Gene ID Conversion Using R Bioconductor and NCBI gene_info

CFDE: Gene Working Group
Biomedical Data Commons Workbench (BDCW)
Metabolomics Workbench DCC

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Motivation

- > Genomic and gene expression data is integral to biomolecular data analysis.
- ➤ The types of identifiers used for genes may differ across different resources (such as GEO, ENCODE, GTEx) providing such data sets.
- The ability to use a single type of gene identifier is crucial for integrating data from two or more resources. Most downstream analysis tools work with (or internally convert IDs to) ENTREZ ID.
- > This gene ID conversion tool will facilitate the use of a common gene identifier.
- ➤ NGS/RNASeq mappers generally provide read-counts (or equivalent quantities) at the **gene-level** or transcript-level.
- ➤ The most common gene IDs are gene **SYMBOL** (e.g., KLF4) AND **Ensembl GENE ID** (e.g., ENSG00000136826.*n*).
- ➤ The most common transcript IDs are NCBI Refseq ID (e.g., NM_001314052.n) or Ensembl Transcript ID (ENST00000420475.n).
- > SYMBOLs are more human readable and change for yet not well studied genes. If a new SYMBOL is assigned to a gene, its current SYMBOL generally gets included in its ALIAS.

User Interface

Web-based interface: A user can specify the name of the organism, enter the list of genes or upload a file containing the gene list and select the ID type for the gene list provided (e.g., SYMBOL or ENTREZ ID). Then, the user can select the types of the converted IDs to be included in the output.

Species (currently human, mouse and rat)

List of genes (Symbol, Ensembl gene ID, etc.; only one ID type in a list)

as a typed list (if just a few genes; white-space or comma separated)

or as a file with one column or row (white-space, comma separated)

or as a file with 2 columns (if additional information is available)

2 columns: GENEID/SYMBOL CHR

The type of gene identifier provided (SYMBOL, ENTREZ GENE ID, ALIAS, ENSEMBL GENE ID, REFSEQ TRANSCRIPT ID).

Challenges

- Most current focus on SYMBOL to ENTREZ ID conversion.
- > Approach
 - □ R packages AnnotationDbi and org.Xy.eg.db are used to convert the IDs.
 - □ If Gene ID type is "SYMBOL or ALIAS" (most suitable choice), then it is searched in SYMBOL first and if not found then it is searched in ALIAS and the results are combined. The SYMBOLs not found in ALIAS too are searched (case-insensitive) in <u>Homologene</u> database to see if it exists in another organism.
- ➤ When searching in ALIAS, a SYMBOL may appear in ALIAS of several genes. For example, AIM1 appears in ALIAS of CRYBG1, AURKB and SLC45A2. Which one is correct? In this case, additional information such as chromosome number will be useful to resolve it to one (hopefully) or two genes.
- For a very small set of genes, a SYMBOL may give two different ENTREZ IDs. For example, searching for MEMO1 on NCBI gene website (https://www.ncbi.nlm.nih.gov/gene/?term=human+MEMO1) lists:

Name/Gene ID	Description	Location	Aliases	MIM
☐ <u>MEMO1</u> ID: 51072	mediator of cell motility 1 [Homo sapiens (human)]	Chromosome 2, NC_000002.12 (3186718832011008, complement)	C2orf4, CGI-27, MEMO, NS5ATP7	611786
MEMO1 ID: 7795	Methylation modifier for class I HLA [<i>Homo</i> sapiens (human)]			601201

Use of chromosome number will resolve this.

Gene ID Conversion

Available as a web-based tool: https://bdcw.org/geneid/geneidconv.php

Also as an SmartAPI: https://smart-api.info/ui/e712b9eb07e637a00ae468f757ce2a1f

For integration with other tools and scripts:

Uses php _GET so that gene symbol can be embedded in the URL itself:



Example URL: targeted for programmatic/API access:

URL to use for json output with CLI (e.g., using [curl -L 'URL']; use &View=txt for text output):

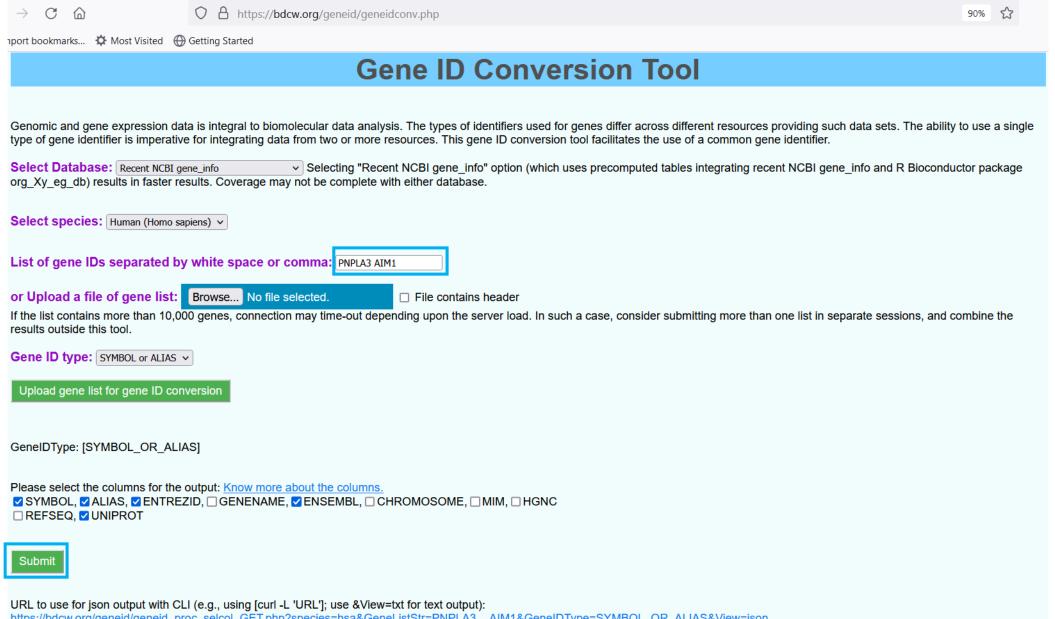
REST format URL:

https://bdcw.org/geneid/rest/species/hsa/GeneIDType/SYMBOL OR ALIAS/GeneListStr/I TPR3 IL6 KLF4/View/json

General purpose page:

https://bdcw.org/geneid/geneidconv.php

- Single and multiple genes
- As a list or via file upload



https://bdcw.org/geneid/geneid_proc_selcol_GET.php?species=hsa&GeneListStr=PNPLA3_AIM1&GeneIDTvpe=SYMBOL_OR_ALIAS&View=ison

General purpose page: https://bdcw.org/geneid/geneidconv.php

The first five rows of gene ID conversion results are given below. To further select, click on HTML. <u>Download txt. HTML JSON.</u>

	SYMBOL_OR_ALIAS	MATCH_SOURCE	ALIAS	SYMBOL	ENTREZID	ENSEMBL	GENENAME	REFSEQ	UNIPROT	KEGG	MARRVEL	LandingPages
113	PNPLA3	SYMBOL	ADPN C22orf20 iPLA(2)epsilon PNPLA3	PNPLA3	80339	ENSG00000100344	patatin like phospholipase domain containing 3	NM_025225	Q9NST1	80339	<u>80339</u>	PNPLA3
114	PNPLA3	SYMBOL	ADPN C22orf20 iPLA(2)epsilon PNPLA3	PNPLA3	80339	ENSG00000100344	patatin like phospholipase domain containing 3	NP_079501	Q9NST1	80339	<u>80339</u>	PNPLA3
1	AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystallin beta- gamma domain containing 1	NM_001371242	Q9Y4K1	202	202	CRYBG1

Disambiguation: AIM1 is alias for CRYBG1, AURKB and SLC45A2

	Select	SYMBOL_OR_ALIAS	MATCH_SOURCE	ALIAS	SYMBOL	ENTREZID	ENSEMBL	GENENAME	REFSEQ	UNIPROT	KEGG	MARRVEL	LandingPages
113		PNPLA3	SYMBOL	ADPN C22orf20 iPLA(2)epsilon PNPLA3	PNPLA3	80339	ENSG00000100344	patatin like phospholipase domain containing 3	NM_025225	Q9NST1	80339	80339	PNPLA3
114		PNPLA3	SYMBOL	ADPN C22orf20 iPLA(2)epsilon PNPLA3	PNPLA3	80339	ENSG00000100344	patatin like phospholipase domain containing 3	NP_079501	Q9NST1	80339	80339	PNPLA3
1		AIM1	ALIAS	AIM1	CRYBG1	202		crystallin beta-gamma domain containing 1	NM_001371242	Q9Y4K1	202	202	CRYBG1
459		AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora kinase B	XP_016880799	Q96GD4	9212	9212	<u>AURKB</u>
473		AIM1	ALIAS	AIM1	<u>AURKB</u>	9212	ENSG00000178999	aurora kinase B	XP_016880800	Q96GD4	9212	9212	<u>AURKB</u>
486		AIM1	ALIAS	AIM1	SLC45A2	<u>51151</u>	ENSG00000164175	solute carrier family 45 member 2	NM_001012509	Q9UMX9	<u>51151</u>	<u>51151</u>	SLC45A2
492		AIM1	ALIAS	AIM1	SLC45A2	<u>51151</u>	ENSG00000164175	solute carrier family 45 member 2	NM_001297417	Q9UMX9	<u>51151</u>	<u>51151</u>	SLC45A2

\rightarrow	\mathbb{C}	0	8	https://bdcw.org/geneid/geneidconv.php

ENSEMBL IDs 80% \$\frac{1}{2}





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Gene ID Conversion Tool

Genomic and gene expression data is integral to biomolecular data analysis. The types of identifiers used for genes differ across different	
esources providing such data sets. The ability to use a single type of gene identifier is imperative for integrating data from two or more resour	се
his gene ID conversion tool facilitates the use of a common gene identifier.	

Select Database:	Recent NCBI gene_info	~	Selecting "Recent NCBI gene_info" option (which uses precomputed tables integrating	
ecent NCBI gene_i	nfo and R Bioconductor package	org	g_Xy_eg_db) results in faster results. Coverage may not be complete with either database	se

e	lect	speci	ies:	Human	(Homo	sapiens)	~

	ist of	gene ID	s separa	ated by	white si	oace or	comma:
--	--------	---------	----------	---------	----------	---------	--------

r U	pload a file of o	ene list:	Browse	No file selected.	☐ File contains head
	pioau a ilie oi ţ	Jene nat.	DIOWSC	INO IIIO SCIECICU.	

the list contains more than 10,000 genes, connection may time-out depending upon the server load. In such a case, consider submitting more nan one list in separate sessions, and combine the results outside this tool.

ene ID type:	ENSEMBL	~

Upload gene list for gene ID conversion

he file genelist_hg38_GenCode_V28_ENSG_subset.txt has been uploaded as uploads/genelist_hg38_GenCode_V28_ENSG_subset.txt. GenelDTvpe: [ENSEMBL]

a file is uploaded and chromosomal coordinates are provided (e.g., SYMBOL CHR START END), specify genome version.

- the file has only two columns (SYMBOL CHR), then there is no need to specify genome version.
- Select closest genome version: Select...

Please select the columns for the output: Know more about the columns.

- 🗸 SYMBOL, 🗆 ALIAS, 💌 ENTREZID, 🗀 GENENAME, 💟 ENSEMBL, 🗀 CHROMOSOME, 🗀 MIM, 🗀 HGNC
- □ REFSEQ, □ UNIPROT

Submit

Ability to further select a gene subset.

The first five rows of gene ID conversion results are given below. To further select, click on HTML Download txt. HTML. JSON.

	ENSEMBL.VER	ENSEMBL	SYMBOL	ENTREZID	LandingPages
1	ENSG00000131781.12	ENSG00000131781	FMO5	2330	FMO5
49	ENSG00000136244.11	ENSG00000136244	<u>IL6</u>	<u>3569</u>	<u>IL6</u>
99	ENSG00000131778.18	ENSG00000131778	CHD1L	<u>9557</u>	CHD1L
294	ENSG00000143167.11	ENSG00000143167	GPA33	10223	GPA33
298	ENSG00000164794.8	ENSG00000164794	KCNV1	27012	KCNV1

Iotal time for gene ID conversion: : 1.497 sec elapsed

	Select	ENSEMBL.VER	ENSEMBL	SYMBOL	ENTREZID
1	✓	ENSG00000131781.12	ENSG00000131781	FMO5	2330
49	✓	ENSG00000136244.11	ENSG00000136244	IL6	3569
99		ENSG00000131778.18	ENSG00000131778	CHD1L	<u>9557</u>
294		ENSG00000143167.11	ENSG00000143167	GPA33	10223
298	✓	ENSG00000164794.8	ENSG00000164794	KCNV1	27012
300		ENSG00000186092.6	ENSG00000186092	OR4F5	<u>79501</u>
302	✓	ENSG00000268020.3	ENSG00000268020	OR4G4P	<u>79504</u>
303		ENSG00000223972.5	ENSG00000223972	DDX11L2	<u>84771</u>
305		ENSG00000143194.12	ENSG00000143194	MAEL	84944
353		ENSG00000198842.9	ENSG00000198842	STYXL2	92235
357		ENSG00000173950.15	ENSG00000173950	XXYLT1	152002
383		ENSG00000284742.1	ENSG00000284742	<u>OR11P1P</u>	<u>282795</u>
384		ENSG00000143195.12	ENSG00000143195	ILDR2	<u>387597</u>
402		ENSG00000196832.4	ENSG00000196832	OR11G2	390439
406		ENSG00000213068.3	ENSG00000213068	RPS17P6	<u>391130</u>
407		ENSG00000226653.3	ENSG00000226653	OR13Z1P	403228
408		ENSG00000179428.2	ENSG00000179428	IL6-AS1	<u>541472</u>
409		ENSG00000226015.2	ENSG00000226015	CCT8P1	<u>644131</u>
410		ENSG00000237613.2	ENSG00000237613	FAM138A	645520
411		ENSG00000184388.5	ENSG00000184388	PABPC1L2B	645974
413		ENSG00000227232.5	ENSG00000227232	WASH7P	<u>653635</u>
414		ENSG00000223972.5	ENSG00000223972	DDX11L16	727856
416		ENSG00000275520.1	ENSG00000275520	FAM236A	100129407

For API-based access to integrate in user's existing tools:

Important: For some genes, multiple IDs may be listed. The users needs to select one or more for further use in a script or on the web.

URLs to use for json output with CLI (e.g., using [curl -L 'URL']; use &View=txt for text output):

https://bdcw.org/geneid/geneid_proc_selcol_GET.php?species=hsa&GeneListStr=AIM1&GeneIDType=SYMBOL_OR_ALIAS&View=txt https://bdcw.org/geneid/geneid_proc_selcol_GET.php?species=hsa&GeneListStr=ITPR3 IL6 KLF4&GeneIDType=SYMBOL_OR_ALIAS&View=json

REST format URL:

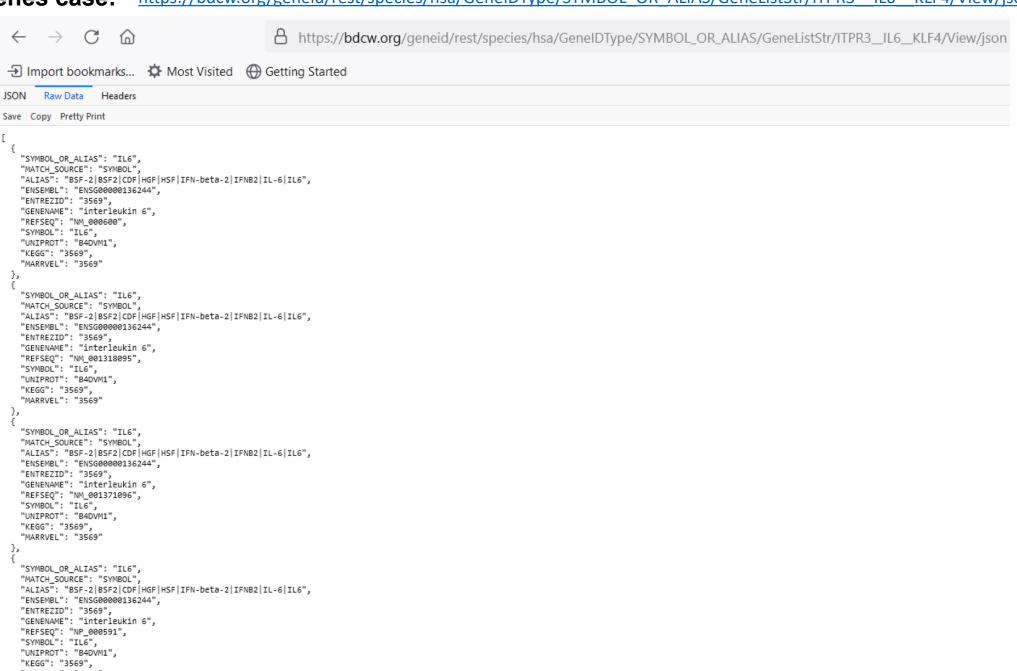
https://bdcw.org/geneid/rest/species/hsa/GeneIDType/SYMBOL OR ALIAS/GeneListStr/AIM1/View/json https://bdcw.org/geneid/rest/species/hsa/GeneIDType/SYMBOL OR ALIAS/GeneListStr/ITPR3 IL6 KLF4/View/json

Single gene case: https://bdcw.org/geneid/geneid_proc_selcol_GET.php?species=hsa&GeneListStr=AIM1&GeneIDType=SYMBOL_OR_ALIAS&View=txt

← → C 🗅	https://bdcw.org/geneid/geneid_proc_selcol_GET.php?species=hsa&GeneListStr=AIM1&GeneIDType=SYMBOL_OR_ALIAS&View=tx							
→ Import bookmarks 🌣 Most Visited 🌐 Getting Started								

SYMBOL_	OR_ALIAS	MATCH_S	OURCE	ALIAS	SYMBOL ENTREZIO)	ENSEMBI	GENENAME		REFSEQ	UNIPROT	KEGG	MARRVEL			
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NM_001	371242	Q9Y4K1	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NM_001	624	Q9Y4K1	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NP_001	358171	Q9Y4K1	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NP_001	615	Q9Y4K1	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	XM_017	010333	Q9Y4K1	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	XP_016	865822	Q9Y4K1	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NM_001	371242	B3KPT0	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NM_001	624	B3KPT0	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NP_001	358171	B3KPT0	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	NP_001	615	B3KPT0	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	XM_017	010333	B3KPT0	202	202
AIM1	ALIAS	AIM1	CRYBG1	202	ENSG00000112297	crystall	in beta	a-gamma doma	ain co	ntaining	1	XP_016	865822	B3KPT0	202	202
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	NM 0012568	334	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM 0012845	526	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM_0013139	950	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM 0013139	951	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM 0013139	952	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM 0013139	953	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM 0013139	954	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NM 0013139	955	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	NM 004217		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	3 NP 0012437	763	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	NP 0012714	155	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999	aurora k	inase B	NP 0013008	381	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999					_	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		_	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_	372	Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999					Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		096GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999					096GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		Q96GD4	9212	9212				
AIM1	ALIAS	AIM1	AURKB	9212	ENSG00000178999			_		096GD4	9212	9212				
AIM1	ALIAS	AIM1	SLC45A2		ENSG00000176333			- -		_	NM 0010:		Q9UMX9	51151	51151	
AIM1	ALIAS	AIM1	SLC45A2		ENSG00000164175	_	_	*			NM 00129		-		51151	
AIM1	ALIAS	AIM1	SLC45A2		ENSG00000164175			•			NM 01618		Q9UMX9	51151	51151	
AIM1	ALIAS	AIM1	SLC45A2		ENSG00000164175						NP 00101		Q9UMX9		51151	
			JECTORE	2222		-		"		_	5010.		6201 IV2	51151	32232	

Multiple genes case: https://bdcw.org/geneid/rest/species/hsa/GeneIDType/SYMBOL OR ALIAS/GeneListStr/ITPR3 IL6 KLF4/View/json



How to use it in a python program:

A python script provides an example of how to use the gene ID conversion program. At the core, a URL-based query is constructed and executed using python packages "requests" and "bs4" (function "BeutifulSoup"). After some processing, the results are available as a pandas dataframe.

https://bdcw.org/geneid/fetch_php_page.py