

Quick Guide of AS-ALPS

Front Page

On the search form in the top right corner, you can search for a cluster of transcripts produced by AS from a single gene, in either human or mouse, specified by either Entrez Gene ID (e.g., 10681) or Transcript ID in H-InvDB, FANTOM or RefSeq (e.g., NM_016194).

→ Go to Page 3

Alternatively, you can move to "Advanced Search" page by clicking the hyper-link below the search box.

→ Go to Page 2

AS-ALPS ver. 1.0

Dataset : Human

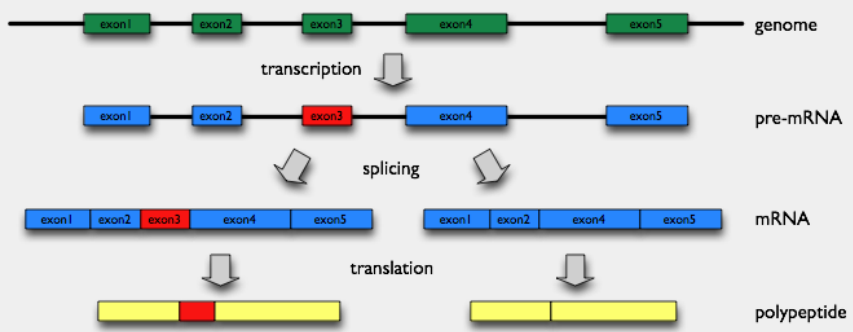
Search for

[Advanced Search](#)

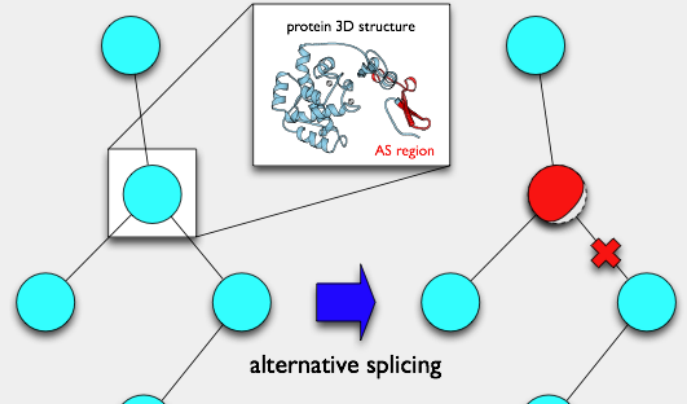
QuickGuide : [\(pdf\)](#), [\(html\)](#)

About this database

Alternative splicing (AS) is a molecular mechanism that produces multiple proteins from a single gene, and is thought to increase the number of proteins extensively over the number of genes in a genome, which could be required to generate complexity of higher eukaryote including human. However, what roles most of AS variants play are not known.



This database, AS-ALPS (Alternative Splicing-induced ALteration of Protein Structure), is aimed at providing useful information to analyze effect of AS on protein interaction and network through alteration of protein structure.



Advanced Search Page 1

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Dataset : Human

Search for

[Advanced Search](#)

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Search AS variant clusters by annotation

DataSet		Human	
Protein name-related keywords		<input type="text"/> (e.g. dna binding)	
Attributes of AS regions (regions of amino acid sequences changed by AS)			
<input type="checkbox"/> Position	<input type="radio"/> N-terminal		
	<input type="radio"/> Middle		
	<input type="radio"/> C-terminal		
	<input type="checkbox"/> Type	<input type="radio"/> Insertion/Deletion	
		<input type="radio"/> Substitution	
	<input type="checkbox"/> 3D structure	<input type="radio"/> Available	
		Inclusion of hydrophobic core residues	<input type="checkbox"/> Yes
Inclusion of interaction sites with -		<input type="checkbox"/> Protein <input type="checkbox"/> DNA/RNA <input type="checkbox"/> Ligand/Substrate	
<input type="radio"/> AND <input type="radio"/> OR			
<input type="radio"/> Not available			
KEGG pathway-related keywords		<input type="text"/> (e.g. glycolysis)	
<input type="button" value="Search"/>		<input type="button" value="Reset"/>	

1. Search by annotation

You can combine the search conditions below.

- (1) human or mouse
- (2) protein name
- (3) attributes of AS regions:
 - (a) position
 - (b) type
 - (c) if 3D info is available,
 - (i) whether AS regions include hydrophobic core residues
 - (ii) what kind of interaction sites AS regions include
- (4) KEGG pathway name

→ Go to Page 3

2. Search by sequence

→ Go to Page 12

Search AS variant clusters by sequence

DataSet: Human


Enter a query amino acid sequence in FASTA format:

Search Results Page

In this table of search results, annotated info is briefly shown for each cluster of transcripts produced by AS from a single gene.

Click a button in the rightmost column to view AS variants in a cluster.

→ Go to Page 4



Dataset : Human
Search Transcript ID for Go
[Advanced Search](#)

QuickGuide : [\(pdf\)](#), [\(html\)](#)

4 clusters of AS variants was found

1 - 4 of 4

Cluster No.	species	chr	KEGG pathway	Entrez Gene ID	protein name	No. of unique AS variants	Attributions of AS regions			
							position	type	3D structure	
1	human	12	<ul style="list-style-type: none"> Taste transduction 	2784	guanine nucleotide-binding protein, beta-3subunit	5	N-ter Mid C-ter	INDEL SUB	Available	View AS variants
2	human	7	-	2783	guanine nucleotide-binding protein, beta-2subunit	9	N-ter C-ter	INDEL SUB	Available	View AS variants
3	human	5	-	10399	guanine nucleotide binding protein (G protein),beta polypeptide 2-like 1	17	N-ter C-ter	INDEL SUB	Available	View AS variants
4	human	15	-	10681	guanine nucleotide-binding protein, beta-5subunit isoform b	8	N-ter Mid C-ter	INDEL SUB	Available	View AS variants

1 - 4 of 4

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You can see here a schematic representation of alignments between a chromosome and transcripts.

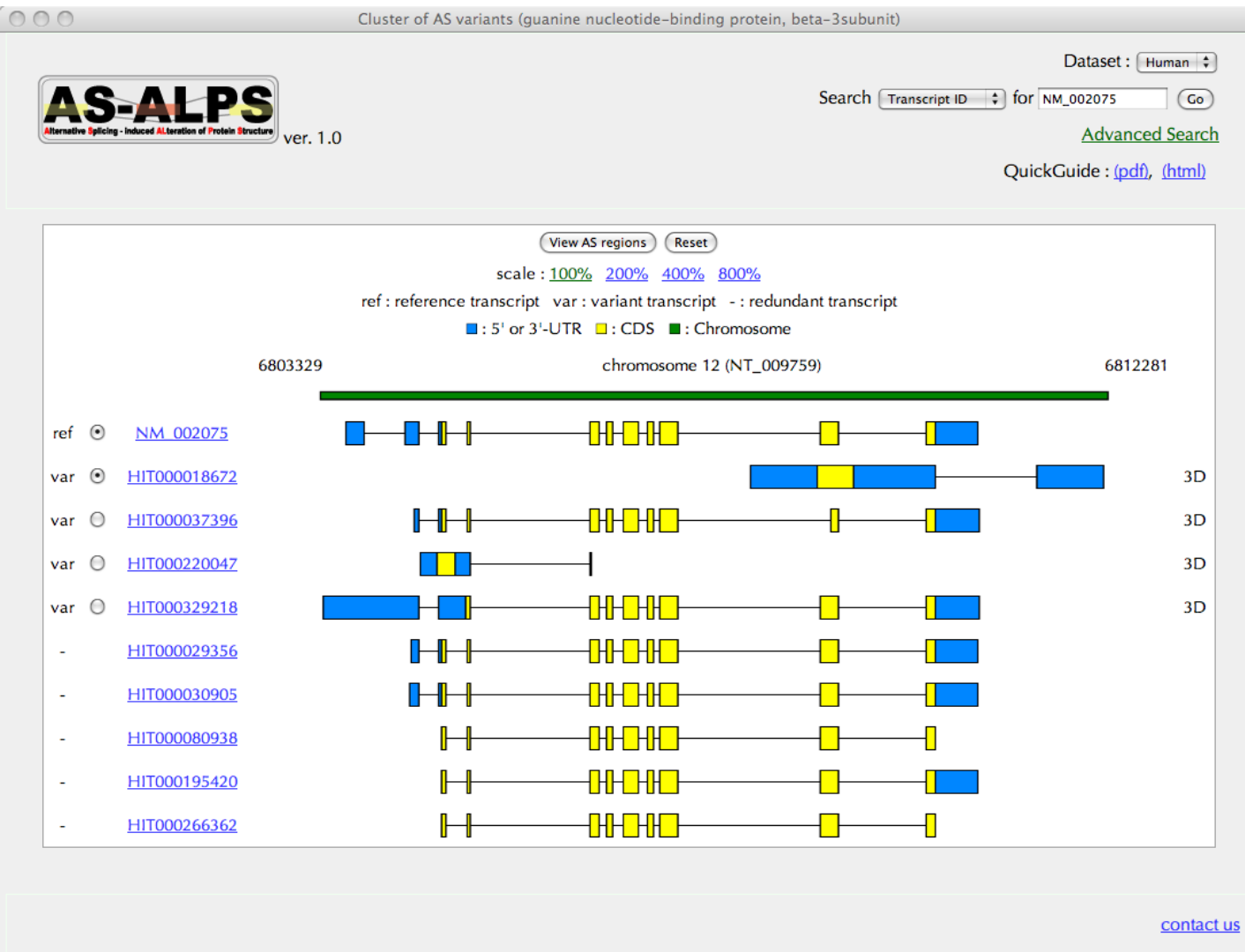
AS regions are detected by a comparison between a pre-defined reference transcript and one of other variant transcripts.

Select one variant transcript in the leftmost column and push the "View AS regions" button.

→ Go to Page 6

Click a transcript ID to see its details

→ Go to Page 5



Transcript Details Page

For each transcript requested, you can check the region of CDS (coding sequence), the status of truncation at the 5' and 3' termini, a possibility of NMD (nonsense-mediated mRNA decay) judged from the sequence, the translated amino acid sequence and the exon composition.

Cluster of AS variants (...) HIT000037396

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Alternative Splicing - induced ALteration of Protein Structure

Dataset : Human

Search Transcript ID for HIT000037396 Go


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details of transcript

transcript ID	accession ID	transcript length	CDS		truncation	amino acid length
HIT000037396	BC015920	1510	90	983	No	297

CDS



possibility of NMD No

amino acid sequence
 MGEMEQLRQEAEQLKKQIADARKACADVTLAELVSGLEVGRVQMRTRTLRLGHLAKIYA
 MHWATDSKLLVSASQDGKLIWDSYTTNKVHAIPLRSSWVMTCAYPGSGNFVACGGLDNNM
 CSIYNLKSREGNVKVSRELSAHTGYLSCCRFLDDNNIVTSSGDTTCALWDIETGQQTVE
 VGHTGDCMSLAVSPDFNLFIGACDASAKLWDVREGTCRQFTTGCHESDINAICFFSLSGR
 LLFAGYDDFNCNVWDSMKSERVGILSGHDNRVSLGVTADGMAVATGSWDSFLKIWN

exon structure

	transcript (HIT000037396)			genome (NT_009759)		
	start	end	length	start	end	length
exon 1	1	59	59	6804400	6804458	59
exon 2	60	146	87	6804683	6804769	87
exon 3	147	185	39	6805011	6805049	39
exon 4	186	292	107	6806395	6806501	107
exon 5	293	356	64	6806599	6806662	64
exon 6	357	519	163	6806788	6806950	163
exon 7	520	586	67	6807057	6807123	67

AS regions View Page

You can see here AS regions detected by a comparison between a pre-defined reference transcript and one variant transcript selected in the previous page.

For the substitution type, identity between amino acid sequences of AS regions mutually replaced is shown if comparable.

Click the button to show results of 3D structure assignment to AS regions.

→ Go to Page 7

AS regions from comparison between NM_016194 and HIT000024633 (guanine nucleotide-binding protein, beta-5subunit isoform b)

Dataset : Human

Search for

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AS region No.	transcript ID	region	length	color	position	type	identity	comment
1	NM_016194	1	42	42	N-terminal	deletion	-	-
	HIT000024633	1	1	1				
2	NM_016194	140	209	70	Middle	deletion	-	-
	HIT000024633	97	98	2				

1 : WD40 repeat-like (50 - 394)

ref: NM_016194

var: HIT000024633

1 : WD40_like (8 - 282)

Orange : Constitutive region Blue : Structural domain

Threshold of sequence identity for 3D structure assignment

transcript ID	amino acid sequence
reference: NM_016194	MCDQTFLLVNVFGSCDKCFKQRALRPVFKKSQQLSYCSTCAEIMATEGLHENETLASLKSEAESLKGKLEERAKLHDVELHQVAERVEALGQFVMKTRRTLKGHGKLVLCMDWCKDKRRIVSSSQDGKVIWDSFTTNKEHAVTMPCTWVMACAYAPSGCAIACGGLDNKCSVYPLTFDKNENMAAKKKSVMHTNYLSACSFNTSDMQILTSAGDGTALWDVDSGQLLQSFHGHGADVLCCLDLAPSETGNTFFVSGGCDKKAMVDMRSGCQVQAFETHESDINSVRYYPGDAFASGSDDATCRLYDLRADREVAIYSKESIIFGASSVDVFSLSGRLLFAGYNDYTINVWDVLKGSRSILFCHENRVSTLRVSPDGTAFCSGSWDHTLRVWA
variant: HIT000024633	MATEGLHENETLASLKSEAESLKGKLEERAKLHDVELHQVAERVEALGQFVMKTRRTLKGHGKLVLCMDWCKDKRRIVSSSQDGKVIWDSFTTNKILTSAGDGTALWDVDSGQLLQSFHGHGADVLCCLDLAPSETGNTFFVSGGCDKKAMVDMRSGCQVQAFETHESDINSVRYYPGDAFASGSDDATCRLYDLRADREVAIYSKESIIFGASSVDVFSLSGRLLFAGYNDYTINVWDVLKGSRSILFCHENRVSTLRVSPDGTAFCSGSWDHTLRVWA

Assigned 3D Structures Page

You can see here a schematic representation of alignments between a pair of transcripts and PDB entries with shown sequence identities to the reference transcript.

You can also check in the "annotation" column, whether each PDB entry includes interaction info.

Click a button in the rightmost column to see details of interaction info.
→ Go to Page 8

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Dataset : Human

Search for

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Deletion (NM_016194: 1-42 <-> HIT000024633: 1-1)
Deletion (NM_016194: 140-209 <-> HIT000024633: 97-98)

var: [HIT000024633](#) 283

ref: [NM_016194](#) 395

[reverse](#)

		ref region	PDB region	identity	annotation	
1GP2 :B	339		54-394 3-338	52.8 %	subunit	<input type="button" value="Details"/>
1TBC :A	340		54-394 4-339	52.8 %		<input type="button" value="Details"/>
1TBC :C	340		54-394 4-339	52.8 %		<input type="button" value="Details"/>
1XHM :A	339		54-394 3-338	52.8 %	subunit	<input type="button" value="Details"/>
1GG2 :B	339		54-394 3-338	52.8 %	subunit	<input type="button" value="Details"/>
1OMW :B	339		54-394 3-338	52.8 %	subunit	<input type="button" value="Details"/>
1TBC :B	340		54-394 4-339	52.8 %	subunit	<input type="button" value="Details"/>
1TBC :D	340		54-394 4-339	52.8 %		<input type="button" value="Details"/>
2BCI :B	339		54-394 3-338	52.8 %	subunit	<input type="button" value="Details"/>
1B9X :A	340		54-394 4-339	52.5 %	subunit	<input type="button" value="Details"/>
1GOT :B	339		54-394 3-338	52.5 %	subunit	<input type="button" value="Details"/>
1AOR :B	339		54-394 3-338	52.5 %	subunit	<input type="button" value="Details"/>
1B9Y :A	340		54-394 4-339	52.5 %	subunit	<input type="button" value="Details"/>
2TRC :B	340		54-394 4-339	52.5 %	subunit	<input type="button" value="Details"/>

1GP2B

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Dataset: Human

Search for

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PDB ID	1GP2:B
amino acid length	339
protein name	G PROTEIN GI BETA 1
species	Bos taurus
resolution	2.300

Subunit interaction residues

■ : alpha helix □ : beta strand ▣ : 3-10 helix

Summary

AS region 1 (89-155), type : deletion		
Protein interaction residues with chain A (G PROTEIN GI ALPHA 1)	23/54	details
Hydrophobic core2	5/9	details
Hydrophobic core3	4/6	details
Hydrophobic core4	1/6	details

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3D Structure-related Interaction Info Page

Interaction sites are depicted with triangles below a bar, in which AS regions are coloured by their types (deletion: red, insertion: cyan, substitution: violet) and constitutive regions are in light orange.

Click the button to show AS regions in 3D structure.

→ Go to Page 9

The lower half of the page summarizes the number of residues relating to interaction and hydrophobic core.

Details of interaction sites

→ Go to Page 10

Details of core residues

→ Go to Page 11

AS regions on 3D structure of 1GP2B

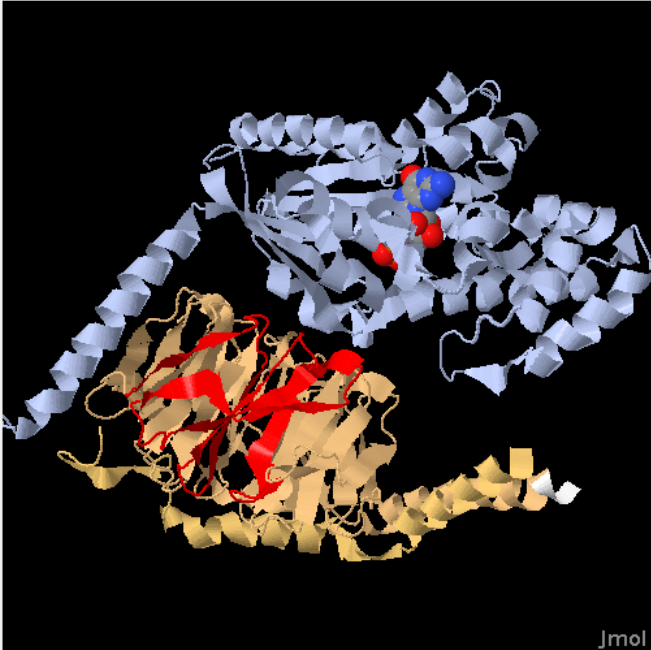
Dataset : Human

Search for

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ver. 1.0



☒ cartoon ☐ CPK
☐ spin
☐ show only a relevant protein chain
☐ show AS region 1 (89-155) by CPK model
☐ show subunit interaction residues by stick model
☐ show all core residues
☐ show core1 by CPK model
☐ show core2 by CPK model
☐ show core3 by CPK model
☐ show core4 by CPK model
☐ show core5 by CPK model
☐ show core6 by CPK model
☐ show core7 by CPK model

Jmol

AS regions on 3D Structure Page

A relevant protein chain with AS regions is shown in the same colours as in page 8.

Other interacting proteins and DNA/RNAs, if any, are also shown in different colours.

You can check locations of hydrophobic cores by clicking check boxes below the 3D viewer.

Details of interaction sites extracted from 3D structure

Residues included in a relevant AS region are coloured by its type (deletion: red, insertion: cyan, substitution: violet).

Details of Subunit interaction residues (1GP2B)

Dataset : Human

Search for

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Alternative Splicing - Induced Alteration of Protein Structure

AS region (89-155), type : deletion

contact residue	residue number	chain ID	contact residue	residue number	chain ID	Num. of pair of contacts
Arg	52	B	Asp	20	A	6
Arg	52	B	Arg	24	A	3
Gly	53	B	Leu	23	A	1
Leu	55	B	Gly	27	A	1
Lys	57	B	His	213	A	1
Lys	57	B	Glu	216	A	3
Tyr	59	B	His	213	A	3
Tyr	59	B	Cys	214	A	4
Gln	75	B	Cys	214	A	4
Lys	78	B	Asp	26	A	1
Ile	80	B	Leu	23	A	1
Asn	88	B	Ala	12	A	3
Asn	88	B	Val	13	A	6
Asn	88	B	Ser	16	A	3
Lys	89	B	Arg	15	A	1
Lys	89	B	Ser	16	A	5
Lys	89	B	Ile	19	A	4

Details of hydrophobic core residues extracted from 3D structure

Residues included in a relevant AS region are coloured by its type (deletion: red, insertion: cyan, substitution: violet).

Details of Core residues (1GP2B)

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Alternative Splicing - Induced ALteration of Protein Structure

Dataset : Human

Search Transcript ID for NM_016194 Go

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AS region (89-155), type : deletion

core ID	core residue	residue number
core2	Val	71
core2	Ala	73
core2	Leu	79
core2	Ile	81
core2	Ile	93
core2	Leu	95
core2	Val	100
core2	Val	112
core2	Leu	126

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AS-ALPS

Protein name-related keywords (e.g. dna binding)

Attributes of AS regions (regions of amino acid sequences changed by AS)

<input type="checkbox"/> Position	<input type="radio"/> N-terminal	
	<input type="radio"/> Middle	
	<input type="radio"/> C-terminal	
<input type="checkbox"/> Type	<input type="radio"/> Insertion/Deletion	
	<input type="radio"/> Alteration	
<input type="checkbox"/> 3D structure	<input type="radio"/> Available	
	Inclusion of hydrophobic core residues	<input type="checkbox"/> Yes
	Inclusion of interaction sites with -	<input type="checkbox"/> Protein <input type="checkbox"/> DNA/RNA <input type="checkbox"/> Ligand/Substrate <input type="radio"/> AND <input type="radio"/> OR
	<input type="radio"/> Not available	

KEGG pathway-related keywords (e.g. glycolysis)

Advanced Search Page 2

Search by sequence

You can search entries in this database by amino acid sequence. The search engine is FASTA.

→ Go to Page 13

Search AS variant clusters by sequence

DataSet:

Enter a query amino acid sequence in FASTA format:

```

MCDQTFVLNVFGSCDKCFKQRLRPVFKKSQQLSYCSTCAEIMATEGLHENETLASLKSE
AESLKGKLEERAKLHDVELHQVAERVEALGFVMTKTRTLKGHNKVLCDWCKDKRRI
VSSSQDGKVVWDSFTTNKEHAVTMPCTWVMACAYAPSGCAIACGGLDNKCSVYPLTFDK
NENMAAKKKSVMHTNYLSACSFNSDMQILTASGDGTALWDVSEGLQSFHGHGADV
LCLDLAPSETGNTFVSGGCDKKAMVWDMRSGQCQAFETHESDINSVRYYPGDAFASGS
DDATCRLYDLRADREVAIYSKESIIFGASSVDLSGLRLLFAGYNDYTINVVDVLKGSRV
SILFGHENRVSTLRVSPDGTAFCSGSWDHTLRVWA
  
```

FASTA Search Result Page 1

You can see here a list of transcripts found by FASTA search.

Click one of transcript IDs in the leftmost column.

→ Go to Page 3

If you would like to check sequence alignments, click "align" in the rightmost column.

→ Go to Page 14

FASTA Search Result

Dataset : Human

Search for

[Advanced Search](#)

Quick Guide : [\(pdf\)](#), [\(html\)](#)

Search Result

FASTA searches a protein or DNA sequence data bank
version 34.26.5 April 26, 2007
Please cite:
W.R. Pearson & D.J. Lipman PNAS (1988) 85:2444-2448

FASTA (3.5 Sept 2006) function [optimized, BL50 matrix (15:-5)] ktup: 2
join: 37, opt: 25, open/ext: -10/-2, width: 16
Scan time: 22.960

The best scores are:

Transcript ID	Score	Align
NM_016194		
HIT000076644		align
NM_006578		
HIT000062440		align
HIT000301841		align
HIT000080939		align
HIT000304087		align
HIT000036456		align
HIT000024633		align
HIT000045575		align
NM_002074		
HIT000321048		align
HIT000274391		align
HIT000032499		align
HIT000298500		align
HIT000034130		align
HIT000272088		align
HIT000100225		align
HIT000302622		align
HIT000270979		align
HIT000031765		align
HIT000267634		align
HIT000080936		align
NM_021629		
HIT000004364		align
HIT000076642		align
HIT000029967		align
HIT000270214		align
NM_005273		
HIT000262878		align

>>>QUERY, 395 aa vs ./as_seq_hs.fasta library

>>NM_016194 (395 aa)

initn: 2698 initl: 2698 opt: 2698 Z-score: 2730.4 bits: 514.0 E(): 5.9e-145
Smith-Waterman score: 2698; 100.000% identity (100.000% similar) in 395 aa
overlap (1-395:1-395)

FASTA Search Result Page 2

You can see here an alignment between your query sequence and a sequence picked up from this database by FASTA search.

```

FASTA Search Result
>>NM_016194 (395 aa)
  initn: 2698 initl: 2698 opt: 2698 Z-score: 2730.4 bits: 514.0 E(): 5.9e-145
Smith-Waterman score: 2698; 100.000% identity (100.000% similar) in 395 aa
overlap (1-395:1-395)

>NM_016 1- 395:-----:
      10      20      30      40      50      60      70      80
QUERY MCDQTFLLVNVFGSCDKCFKQALRPVFKKSQQLSYCSTCAEIMATEGLHENETLASLKSEAESLKGKLEERAKLHDVEL
NM_016 MCDQTFLLVNVFGSCDKCFKQALRPVFKKSQQLSYCSTCAEIMATEGLHENETLASLKSEAESLKGKLEERAKLHDVEL
      10      20      30      40      50      60      70      80
      90     100     110     120     130     140     150     160
QUERY HQVAERVEALGQFVMKTRRTLKGHGKVKLCMDWCKDKRRIVSSSQDGKVIWVDSFTTNKEHAVTMPCTWVMACAYAPSGC
NM_016 HQVAERVEALGQFVMKTRRTLKGHGKVKLCMDWCKDKRRIVSSSQDGKVIWVDSFTTNKEHAVTMPCTWVMACAYAPSGC
      90     100     110     120     130     140     150     160
      170     180     190     200     210     220     230     240
QUERY AIACGGLDNKC SVYPLTFDKNENMAAKKKSVMHTNYLSACSFTNSDMQILTAGSGDGTALWDVESGQLLQSFHGHGADV
NM_016 AIACGGLDNKC SVYPLTFDKNENMAAKKKSVMHTNYLSACSFTNSDMQILTAGSGDGTALWDVESGQLLQSFHGHGADV
      170     180     190     200     210     220     230     240
      250     260     270     280     290     300     310     320
QUERY LCLDLAPSETGNTFVSGGCDKKAMVWDMRSGQCQVAFETHESDINSVRYYPGDAFASGSDDATCRLYDLRADREVAIYS
NM_016 LCLDLAPSETGNTFVSGGCDKKAMVWDMRSGQCQVAFETHESDINSVRYYPGDAFASGSDDATCRLYDLRADREVAIYS
      250     260     270     280     290     300     310     320
      330     340     350     360     370     380     390
QUERY KESIIFGASSVDFSLSGRLLFAGYNDYTINVDVLKGSRVSIILFGHENRVSTLRVSPDGTAFCSGSDHDLTRVWA
NM_016 KESIIFGASSVDFSLSGRLLFAGYNDYTINVDVLKGSRVSIILFGHENRVSTLRVSPDGTAFCSGSDHDLTRVWA
      330     340     350     360     370     380     390

>>HIT000076644 (396 aa)
  initn: 2698 initl: 2698 opt: 2698 Z-score: 2730.3 bits: 514.0 E(): 5.9e-145
Smith-Waterman score: 2698; 100.000% identity (100.000% similar) in 395 aa
overlap (1-395:1-395)

>HIT000 1- 395:-----:
      10      20      30      40      50      60      70      80
QUERY MCDQTFLLVNVFGSCDKCFKQALRPVFKKSQQLSYCSTCAEIMATEGLHENETLASLKSEAESLKGKLEERAKLHDVEL
HIT000 MCDQTFLLVNVFGSCDKCFKQALRPVFKKSQQLSYCSTCAEIMATEGLHENETLASLKSEAESLKGKLEERAKLHDVEL
      10      20      30      40      50      60      70      80
      90     100     110     120     130     140     150     160
QUERY HQVAERVEALGQFVMKTRRTLKGHGKVKLCMDWCKDKRRIVSSSQDGKVIWVDSFTTNKEHAVTMPCTWVMACAYAPSGC
HIT000 HQVAERVEALGQFVMKTRRTLKGHGKVKLCMDWCKDKRRIVSSSQDGKVIWVDSFTTNKEHAVTMPCTWVMACAYAPSGC
      90     100     110     120     130     140     150     160
      170     180     190     200     210     220     230     240
QUERY AIACGGLDNKC SVYPLTFDKNENMAAKKKSVMHTNYLSACSFTNSDMQILTAGSGDGTALWDVESGQLLQSFHGHGADV
HIT000 AIACGGLDNKC SVYPLTFDKNENMAAKKKSVMHTNYLSACSFTNSDMQILTAGSGDGTALWDVESGQLLQSFHGHGADV
      170     180     190     200     210     220     230     240
      250     260     270     280     290     300     310     320

```