 chr:2:162,318,703)

- Association Results
- Association Details
- pQTL Results
- pQTL Details
- eQTL Results
- Variants of Interest

LD Summary	Name	Non-Reference Allele	Location	Gene	Protein Change	Polyphen	f(FE)	f(Max)
<input checked="" type="checkbox"/>	rs141572054	T	162,359,145	GGA	p.Arg186Ter		0.002	0.012
<input checked="" type="checkbox"/>	rs146161584	G	162,359,091	GGA	p.Ser168Gly	possibly damaging	0.000	0.010
<input type="checkbox"/>	rs759197923	T	162,352,402	GGA	p.Tyr607Ile	probably damaging	0.000	0.001
<input checked="" type="checkbox"/>	rs17783344	G	162,352,383	GGA	p.Ser80Ala	benign	0.126	0.209
<input type="checkbox"/>	rs754396914	C	162,352,371	GGA	p.Cys76Arg	probably damaging	0.000	0.001
<input type="checkbox"/>	rs79545841	T	162,352,367	GGA	p.Gln64ArgHis	probably damaging	0.004	0.004
<input type="checkbox"/>	rs545281061	-	162,347,739	GGA	p.Gln64ArgHisTer14		0.000	0.004
<input type="checkbox"/>	rs774170639	A	162,347,656	GGA	p.Asp36Asn	benign	0.000	0.001
<input type="checkbox"/>	rs1403164634	G	162,318,102	IFIH1	p.Gly69Ala	benign	0.000	0.001
<input type="checkbox"/>	rs147278787	A	162,318,079	IFIH1	p.Arg77Trp	possibly damaging	0.001	0.003

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- Functional Variants Details
- Expression Data Details
- Protein and Structure Chemistry Details
- Clinical Results
- Clinical Data Details
- References (18 out of 23 not yet reviewed)

Reviewed	1st Author	Year	Journal/Site/File	Title/Description	Curator Comment	
<input checked="" type="checkbox"/>		2019	https://clic.io/repurposing-app	Curated information about drugs and drug targets from Broad.	Enter comment	Show/Add Details X
<input checked="" type="checkbox"/>	Li Q	2017	Pharmacogenet. Genomics	Genome-wide association study of paliperidone efficacy.	Enter comment	Show/Add Details
<input checked="" type="checkbox"/>		2017	http://www.nslablab.io/blog/2017/7/19/rapid-growth-of-the-uk-biobank-phenotypes-for-337000-samples-in-the-uk-biobank	UK Biobank Rapid GWAS	Enter comment	Show/Add Details X
<input checked="" type="checkbox"/>	Pettersson M	2017	Am. J. Med. Genet. A	Further evidence for specific IFIH1 mutation as a cause of Singleton-Merten syndrome with phenotypic heterogeneity.	Enter comment	Show/Add Details X
<input checked="" type="checkbox"/>	Gorman JA	2017	Nat. Immunol.	The A946T variant of the RNA sensor IFIH1 mediates an interferon program that limits viral infection but increases the risk for autoimmunity.	Enter comment	Show/Add Details (1) X
<input type="checkbox"/>	Bronson PG	2016	Nat. Genet.	Common variants at PVT1, ATG13, AMBRA1, AH1 and CLEC16A are associated with selective IgA deficiency.	Enter comment	Show/Add Details
<input type="checkbox"/>	Morris DL	2016	Nat. Genet.	Genome-wide association meta-analysis in Chinese and European individuals identifies ten new loci associated with systemic lupus erythematosus.	Enter comment	Show/Add Details
<input type="checkbox"/>	Jin Y	2016	Nat. Genet.	Genome-wide association studies of autoimmune vitiligo identify 23 new risk loci and highlight key pathways and regulatory variants.	Enter comment	Show/Add Details
<input type="checkbox"/>	Bentham J	2015	Nat. Genet.	Genetic association analysis implicates aberrant regulation of innate and adaptive immunity genes in the pathogenesis of systemic lupus erythematosus.	Enter comment	Show/Add Details
<input type="checkbox"/>	Yin X	2015	Nat Commun	Genome-wide meta-analysis identifies multiple novel associations and ethnic heterogeneity of psoriasis susceptibility.	Enter comment	Show/Add Details

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Add PubMed Reference Add File Reference Add Web Reference Add bioRxiv Reference

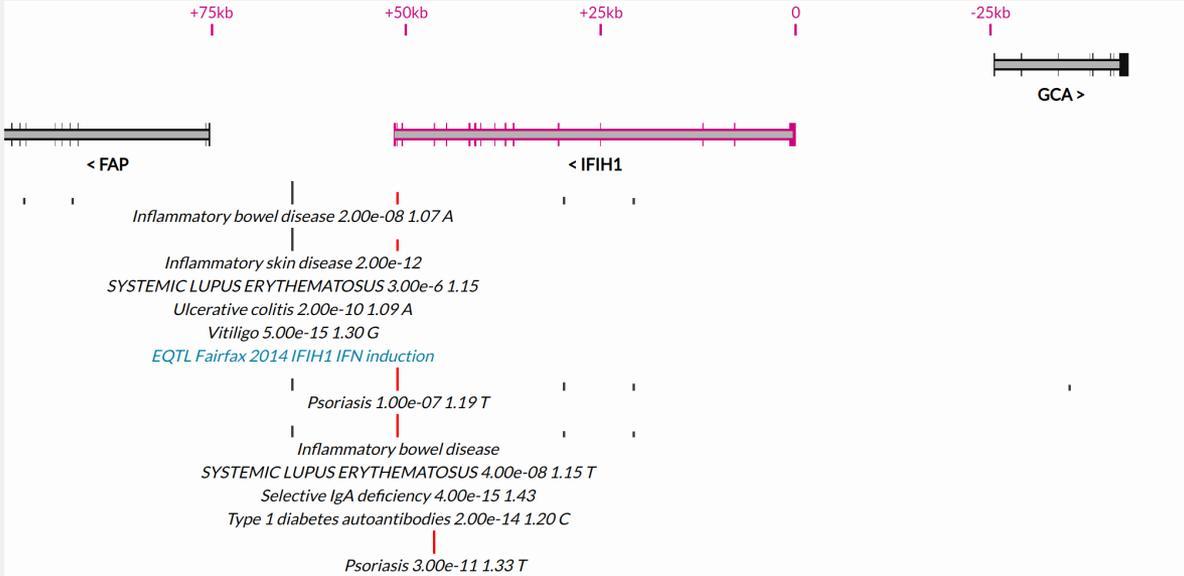
Interactive graphical view of curated data within a genomic context.

Text-based/tabular view of curated data.

Interactive graphical view of curated data within a genomic context...

Interactive
graphical view of
curated data within
a genomic context.

Overview



Psoriasis 1.00e-07 1.19 T

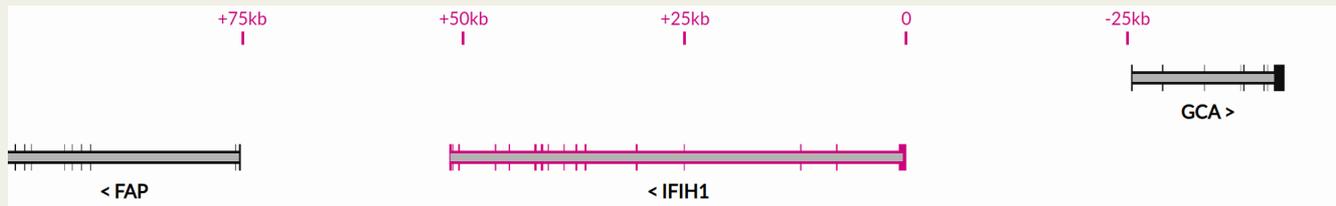
Credible Set: LD 1000GENOMES:phase_3:AMR

Marker	r2	Coding
rs2111485	0.922	
rs1990760		Y
rs10555400	0.734	
rs13023380	0.674	
rs7587426	0.616	

View Mode Credible Sets LD Summary (0.6 r2 threshold, 1KGp3:EUR)

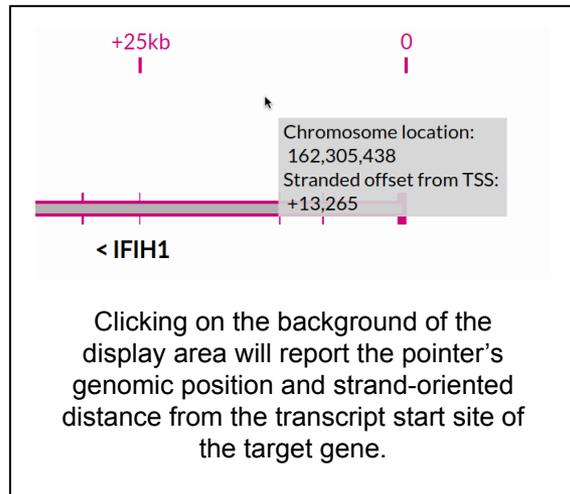
Hide non-coding genes

Interactive graphical view of curated data within a genomic context.

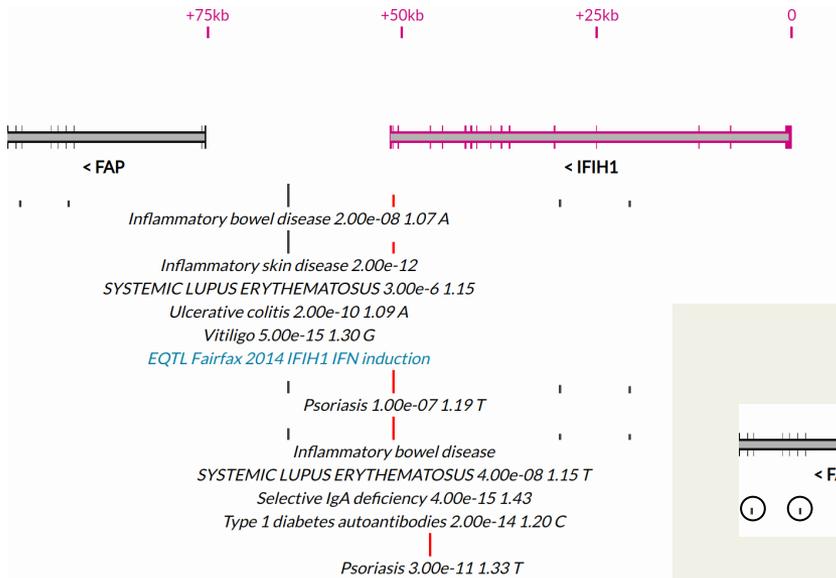


The display area shows a 50-kb neighborhood around the Ensembl canonical transcript for the target gene.

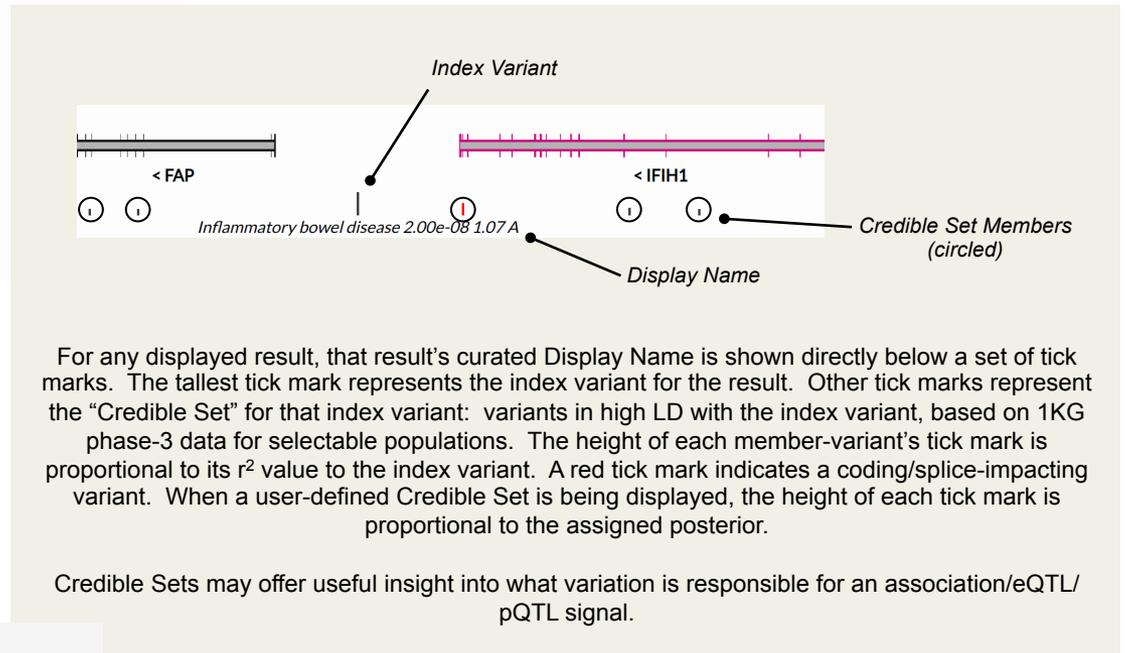
The genomic ruler is oriented relative to the start of the target-gene transcript.



Interactive graphical view of curated data within a genomic context.



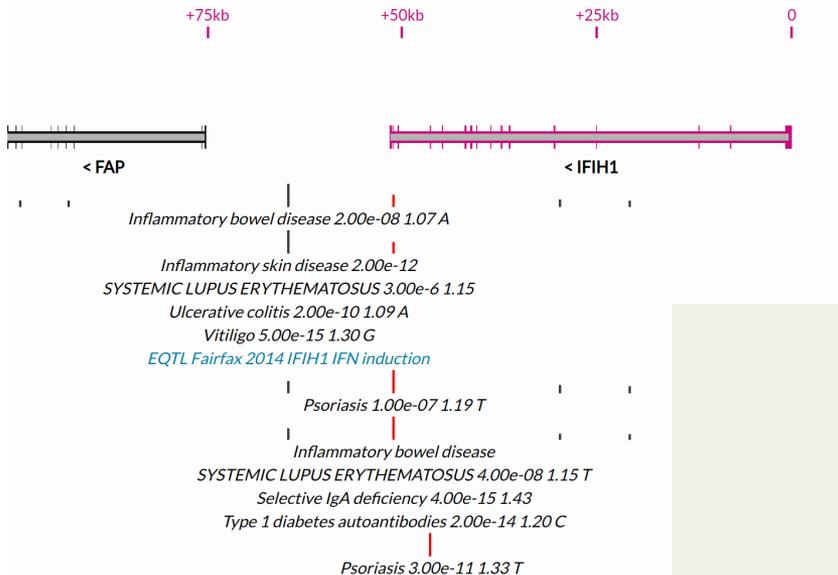
Curated association/eQTL/pQTL results are shown below any gene annotations.



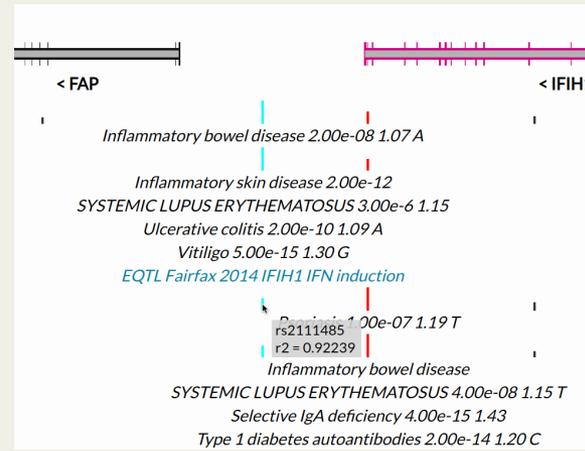
Inflammatory skin disease 2.00e-12
 SYSTEMIC LUPUS ERYTHEMATOSUS 3.00e-6 1.15
 Ulcerative colitis 2.00e-10 1.09 A
 Vitiligo 5.00e-15 1.30 G
 EQTL Fairfax 2014 IFIH1 IFN induction

All results with the same index variant and Credible Set share a single set of tick marks.

Interactive graphical view of curated data within a genomic context.



Curated association/eQTL/pQTL results are shown below along gene annotations.



Hovering the cursor over a variant's tick mark will display its name and its r^2 value to the index variant. Additionally, the same variant will be shaded teal where it appears in other association/eQTL/pQTL results.

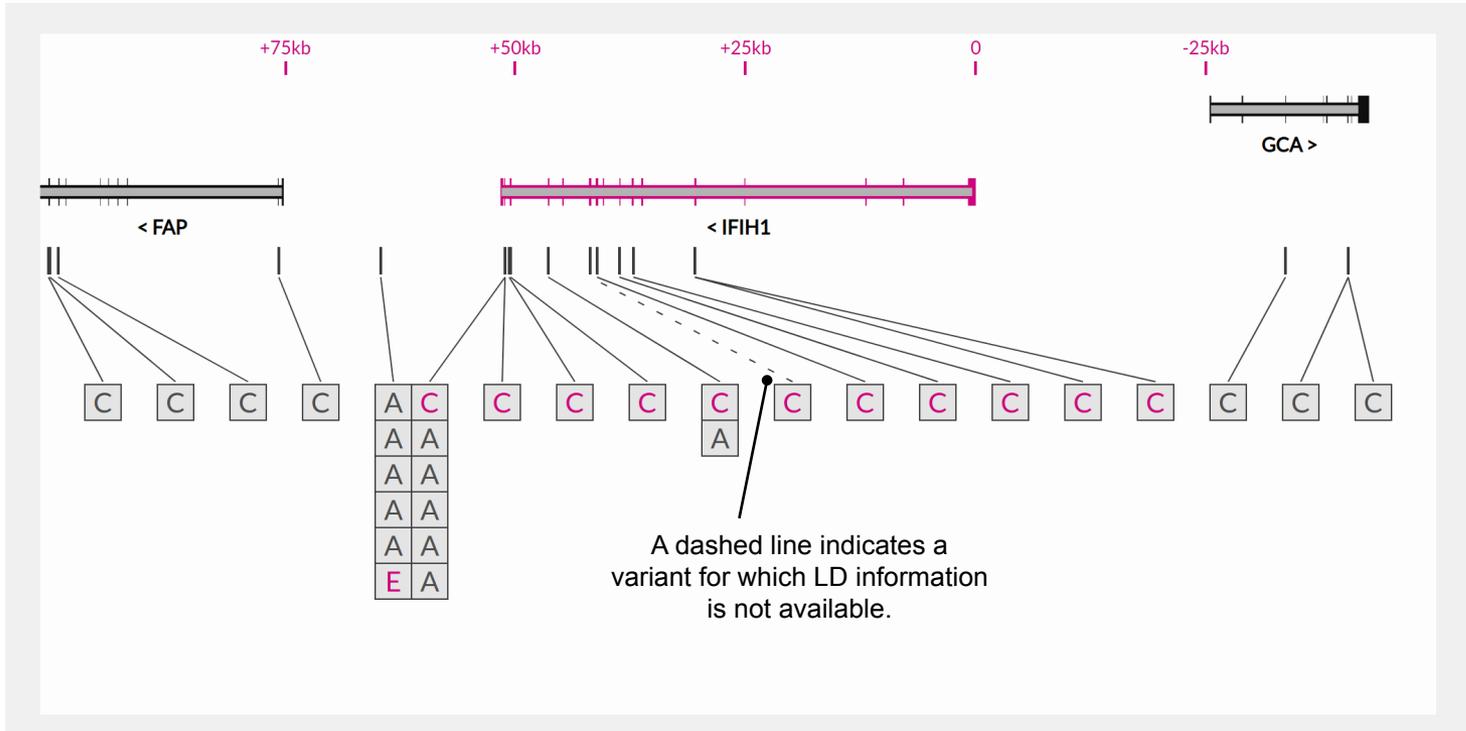
Psoriasis 1.00e-07 1.19 T
Credible Set: LD 1000GENOMES:phase_3:AMR

Marker	r2	Coding
rs2111485	0.922	
rs1990760		Y
rs10555400	0.734	
rs13023380	0.674	
rs7587426	0.616	

Clicking on the Display Name for an association/eQTL/pQTL result will reveal a dismissable table just below the display area summarizing the index variant and its Credible Set.

Interactive graphical view of curated data within a genomic context.

LD-summary graphical perspective



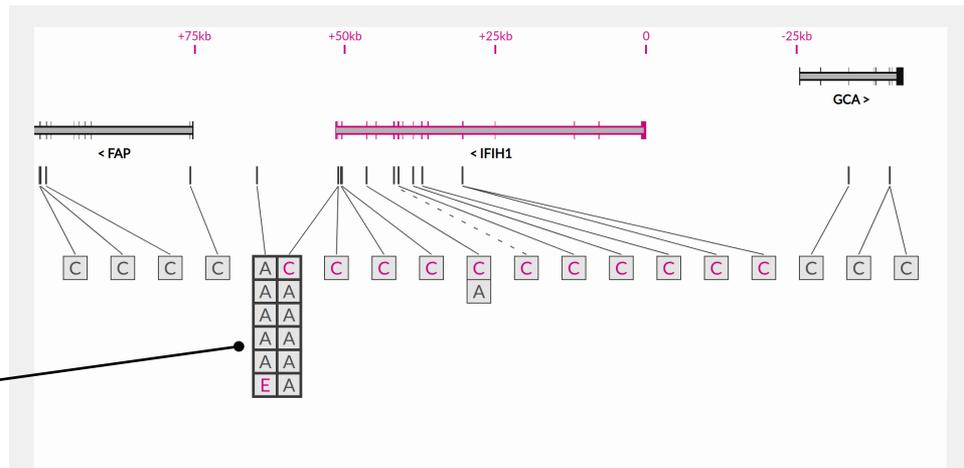
Each box represents a particular association result (A), allele coding impact (C), eQTL result (E), or pQTL result (P). Results for the same variant are stacked vertically.

Variants are placed into groups where all members have $r^2 \geq 0.6$ (1KGp3:EUR) to at least one other member of the group.

eQTL results and allele coding impacts assigned to the target gene are highlighted in magenta.

Interactive graphical view of curated data within a genomic context.

LD-summary graphical perspective



Clicking on an LD group will reveal a dismissable table below the display area summarizing the events in that group.

Marker	Finding	Allele	Gene	Coding Impact	Phenotype	Tissue	Freq	OR/Beta
rs2111485	Assoc.				Systemic lupus erythematosus		0.398 sum(MAF) 1KGp3:EUR	1.15
rs2111485	Assoc.	A			Ulcerative colitis		0.398 sum(MAF) 1KGp3:EUR	1.09
rs2111485	Assoc.				Inflammatory skin disease		0.398 sum(MAF) 1KGp3:EUR	
rs2111485	Assoc.	G			Vitiligo		0.398 sum(MAF) 1KGp3:EUR	1.30
rs2111485	Assoc.	A			Inflammatory bowel disease		0.398 sum(MAF) 1KGp3:EUR	1.07
rs2111485	eQTL		IFIH1			Monocytes	0.398 sum(MAF) 1KGp3:EUR	5.85
rs1990760	C. Variant	T	IFIH1	p.Ala946Thr			0.459 ExAC:ALL	
rs1990760	Assoc.				Selective IgA deficiency		0.395 sum(MAF) 1KGp3:EUR	1.43
rs1990760	Assoc.	T			Systemic lupus erythematosus		0.395 sum(MAF) 1KGp3:EUR	1.15
rs1990760	Assoc.				Inflammatory bowel disease		0.395 sum(MAF) 1KGp3:EUR	
rs1990760	Assoc.	T			Psoriasis		0.395 sum(MAF) 1KGp3:EUR	1.19
rs1990760	Assoc.	C			Type 1 diabetes autoantibodies		0.395 sum(MAF) 1KGp3:EUR	1.20

Interactive graphical view of curated data within a genomic context.

Text-based/tabular view of curated data...

Text-based/tabular
view of curated
data.

Overview

Summary (IFIH1 TSS Ensembl 93:chr2:162,318,703)

- Association Results
- Association Details
- pQTL Results
- pQTL Details
- eQTL Results
- Variants of Interest

LD Summary	Name	Non-Reference Allele	Location	Gene	Protein Change	Polyphen	f(NFE)	f(Max)
<input checked="" type="checkbox"/>	rs141572054	T	162,359,145	GCA	p.Arg186Ter		0.002	0.012
<input checked="" type="checkbox"/>	rs146161584	G	162,359,091	GCA	p.Ser168Gly	possibly damaging	0.000	0.010
<input type="checkbox"/>	rs758193923	T	162,352,402	CCA	p.Tyr86Phe	probably damaging	0.000	0.001
<input checked="" type="checkbox"/>	rs17783344	G	162,352,383	GCA	p.Ser80Ala	benign	0.126	0.209
<input type="checkbox"/>	rs754396914	C	162,352,371	GCA	p.Cys76Arg	probably damaging	0.000	0.001
<input type="checkbox"/>	rs79565841	T	162,352,367	GCA	p.Gln74His	probably damaging	0.004	0.004
<input type="checkbox"/>	rs545281061	-	162,347,739	GCA	p.Gln64ArgfsTer14		0.000	0.004
<input type="checkbox"/>	rs774170639	A	162,347,656	GCA	p.Asp36Asn	benign	0.000	0.001
<input type="checkbox"/>	rs1403164634	G	162,318,102	IFIH1	p.Gly69Ala	benign	0.000	0.001
<input type="checkbox"/>	rs147278787	A	162,318,079	IFIH1	p.Arg77Trp	possibly damaging	0.001	0.003

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- Functional Variants Details
- Expression Data Details
- Protein and Structure Chemistry Details
- Clinical Results
- Clinical Data Details
- References (18 out of 23 not yet reviewed)

Reviewed	1st Author	Year	Journal/Site/File	Title/Description	Curator Comment	
<input checked="" type="checkbox"/>		2019	https://clue.io/repurposing-app	Curated information about drugs and drug targets from Broad.	Enter comment	Show/Add Details X
<input checked="" type="checkbox"/>	Li Q	2017	Pharmacogenet. Genomics	Genome-wide association study of palliperidone efficacy.	Enter comment	Show/Add Details
<input checked="" type="checkbox"/>		2017	http://www.nealelab.is/blog/2017/7/19/rapid-gwas-of-thousands-of-phenotypes-for-337000-samples-in-the-uk-biobank	UK Biobank Rapid GWAS	Enter comment	Show/Add Details X
<input checked="" type="checkbox"/>	Petterson M	2017	Am. J. Med. Genet. A	Further evidence for specific IFIH1 mutation as a cause of Singleton-Merten syndrome with phenotypic heterogeneity.	Enter comment	Show/Add Details X
<input checked="" type="checkbox"/>	Gorman JA	2017	Nat. Immunol.	The A946T variant of the RNA sensor IFIH1 mediates an interferon program that limits viral infection but increases the risk for autoimmunity.	Enter comment	Show/Add Details (1) X
<input type="checkbox"/>	Bronson PG	2016	Nat. Genet.	Common variants at PVT1, ATG13-AMBRA1, AH11 and CLEC16A are associated with selective IgA deficiency.	Enter comment	Show/Add Details
<input type="checkbox"/>	Morris DL	2016	Nat. Genet.	Genome-wide association meta-analysis in Chinese and European individuals identifies ten new loci associated with systemic lupus erythematosus.	Enter comment	Show/Add Details
<input type="checkbox"/>	Jin Y	2016	Nat. Genet.	Genome-wide association studies of autoimmune vitiligo identify 23 new risk loci and highlight key pathways and regulatory variants.	Enter comment	Show/Add Details
<input type="checkbox"/>	Bentham J	2015	Nat. Genet.	Genetic association analyses implicate aberrant regulation of innate and adaptive immunity genes in the pathogenesis of systemic lupus erythematosus.	Enter comment	Show/Add Details
<input type="checkbox"/>	Yin X	2015	Nat Commun	Genome-wide meta-analysis identifies multiple novel associations and ethnic heterogeneity of psoriasis susceptibility.	Enter comment	Show/Add Details

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[Add Pubmed Reference](#)
[Add File Reference](#)
[Add Web Reference](#)
[Add bioRxiv Reference](#)

Text-based/tabular view of curated data.

13 different sections hold curated data

- ▶ Summary (IFIH1 TSS Ensembl 93:chr2:162,318,703)
- ▶ Association Results
- ▶ Association Details
- ▶ pQTL Results
- ▶ pQTL Details
- ▶ eQTL Results
- ▶ Variants of Interest
- ▶ Functional Variants Details
- ▶ Expression Data Details
- ▶ Protein and Structure Chemistry Details
- ▶ Clinical Results
- ▶ Clinical Data Details
- ▶ References (18 out of 23 not yet reviewed)

Text-based/tabular
view of curated
data.

Summary section

The Ensembl version against which the data were curated is shown here, along with the start position of the canonical transcript for the target gene.

▼ Summary (IFIH1 TSS Ensembl 93:chr2:162,318,703)

Enter summary

Text additions/changes in this area are automatically saved to the target-gene database.

Text-based/tabular
view of curated
data.

Association Results section

Display Name	Phenotype	Source	Year	Index Variant	Allele	Pvalue	OR/Beta	Credible Set	Show	Curator Comment
Inflammatory bowel disease	Inflammatory bowel disease	Liu JZ; Association analyses identify 38 susceptibility loci for inflammatory bowel disease and highlight shared genetic risk across populations.	2015	rs1990760		4.00e-10		LD 1KGp3:EUR (3)	<input checked="" type="checkbox"/>	Enter comment
Inflammatory bowel disease 2.00e-08 1.07 A	Inflammatory bowel disease	Jostins L; Host-microbe interactions have shaped the genetic architecture of inflammatory bowel disease.	2012	rs2111485	A	2.00e-08	1.07	LD 1KGp3:AMR (5)	<input checked="" type="checkbox"/>	Enter comment
Inflammatory skin disease 2.00e-12	Inflammatory skin disease	Baurecht H; Genome-wide comparative analysis of atopic dermatitis and psoriasis gives	2015	rs2111485		2.00e-12		LD 1KGp3:EUR (1)	<input checked="" type="checkbox"/>	Enter comment

Editable Display Name used to label the result in the display area. Changes are automatically saved to the target-gene database.

Alleles with an "Audited" label are those for which strandedness is believed to be expressed as genome-forward-strand, based on data provided.

Dropdown menu to select the applicable Credible Set. The menu can also be used to create a custom Credible Set.

- LD 1KGp3:EUR (1)
- Unset
- None
- LD 1KGp3:AFR (1)
- LD 1KGp3:AMR (5)
- LD 1KGp3:EAS (3)
- LD 1KGp3:EUR (1)
- LD 1KGp3:SAS (0)
- Add custom

Toggle to indicate if a result should be shown in the display area (Credible Set must not be "Unset" for a result to be shown).

Editable curator comment. Changes are automatically saved to the target-gene database.

Marder score) 3.00e-06 5.87 T	schizophrenia (negative Marder score)									
Selective IgA deficiency 7.0e-10 1.49	Immunoglobulin A	Ferreira RC; Association of IFIH1 and other autoimmunity risk alleles with selective IgA deficiency.	2010	rs1990760		7.00e-10	1.49	LD 1KGp3:AMR (4)	<input type="checkbox"/>	Enter comment

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Add Association

Button to add a new association result (see next page).

Text-based/tabular view of curated data.

Association Results section

Adding an association result

Desired reference must already have been added to the References section to appear in the Source dropdown menu (see later description of the References section).

Add Association



New Association

Phenotype:

Source:

Index variant:

Allele (optional):

Pvalue:

OR/Beta:

Enter comment
Comment:

Save

highlights the role of innate immunity.



Some new phenotype 1.00e-09 1.19 T	Some new phenotype	Curated information about drugs and drug targets from Broad.	2019	rs1990760	T	1.00e-09	1.19	Unset	<input type="checkbox"/>	This is a made-up example.	X
---------------------------------------	--------------------	---	------	-----------	---	----------	------	-------	--------------------------	----------------------------	---

New row added to association-results table.

Button to remove association result.

Text-based/tabular view of curated data.

Association Results section

Creating a custom Credible Set

Display Name	Phenotype	Source	Year	Index Variant	Allele	Pvalue	OR/Beta	Credible Set	Show	Curator Comment
Inflammatory bowel disease	Inflammatory bowel disease	Liu JZ; Association analyses identify 38 susceptibility loci for inflammatory bowel disease and highlight shared genetic risk across populations.	2015	rs1990760		4.00e-10		LD 1KGp3:EUR (3)	<input checked="" type="checkbox"/>	Enter comment
Inflammatory bowel disease 2.00e-08 1.07 A	Inflammatory bowel disease	Jostins L; Host-microbe interactions have shaped the genetic architecture of inflammatory bowel disease.	2012	rs2111485	A	2.00e-08	1.07	LD 1KGp3:AMR (5)	<input checked="" type="checkbox"/>	Enter comment
Inflammatory skin disease 2.00e-12	Inflammatory skin disease	Baurecht H; Genome-wide comparative analysis of atopic dermatitis and psoriasis gives	2015	rs2111485		2.00e-12		LD 1KGp3:EUR (1)	<input checked="" type="checkbox"/>	Enter comment

Custom Credible Set

Name of credible set:

Index-variant (rs2111485) posterior:

Other credible-set members:

Marker	Posterior
rs1990760	0.11
Eg. rs123	Eg. 0.123

Add Row

Define custom Credible Set.

Select "Add custom" dropdown option.

- LD 1KGp3:EUR (1)
- Unset
- None
- LD 1KGp3:AFR (1)
- LD 1KGp3:AMR (5)
- LD 1KGp3:EAS (3)
- LD 1KGp3:EUR (1)
- LD 1KGp3:SAS (0)
- Add custom

- LD 1KGp3:EUR (1)
- Unset
- None
- LD 1KGp3:AFR (1)
- LD 1KGp3:AMR (5)
- LD 1KGp3:EAS (3)
- LD 1KGp3:EUR (1)
- LD 1KGp3:SAS (0)
- MyCredibleSet (1)
- Delete custom

The custom Credible Set and ability to delete the custom Credible Set are now selectable options in the dropdown menu.

Text-based/tabular view of curated data.

pQTL Results and eQTL Results sections

These sections are very similar to the Association Results section

pQTL Results													
eQTL Results													
Display Name	Tissue	Gene	Source	Year	Index Variant	Pvalue	Beta	Effect Allele	Credible Set	Show	Curator Comment		
EQTL GTeX FAP Nerve Tibial 2.55e-05	Nerve Tibial	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs3788967	2.55e-05	0.56		Unset	<input type="checkbox"/>	Tissue pval threshold = 4.11e-05		0
EQTL GTeX FAP Artery Tibial 5.19e-07	Artery Tibial	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs3827491	5.19e-07	0.43		Unset	<input type="checkbox"/>	Tissue pval threshold = 3.54e-05		0
EQTL GTeX FAP Cells Transformed fibroblasts 1.67e-06	Cells Transformed fibroblasts	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs6746339	1.67e-06	0.34		Unset	<input type="checkbox"/>	Tissue pval threshold = 3.54e-05		0
EQTL GTeX FAP Cells Transformed fibroblasts 7.80e-06	Cells Transformed fibroblasts	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs12468578	7.80e-06	0.36		Unset	<input type="checkbox"/>	Tissue pval threshold = 3.54e-05		0
EQTL GTeX FAP Thyroid 2.67e-05	Thyroid	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs2389683	2.67e-05	0.45		Unset	<input type="checkbox"/>	Tissue pval threshold = 3.29e-05		0
EQTL GTeX FAP Thyroid 2.91e-05	Thyroid	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs11455810	2.91e-05	0.44		Unset	<input type="checkbox"/>	Tissue pval threshold = 3.29e-05		0
EQTL GTeX FAP Thyroid 4.53e-06	Thyroid	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs16846600	4.53e-06	0.51		Unset	<input type="checkbox"/>	Tissue pval threshold = 3.29e-05		0
EQTL GTeX FAP Muscle Skeletal 1.58e-05	Muscle Skeletal	FAP	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs35544136	1.58e-05	0.67		Unset	<input type="checkbox"/>	Tissue pval threshold = 1.85e-05		0
EQTL GTeX GCA Adrenal Gland 6.34e-06	Adrenal Gland	GCA	GTEx Consortium; Human genomics. The Genotype-Tissue Expression (GTEx) pilot analysis: multitissue gene regulation in humans.	2015	rs34977319	6.34e-06	-0.77		Unset	<input type="checkbox"/>	Tissue pval threshold = 1.75e-05		0
EQTL Fairfax 2014 IFIH1 IFN induction	Monocytes	IFIH1	Fairfax BP: Innate immune activity conditions the effect of regulatory variants upon monocyte gene expression.	2014	rs2111485	1.19e-08	5.85		LD 1KGp3:EUR (1)	<input checked="" type="checkbox"/>	Enter comment		10

The eQTL Results table contains an extra column showing the count of displayed association and pQTL results that have intersecting Credible-Set members with the 1KGp3:EUR Credible Set of each eQTL result.

Text-based/tabular view of curated data.

Variants of Interest section

In this section are listed those variant alleles which confer a coding or likely splice change to any gene in the neighborhood of the target gene

▼ Variants of Interest								
LD Summary	Name	Non-Reference Allele	Location	Gene	Protein Change	Polyphen	f(NFE)	f(Max)
<input checked="" type="checkbox"/>	rs141572054	T	162,359,145	GCA	p.Arg186Ter		0.002	0.012
<input checked="" type="checkbox"/>	rs146161584	G	162,359,091	GCA	p.Ser168Gly	possibly damaging	0.000	0.010
<input type="checkbox"/>	rs758193923	T	162,352,402	GCA	p.Tyr86Phe	probably damaging	0.000	0.001
<input checked="" type="checkbox"/>	rs17783344	G	162,352,383	GCA	p.Ser80Ala	benign	0.126	0.209
<input type="checkbox"/>	rs754396914	C	162,352,371	GCA	p.Cys76Arg	probably damaging	0.000	0.001
<input type="checkbox"/>	rs79565841	T	162,352,367	GCA	p.Gln74His	probably damaging	0.004	0.004
<input type="checkbox"/>	rs545281061	-	162,347,739	GCA	p.Gln64ArgfsTer14		0.000	0.004
<input type="checkbox"/>	rs774170639	A	162,347,656	GCA	p.Asp36Asn	benign	0.000	0.001
<input type="checkbox"/>	rs1403164634	G	162,318,102	IFIH1	p.Gly69Ala	benign	0.000	0.001
<input type="checkbox"/>	rs147278787	A	162,318,079	IFIH1	p.Arg77Trp	possibly damaging	0.001	0.003

Page: 1 1 to 10 of 68 rows 10

Toggle for inclusion of the variant in the LD-Summary perspective of the display area.

To be included in this table, an allele must have a frequency of at least 0.001 in one of the ExAC sub-populations.

Text-based/tabular view of curated data.

Clinical Results section

▼ Clinical Results

Phenotype/Gene Associations

Phenotype	Gene	Source	Curator Comment
SINGLETON-MERTEN SYNDROME 1	IFIH1	MIM morbid	Enter comment
Aicardi-Goutieres syndrome 7	IFIH1	MIM morbid	Enter comment

Page: 1 1 to 2 of 2 rows 10

Phenotype/Allele Associations

Phenotype	Variant	Unaudited Risk Allele	Clinical Significance	Transcript-Impact Genes	Source
SINGLETON-MERTEN SYNDROME 1	rs6748554	G	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs13418718	A	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs35337543	G	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs35667974	C	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs41399348	T	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs74162087	T	uncertain significance	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs74162089	A	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs79324540	T	uncertain significance	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs140562355	T	benign	IFIH1	ClinVar
SINGLETON-MERTEN SYNDROME 1	rs143870870	C	benign	IFIH1	ClinVar

Page: 1 1 to 10 of 34 rows 10

Editable curator comment. Changes are automatically saved to the target-gene database.

Text-based/tabular view of curated data.

References section

▼ References (18 out of 23 not yet reviewed)

Reviewed	1st Author	Year	Journal/Site/File	Title/Description	Curator Comment		
<input checked="" type="checkbox"/>		2019	https://clue.io/repurposing-app	Curated information about drugs and drug targets from Broad.	Enter comment	Show/Add Details	X
<input checked="" type="checkbox"/>	Li Q	2017	Pharmacogenet. Genomics	Genome-wide association study of paliperidone efficacy.	Enter comment	Show/Add Details	
<input checked="" type="checkbox"/>		2017	http://www.nealelab.is/blog/2017/7/19/rapid-gwas-of-thousands-of-phenotypes-for-337000-samples-in-the-uk-biobank	UK Biobank Rapid GWAS	Enter comment	Show/Add Details	X
<input checked="" type="checkbox"/>	Petterson M	2017	Am. J. Med. Genet. A	Further evidence for specific IFIH1 mutation as a cause of Singleton-Merten syndrome with phenotypic heterogeneity.	Enter comment	Show/Add Details	X
<input checked="" type="checkbox"/>	Gorman JA	2017	Nat. Immunol.	The A946T variant of the RNA sensor IFIH1 mediates an interferon program that limits viral infection but increases the risk for autoimmunity.	Enter comment	Show/Add Details (1)	X
<input type="checkbox"/>	Bronson PG	2016	Nat. Genet.	Common variants at PVT1, ATG13-AMBRA1, AHI1 and CLEC16A are associated with selective IgA deficiency.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Morris DL	2016	Nat. Genet.	Genome-wide association meta-analysis in Chinese and European individuals identifies ten new loci associated with systemic lupus erythematosus.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Jin Y	2016	Nat. Genet.	Genome-wide association studies of autoimmune vitiligo identify 23 new risk loci and highlight key pathways and regulatory variants.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Bentham J	2015	Nat. Genet.	Genetic association analyses implicate aberrant regulation of innate and adaptive immunity genes in the pathogenesis of systemic lupus erythematosus.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Yin X	2015	Nat Commun	Genome-wide meta-analysis identifies multiple novel associations and ethnic heterogeneity of psoriasis susceptibility.	Enter comment	Show/Add Details	

Page: 1 1 to 10 of 23 rows

Add PubMed Reference Add File Reference Add Web Reference Add bioRxiv Reference

Editable curator comment. Changes are automatically saved to the target-gene database.

Button to remove user-added references.

Toggles to show/hide Detail elements attached to each reference (see next page). Number of Detail elements shown in parentheses.

Checkboxes to declare references having been reviewed.

Buttons to add new references of different types.

Text-based/tabular view of curated data.

References section

Graphic elements (Details) can be saved with each reference

References (18 out of 23 not yet reviewed)

Reviewed	1st Author	Year	Journal/Site/File	Title/Description	Curator Comment		
<input checked="" type="checkbox"/>		2019	https://clue.io/repurposing-app	Curated information about drugs and drug targets from Broad.	Enter comment	Show/Add Details	X
<input checked="" type="checkbox"/>	Li Q	2017	Pharmacogenet. Genomics	Genome-wide association study of paliperidone efficacy.	Enter comment	Show/Add Details	
<input checked="" type="checkbox"/>		2017	http://www.nealelab.is/blog/2017/7/19/rapid-gwas-of-thousands-of-phenotypes-for-337000-samples-in-the-uk-biobank	UK Biobank Rapid GWAS	Enter comment	Show/Add Details	X
<input checked="" type="checkbox"/>	Pettersson M	2017	Am. J. Med. Genet. A	Further evidence for specific IFIH1 mutation as a cause of Singleton-Merten syndrome with phenotypic heterogeneity.	Enter comment	Show/Add Details	X
<input checked="" type="checkbox"/>	Gorman JA	2017	Nat. Immunol.	The A946T variant of the RNA sensor IFIH1 mediates an interferon program that limits viral infection but increases the risk for autoimmunity.	Enter comment	Hide Details (1)	

Functional data linking rs1990760 (A946T) and protection from infection challenge in mice.

Section Assignment

Functional

Zoomable thumbnail of Detail.

Remove

Editable description field. Changes are automatically saved to the target-gene database.

Button to remove Detail

Button to add new Detail (see next page).

	Author	Year	Journal/Site/File	Title/Description	Curator Comment		
<input type="checkbox"/>	Bronson PG	2016	Nat. Genet.	Common variants at PVT1, ATG13-AMBRA1, AH1 and CLEC16A are associated with selective IgA deficiency.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Morris DL	2016	Nat. Genet.	Genome-wide association meta-analysis in Chinese and European individuals identifies ten new loci associated with systemic lupus erythematosus.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Jin Y	2016	Nat. Genet.	Genome-wide association studies of autoimmune vitiligo identify 23 new risk loci and highlight key pathways and regulatory variants.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Bentham J	2015	Nat. Genet.	Genetic association analyses implicate aberrant regulation of innate and adaptive immunity genes in the pathogenesis of systemic lupus erythematosus.	Enter comment	Show/Add Details	
<input type="checkbox"/>	Yin X	2015	Nat Commun	Genome-wide meta-analysis identifies multiple novel associations and ethnic heterogeneity of psoriasis susceptibility.	Enter comment	Show/Add Details	

Page: 1 1 to 10 of 23 rows 10

Add PubMed Reference Add File Reference Add Web Reference Add bioRxiv Reference

Dropdown menu to select the applicable section in which the Detail should appear.

- Functional
- None
- Association
- Clinical
- Expression
- Functional
- PQTL
- Protein

Text-based/tabular view of curated data.

References section

Adding a Detail

Add Detail



New Detail

Enter description

Detail graphic can be dragged here or pasted from the system clipboard.

Save

Text-based/tabular
view of curated
data.

Notebook utilities...

Overview page of available Target Gene Notebooks

This is the TGN server's index.html page

Target Gene Notebook

User Guide Manage Tags About

IBD Chol Metab CAD

Available Notebooks

Gene	Chr	Mb Location	Neighborhood Coding Genes	References Status	Last Modified Time	Tags
IFIH1	2	162.3	FAP GCA GCG KCNH7	18 out of 23 not reviewed	2019-04-11T19:04:36.353346Z	IBD
PCSK9	1	55.1	BSND TMEM61 USP24	22 out of 22 not reviewed	2019-04-13T18:43:48.814601Z	Chol Metab CAD

Links to available Target Gene Notebooks.

Tags assigned to each Target Gene Notebook

Link to Tag-management tools (see next page).

Toggle to filter TGN list according to assigned Tags.

Target Gene Notebook

User Guide Manage Tags About

IBD Chol Metab CAD

Available Notebooks

Gene	Chr	Mb Location	Neighborhood Coding Genes	References Status	Last Modified Time	Tags
IFIH1	2	162.3	FAP GCA GCG KCNH7	18 out of 23 not reviewed	2019-04-11T19:04:36.353346Z	IBD

Notebook utilities.

Tag-management tools

Manage Tags



Target Gene Notebook User Guide About

Existing Tags

Tag	Tag Class	Short Name	Long Name	Description	
CAD	Disease association	CAD	Coronary artery disease		Edit Delete
Chol Metab	Pathway	Chol Metab	Cholesterol metabolism		Edit Delete
IBD	Disease association	IBD	Inflammatory Bowel Disease		Edit Delete

Button to create a new Tag. New Tag

Assign by Tag

Push Web Reference to Notebooks

Assign by TGN

Button to assign specific Tags to Target Gene Notebooks.

Button to add web references to notebooks *en masse*.

Button to assign specific Target Gene Notebooks to Tags.

Tag-management tools

Creating a Tag

New Tag



Existing Tags

Create/Edit Tag

Class:

Short name:

Long name:

Tag Color:

Enter description

Description:

Save

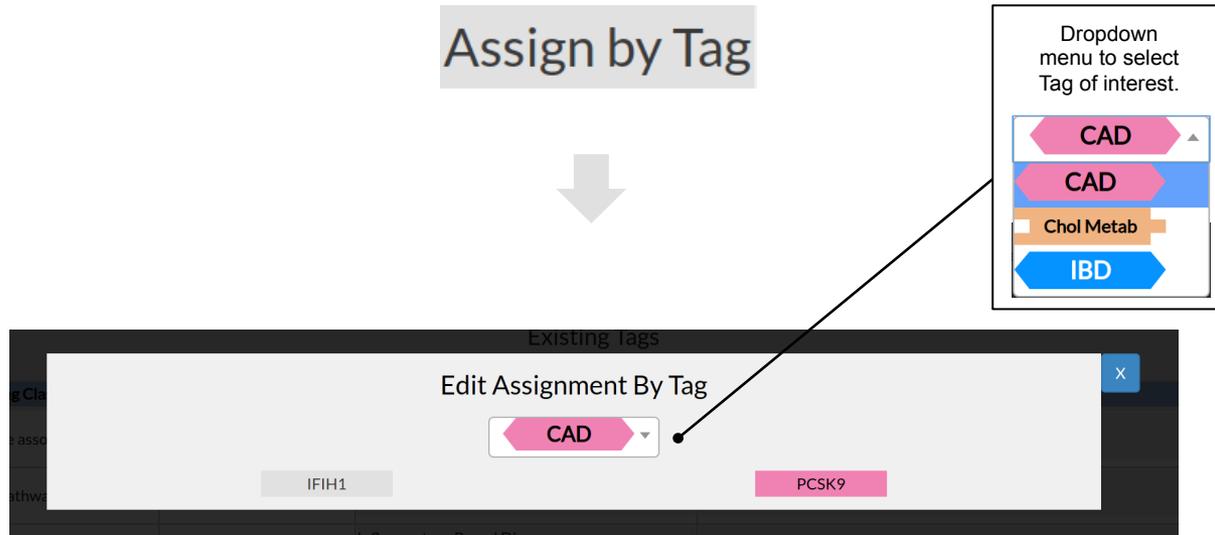
Dropdown menu to select Tag class, which determines Tag shape.

- Please select
- Collaboration
- Disease association
- Pathway
- Point person
- Project name
- Publication
- Site
- Statistic

Button to open color selector.

Tag-management tools

Assigning specific Tags to Target Gene Notebooks



Use Notebook buttons to toggle Tag assignment: if the Notebook button is the same color as the Tag, then that Tag is assigned to the Notebook.

Tag-management tools

Assigning specific Target Gene Notebooks to Tags

Assign by TGN

↓

Dropdown menu to select Target Gene Notebook of interest.

IFIH1
IFIH1
PCSK9

Existing tags

Edit Assignment By TGN

IFIH1

CAD Chol Metab IBD

Use Tag buttons to toggle Tag assignment: if the Tag button is the same color as the Tag, then the Notebook is assigned to that Tag.

Tag-management tools

Pushing a web reference to existing Notebooks
(yes, we know this is technically not Tag management)

Push Web Reference to Notebooks



Existing Tags

Push Web Reference to Notebooks

Year:

Enter title

Select All De-Select All

IFIH1 PCSK9

Toggle buttons to indicate Notebooks to receive the web reference. Those Notebooks shaded blue will be sent the reference.

Notebooks that already have the web reference will ignore the push.