

BMG 744

# Bioinformatics/Genomics/Proteomics

The “ics” revolution  
March 4, 2003



# Contact Information

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## Molecular and Genetic Bioinformatics Facility

### General Information

- [UAB Bioinformatics Resources](#)
- [Sequence Analysis at UAB](#)
- [MIC753 - "Practical Applications of Sequence Analysis"](#)
- [CIS 640 - Bioinformatics I: "Lectures on Practical Bioinformatics" pdf](#)

### Genomic Sequencing

- [The Poxvirus Bioinformatics Resource](#)
- [The \*Streptococcus pneumoniae\* genome diversity project](#)
- [The \*Streptococcus pneumoniae\* strain SpR6 genome sequencing project](#)
- [The \*Ureaplasma urealyticum\* genomic sequencing project](#)

### UAB Only (Password required. Call or Email Elliot for access)

- [GCG at UAB](#)
- [SeqWeb - Web interface to GCG](#)
- [GCG 10 Documentation](#)
- [GCG 10 Documentation - Downloadable pdf files](#)

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# Definitions

(From NIH)

## ➤ Bioinformatics

- Research, development, or application of computational tools and approaches for expanding the use of biological, medical, behavioral or health data, including those to acquire, store, organize, archive, analyze, or visualize such data.

## ➤ Computational Biology

- The development and application of data-analytical and theoretical methods, mathematical modeling and computational simulation techniques to the study of biological, behavioral, and social systems.



# Elliot's Definition of Bioinformatics

- Computer-aided analysis of biological information
  
- Caveat:
  - In the end, bioinformatics (a.k.a. computers) can only help in making inferences concerning biological processes.
  - These inferences (or hypotheses) have to be tested in the laboratory



# Why Bioinformatics?

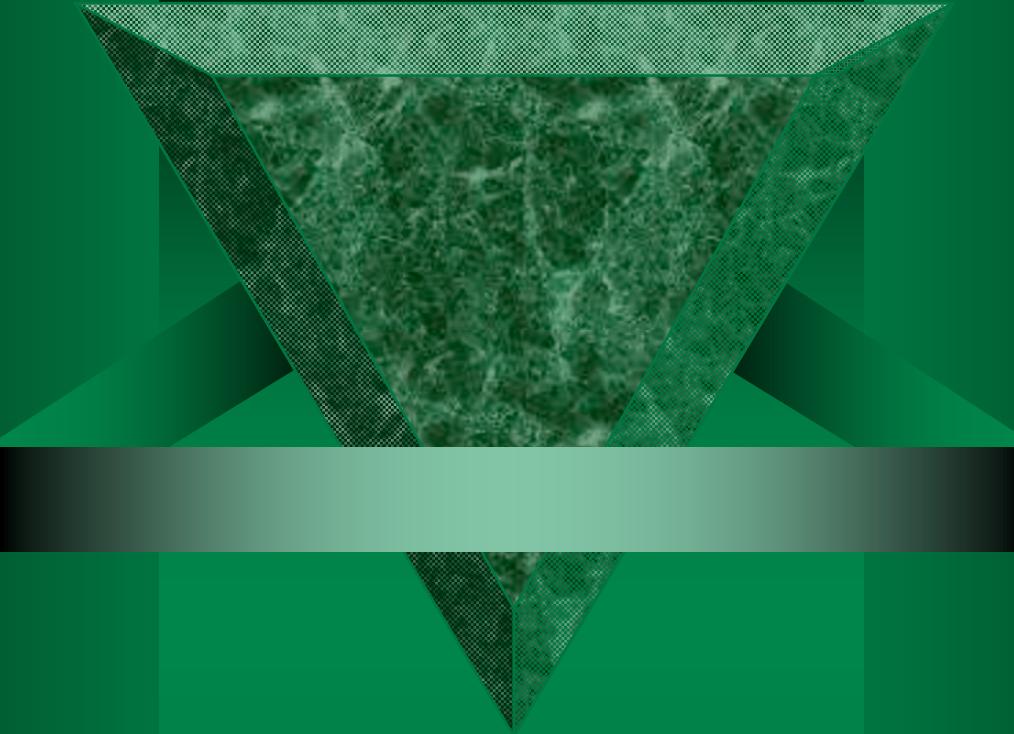
- Assists in the understanding of the basic genetic components of living systems.
- Understanding Genetics
  - Sequence
  - Genetic organization
  - Comparative analysis
  - Variation
- Structure Prediction
- Functional Inference
  - Enzymatic/Regulatory/Structural
  - Expression
  - Protein interactions



# Why Bioinformatics Now?

## ➤ Data overload

- Genomics
- Gene expression arrays
- Proteomics
- Structural biology
- Future biological revolutions



# Genomics



# The Human Genome Project

- Mapping and Sequencing the Genomes of Model Organisms
- Data Collection and Distribution
- Ethical, Legal, and Social Considerations
- Research Training
- Technology Development
- Technology Transfer



# Genomes of Humans and their “cousins”

- Eukaryotic
- Prokaryotic
- Archaea
- Viruses

# Human Genome Sequencing as of 1/5/2003

95.8% finished; chromosomes 6, 7, 13, 20, 21, 22 and Y are considered complete

Chr.	Tiling Path	Sequenced Clones (%)		
		Clones	Draft	Finished
1	2202	71 (3.2)	2089	(94.9)
2	1966	14 (0.7)	1948	(99.1)
3	1740	82 (4.7)	1630	(93.7)
4	1629	47 (2.9)	1572	(96.5)
5	1775	54 (3.0)	1700	(95.8)
6	1795	2 (0.1)	1790	(99.7)
7	1526	1 (0.1)	1514	(99.2)
8	1239	79 (6.4)	1133	(91.4)
9	999	22 (2.2)	963	(96.4)
10	1132	6 (0.5)	1117	(98.7)
11	1147	43 (3.7)	1088	(94.9)
12	1140	74 (6.5)	1044	(91.6)
13	854	0 (0.0)	854	(100.0)
14	655	1 (0.2)	642	(98.0)
15	710	68 (9.6)	618	(87.0)
16	725	44 (6.1)	672	(92.7)
17	692	132 (19.1)	541	(78.2)
18	600	6 (1.0)	585	(97.5)
19	861	9 (1.0)	852	(99.0)
20	632	0 (0.0)	632	(100.0)
21	473	0 (0.0)	472	(99.8)
22	527	0 (0.0)	527	(100.0)
X	1588	37 (2.3)	1508	(95.0)
Y	200	0 (0.0)	200	(100.0)
total	26807	792 (3.0)	25691	(95.8)



# Genome Project Organization

- Cloning
- Mapping
- Sequencing
- Annotation
- Analysis



# Bioinformatic Information Flow

- “Raw” data generation
  - Sequence generation and assembly
- Analytical tools
  - Pattern matching
- Database generation
  - Construction and data import
- Visualization (publication) of results
  - Static: Table or graph
  - Dynamic: Web page/Java applet



# Annotation and Analysis

## ➤ Gene prediction

- Identify patterns characteristic of ORFs

## ➤ Functional assignment

- Similarity searching

## ➤ Metabolic pathway modeling

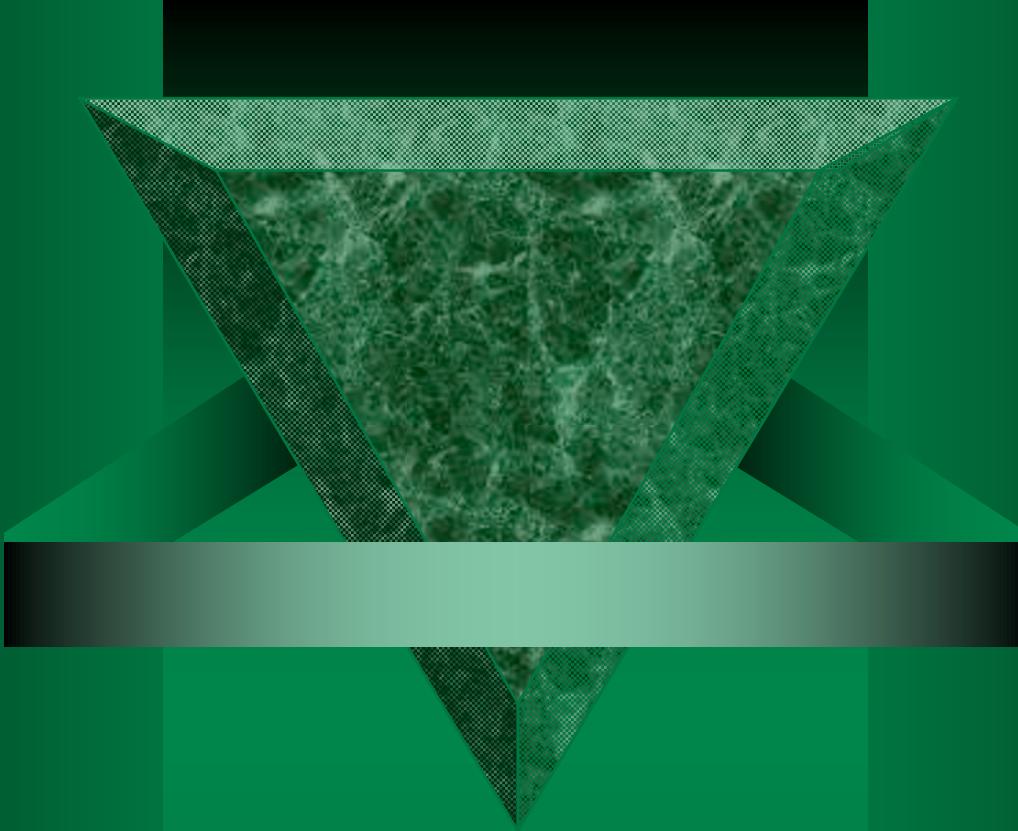
## ➤ Comparative analysis

- Identification and comparison with related genes



# Comparative Genomics

- “Similar” sequences
  - Sequences related by primary sequence similarity
- Homologs
  - Sequences related by evolution
  - Orthologs
    - Related due to speciation
  - Paralogs
    - Related due to gene duplication



# Bioinformatic Databases

Something to compare against



# Sequence Databases

## ➤ DNA

- Genbank (NCBI)
- EMBL
- DDBJ

## ➤ Protein

- PIR
- Swiss-Prot
- Swiss-Prot TrEMBL
- PDB



# Other Databases

## ➤ Structural

- Protein Data Bank (PDB): <http://www.rcsb.org/pdb/>

## ➤ Expression

- Microarray Gene Expression Data Society (MGED):  
<http://www.mged.org/>
- Gene Expression Omnibus (GEO – NCBI)

## ➤ Proteomic

- Mascot: <http://www.matrixscience.com/>

## ➤ Metabolism

- BioCyc: <http://biocyc.org/>

## ➤ Ontology

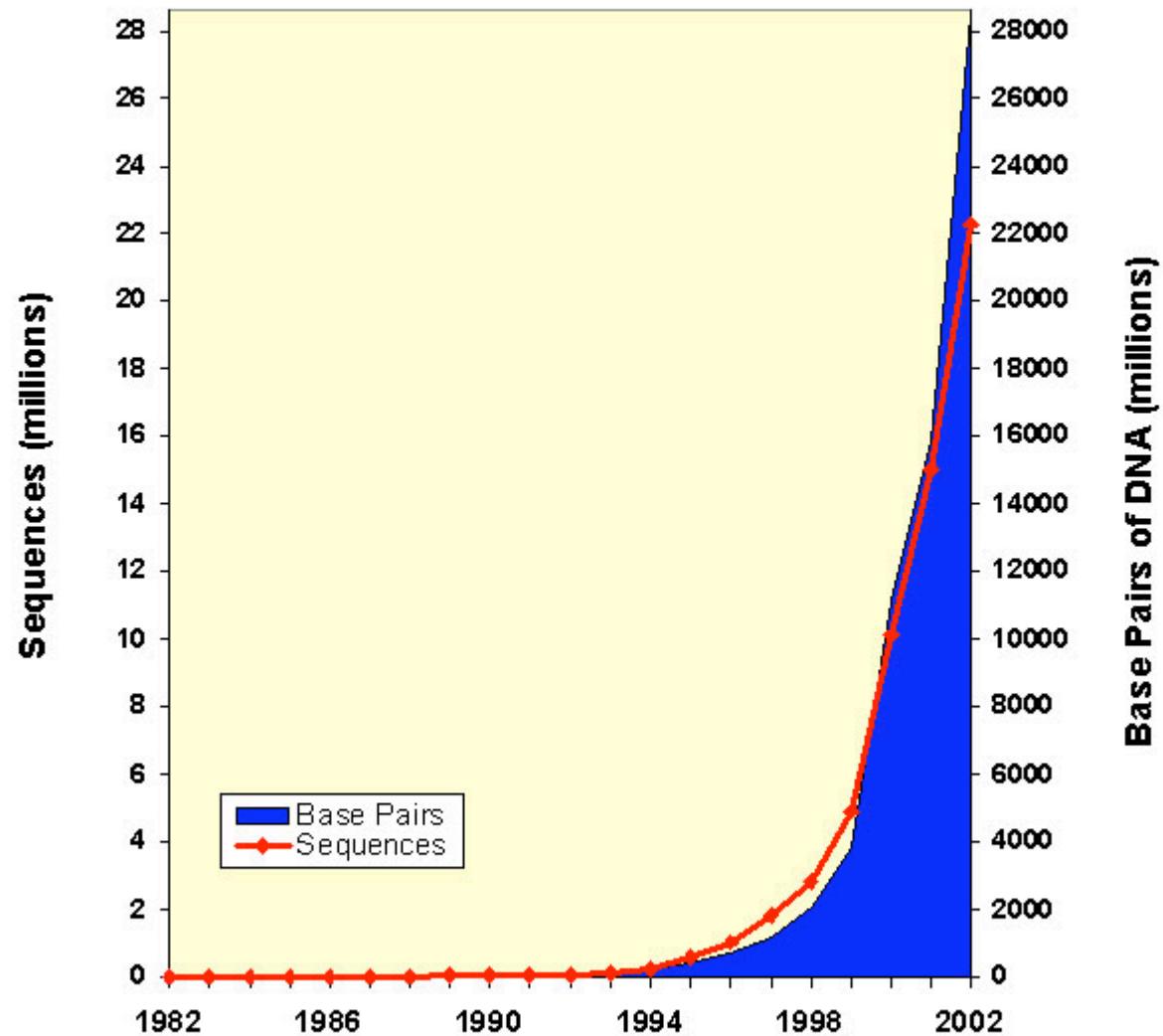
- Gene Ontology (GO) Consortium: <http://www.geneontology.org/>



# Genbank

- Primary nucleic acid sequence database
- Maintained by NCBI
  - National Center for Biotechnology Information
  - <http://www.ncbi.nlm.nih.gov>
- February 15, 2003; Release 134
  - 29,358,082,791 bases
  - 23,035,823 sequences

## Growth of GenBank





# Genbank Divisions

1. PRI - primate sequences
2. ROD - rodent sequences
3. MAM - other mammalian sequences
4. VRT - other vertebrate sequences
5. INV - invertebrate sequences
6. PLN - plant, fungal, and algal sequences
7. BCT - bacterial sequences
8. VRL - viral sequences
9. PHG - bacteriophage sequences
10. SYN - synthetic sequences
11. UNA - unannotated sequences
12. EST - EST sequences (expressed sequence tags)
13. PAT - patent sequences
14. STS - STS sequences (sequence tagged sites)
15. GSS - GSS sequences (genome survey sequences)
16. HTG - HTGS sequences (high throughput genomic sequences)
17. HTC - unfinished high-throughput cDNA sequencing



# HTGS

## *High Throughput Genomic Sequences*

- ✓ ‘Unfinished’ DNA sequences generated by the high-throughput sequencing centers
- ✓ Phase 0
  - Single-few pass reads of a single clone (not contigs)
- ✓ Phase 1
  - Unfinished, may be unordered, unoriented contigs, with gaps
- ✓ Phase 2
  - Unfinished, ordered, oriented contigs, with or without gaps
- ✓ Phase 3
  - Primary division (Genbank)
  - Finished, no gaps (with or without annotations)



# EST

## ➤ Expressed Sequence Tags

- “Single-pass” cDNA sequences
- Generally representative of the 3’ ends of cDNAs
- More “full-length” ESTs now available

## ➤ NCBI also has a dbEST database

- Same content
- Different format



# STS

## ➤ Sequence Tagged Sites

- Sequence and mapping data
- Short genomic landmark sequences

## ➤ NCBI also has a dbSTS database

- Same content
- Different format



# GSS

- Genome Survey Sequences
- Similar to the EST division, except that its sequences are genomic in origin, rather than cDNA
  - Random “single pass read” genome survey sequences.
  - Cosmid/BAC/YAC end sequences
  - Exon trapped genomic sequences
  - alu PCR sequences
- NCBI also has a dbGSS database
  - Same content
  - Different format



# Other NCBI Databases

- RefSeq
- Unigene
- HomoloGene
- Genomic
- dbSNP



# RefSeq

- NCBI Reference Sequence project
- Provides reference sequence standards for the naturally occurring molecules from chromosomes to mRNAs to proteins
- Stable reference point for:
  - mutation analysis
  - gene expression studies
  - polymorphism discovery
- Accession numbers have two letters, an underscore, and six numbers
  - NM\_123456



# RefSeq...

## ➤ Curated RefSeq

- transcripts and proteins

## ➤ Genome Annotation

- contigs, transcripts, and proteins

## ➤ Complete Genomes

- genomes, chromosomes, and proteins



# Unigene

- ✓ GenBank sequences partitioned into a non-redundant set of gene-oriented clusters
  - Each UniGene cluster contains sequences that represent a unique gene, as well as related information such as the tissue types in which the gene has been expressed and map location.
- ✓ Includes EST and complete cDNA sequences
- ✓ Provides information on differentially-spliced transcripts

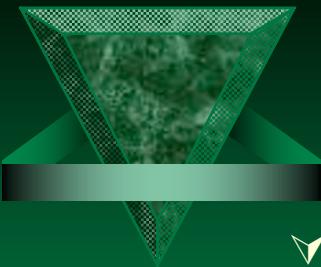
# Unigene Organisms

	Vertebrata	
	Mammalia	
	<a href="#"><i>Bos taurus</i></a> (cow)	12,808 entries
	<a href="#"><i>Homo sapiens</i></a> (human)	128,826 entries
	<a href="#"><i>Mus musculus</i></a> (mouse)	90,444 entries
	<a href="#"><i>Rattus norvegicus</i></a> (rat)	63,253 entries
	<a href="#"><i>Sus scrofa</i></a> (pig)	14,344 entries
	Aves	
	<a href="#"><i>Gallus gallus</i></a> (chicken)	5,068 entries
	Amphibia	
	<a href="#"><i>Xenopus laevis</i></a> (frog)	19,512 entries
	Actinopterygii	
	<a href="#"><i>Danio rerio</i></a> (zebrafish)	16,355 entries
	Urochordata	
	Asciidae	
	<a href="#"><i>Ciona intestinalis</i></a> (sea squirt)	13,674 entries
	Arthropoda	
	Insecta	
	<a href="#"><i>Anopheles gambiae</i></a> (malaria mosquito)	3,270 entries
	<a href="#"><i>Drosophila melanogaster</i></a> (fruit fly)	14,779 entries
	Nematoda	
	Chromadorea	
	<a href="#"><i>Caenorhabditis elegans</i></a>	20,137 entries
	Embryophyta	
	Eudicotyledons	
	<a href="#"><i>Arabidopsis thaliana</i></a> (thale cress)	27,141 entries
	<a href="#"><i>Glycine max</i></a> (soybean)	8,987 entries
	<a href="#"><i>Lycopersicon esculentum</i></a> (tomato)	3,740 entries
	<a href="#"><i>Medicago truncatula</i></a> (barrel medic)	5,729 entries
	Liliopsida	
	<a href="#"><i>Hordeum vulgare</i></a> (barley)	7,944 entries
	<a href="#"><i>Oryza sativa</i></a> (rice)	19,223 entries
	<a href="#"><i>Triticum aestivum</i></a> (wheat)	20,454 entries
	<a href="#"><i>Zea mays</i></a> (maize)	13,512 entries
	Chlorophyta	
	Chlorophyceae	
	<a href="#"><i>Chlamydomonas reinhardtii</i></a>	6,448 entries



# HomoloGene

- ✓ Curated and calculated orthologs and homologs for genes represented in UniGene and LocusLink. Organisms include:
  - *Arabidopsis thaliana*
  - *Bos taurus*
  - *Caenorhabditis elegans*
  - *Danio rerio*
  - *Drosophila melanogaster*
  - *Homo sapiens*
  - *Hordeum vulgare*
  - *Lycopersicon esculentum*
  - *Medicago truncatula*
  - *Mus musculus*
  - *Oryza sativa*
  - *Rattus norvegicus*
  - *Sus scrofa*
  - *Triticum aestivum*
  - *Xenopus laevis*
  - *Zea mays*



# Genomic DBs

- ✓ Human
- ✓ Mouse
- ✓ Rat
- ✓ Zebrafish
- ✓ Drosophila
- ✓ Nematode
- ✓ Plant genomes
- ✓ Yeast
- ✓ Malaria
- ✓ Microbial genomes
- ✓ Viruses
- ✓ Viroids
- ✓ Plasmids
- ✓ Eukaryotic organelles



# dbSNP

## ➤ Single Nucleotide Polymorphisms

- Single base changes
- Small-scale insertions/deletions
- Polymorphic repetitive elements
- Microsatellite variation



# LocusLink

- Provides a single query interface to curated sequence and descriptive information about genetic loci
  - Nomenclature
  - Aliases
  - Sequence accessions
  - Phenotypes
  - EC numbers
  - MIM numbers
  - UniGene clusters
  - Homology
  - Map locations
  - Web sites



# OMIM

- ✓ Online Mendelian Inheritance in Man
- ✓ Database of gene-linked genetic disorders
- ✓ Maintained at Johns Hopkins University
  - Dr. Victor A. McKusick



# Sample OMIM Queries (From the OMIM Help Docs)

- ✓ What human genes are related to hypertension? Which of those genes are on chromosome 17?
- ✓ List the OMIM entries that describe genes on chromosome 10.
- ✓ List the OMIM entries that contain information about allelic variants.
- ✓ Retrieve the OMIM record for the cystic fibrosis transmembrane conductance regulator (CFTR), and link to related protein sequence records via Entrez.
- ✓ Find the OMIM record for the p53 tumor protein, and link out to related information in LocusLink and the p53 Mutation Database.



# EMBL and DDBJ

- European Molecular Biology Laboratory
  - Hinxton, UK
  - <http://www.ebi.ac.uk/>
- DNA Data Bank of Japan
  - Mishima, Japan
  - <http://www.ddbj.nig.ac.jp/>



## Coordination with Genbank

- Prevents duplication
- Genbank enters sequences from U.S. journals and researchers
- EMBL handles European data
- DDBJ handles Asian data
- Data exchanged daily



# Accession Numbers

- Each sequence submitted to a database is assigned a unique primary accession number
- Accession numbers do not change
- If a sequence is merged with another, a new accession number is assigned, and the original number becomes a secondary accession number
- Accession numbers may include version numbers
  - AO2428.2



# The Sequence Record

- Different for each database
- Locus (Name)
- Accession Number
- Keywords
- Description
- Properties
- References
- The Sequence



# GenBank Sample Record

✓ <http://www.ncbi.nlm.nih.gov/Sitemap/samplerecord.html>

analyze% typedata ge:humcftrm  
!!NA\_SEQUENCE 1.0

LOCUS HUMCFTRM 6129 bp mRNA PRI 15-DEC-1989

DEFINITION Human cystic fibrosis mRNA, encoding a presumed transmembrane conductance regulator (CFTR).

ACCESSION M28668

NID g180331

KEYWORDS cystic fibrosis; transmembrane conductance regulator.

SOURCE Human, cDNA to mRNA.

ORGANISM Homo sapiens  
Eukaryotae; mitochondrial eukaryotes; Metazoa; Chordata;  
Vertebrata; Eutheria; Primates; Catarrhini; Hominidae; Homo.

REFERENCE 1 (bases 1 to 6129)

AUTHORS Riordan, J.R., Rommens, J.M., Kerem, B., Alon, N., Rozmahel, R., Grzelczak, Z., Zielenski, J., Lok, S., Plavsic, N., Chou, J.-L., Drumm, M.L., Iannuzzi, M.C., Collins, F.S. and Tsui, L.-C.

TITLE Identification of the cystic fibrosis gene: Cloning and characterization of complementary DNA

JOURNAL Science 245, 1066-1073 (1989)

MEDLINE 89368940

COMMENT A three base-pair deletion spanning positions 1654-1656 is observed in cDNAs from cystic fibrosis patients.

FEATURES Location/Qualifiers

source 1..6129  
/organism="Homo sapiens"  
/db\_xref="taxon:9606"

CDS 133..4575  
/note="cystic fibrosis transmembrane conductance regulator"  
/codon\_start=1  
/db\_xref="PID:g180332"  
/translation="MQRSPLEKASVVSKLFFSWTRPILRKGYRQRLELSDIYQIPSVD  
SADNLSEKLEREWDRRELASKKNPKLINALRRCCFWRFMFYGIFLYLGEVTKA  
VQPLL  
LNRFSSKDIAILDDLLPLTIFDFIQLLLIVIGAIAVVAVLQPYIFVATVPV  
VAFIMLR  
AYFLQTSQQLKQLESEGRSPIFTHLVTSLKGLWTLRAFGRQPYFETLFHK  
KALNLHTAN  
WFLYLSTLRWFQMRIEMIFVIFFIAVTFISILTTGEGEGRVGIIIL  
TLAMNIMSTLQWA  
VNSSIDVDLSLMRSVSRVFKFIDMPTEGKPTKSTKPYKNGQLSKVMII  
IENSHVKKDDIW  
PSGGQMTVKDLTAKYTEGGNAILENISFSISPGQRVGLLGRTGSGK  
STLLSAFLRLLN  
TEGEIQIDGVSWDSITLQQWRKAFGVIPQKVFIFSGTFRKNLD  
PYEQWSDQE  
EIWKVAD  
EVGLRSVIEQFPGKLD  
FVLVDGGCVLSHGHKQLMCLARSVLSKAKILLDEPSAHLD  
PYQIIRRTL  
KQAFADCTVILCEHRIEAMLECQQFLVIEENKVRQYDSI  
QKLLNERSL  
FRQAISPSDRV  
KLFPHRNSSKCKSKPQIAALKEETEEEVQDTRL"

BASE COUNT 1886 a 1181 c 1330 g 1732 t

ORIGIN

HUMCFTRM Length: 6129 April 13, 1998 13:00 Type: N Check: 6781 ..

1 AATTGGAAGC AAATGACATC ACAGCAGGTC AGAGAAAAAG GGTTGAGCGG  
51 CAGGCACCCA GAGTAGTAGG TCTTGCGAT TAGGAGCTTG AGCCCAGACG  
101 GCCCTAGCAG GGACCCCCAGC GCCCGAGAGA CCATGCAGAG GTCGCCTCTG  
151 GAAAAGGCCA GCGTTGTCTC CAAACTTTT TTCAGCTGGA CCAGACCAAT  
201 TTTGAGGAAA GGATACAGAC AGCGCCTGGA ATTGTCAGAC ATATAACCAA  
251 TCCCTTCTGT TGATTCTGCT GACAATCTAT CTGAAAAATT GGAAAGAGAA  
301 TGGGATAGAG AGCTGGCTTC AAAGAAAAAT CCTAAACTCA TTAATGCCCT  
351 TCGGCGATGT TTTTCTGGA GATTTATGTT CTATGGAATC TTTTTATATT  
401 TAGGGGAAGT CACCAAAGCA GTACAGCCTC TCTTACTGGG AAGAACATA  
451 GCTTCCTATG ACCCGGATAA CAAGGAGGAA CGCTCTATCG CGATTATCT

```
analyze% typedata -ref GB_PR:HUMIFNRF1A
```

!!NA\_SEQUENCE 1.0  
LOCUS HUMIFNRF1A 7721 bp DNA PRI 10-NOV-1992  
DEFINITION Homo sapiens interferon regulatory factor 1 gene, complete  
cds.  
ACCESSION L05072  
NID g184648  
KEYWORDS interferon regulatory factor 1.  
SOURCE Homo sapiens Placenta DNA.  
ORGANISM Homo sapiens  
Eukaryotae; mitochondrial eukaryotes; Metazoa; Chordata;  
Vertebrata; Eutheria; Primates; Catarrhini; Hominidae; Homo.  
REFERENCE 1 (bases 1 to 7721)  
AUTHORS Cha, Y., Sims, S.H., Romine, M.F., Kaufmann, M. and  
Deisseroth, A.B.  
TITLE Human interferon regulatory factor 1: intron/exon organization  
JOURNAL DNA Cell Biol. 11, 605-611 (1992)  
MEDLINE 93000481

FEATURES Location/Qualifiers

source 1. .7721  
/organism="Homo sapiens"  
/db\_xref="taxon:9606"  
/tissue\_type="Placenta"  
/map="5q23-q31"

exon 1. .219  
/gene="IRF1"  
/note="putative"  
/number=1

5'UTR join(1. .219,1279. .1287)  
/gene="IRF1"

gene join(1. .219,1279. .1287)  
/gene="IRF1"

intron 220. .1278  
/gene="IRF1"  
/number=1

exon 1279. .1374  
/gene="IRF1"  
/number=2

CDS join(1288. .1374,2738. .2837,3630. .3806,3916. .3965,  
4073. .4202,4386. .4508,5040. .5089,6248. .6383,6670.  
.6794)  
/gene="IRF1"  
/codon\_start=1  
/product="interferon regulatory factor 1"  
/db\_xref="PID:g184649"  
/translation="MPITRMRMRPWLEMQINSNQIPGLIWINKEEMIFQIPWKHAAKH  
GWDINKDACLFRSWAIHTGRYKAGEKEPDPKTWKANFRCAMNSLPDIEEVKDQSRNKG  
SSAVRVYRMLPPLTKNQRKERKSRSRDAKSRAKRGCGDSSPDTFSDGLSSSTLPDD  
HSSYTVPGYMQDLEVEQALTPALSPCAVSSTLPDWHIPVEVVPDSTSDFLYNFQVSPMP  
GTCGAATTGATTTTDEEECKLIDEDIMKILIGESEWHDPMNIVDCKCIVLILNEPDGVKPTSIWVCDPEGCKE"

```

intron    1375. .2737
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          /number=2
exon      2738. .2837
          /gene="IRF1"
          /number=3
intron    2838. .3629
          /gene="IRF1"
          /number=3
exon      3630. .3806
          /gene="IRF1"
          /number=4
intron    3807. .3915
          /gene="IRF1"
          /number=4
exon      3916. .3965
          /gene="IRF1"
          /number=5
intron    3966. .4072
          /gene="IRF1"
          /number=5

            ...
exon      5040. .5089
          /gene="IRF1"
          /number=8
intron    5090. .6247
          /gene="IRF1"
          /number=8
exon      6248. .6383
          /gene="IRF1"
          /number=9
intron    6384. .6669
          /gene="IRF1"
          /number=9
exon      6670. .7656
          /gene="IRF1"
          /number=10
3'UTR     6795. .7656
BASE COUNT 1750 a    1946 c    2253 g    1772 t
ORIGIN

```

```
analyze% typedata -ref est:hum091226f
!!NA_SEQUENCE 1.0
LOCUS      HUM091226F    152 bp     mRNA          EST        02-APR-1996
DEFINITION Homo sapiens retinal fovea EST HFV091226 sequence.
ACCESSION  L48850
NID        g1254959
KEYWORDS   EST; expressed sequence tag.
SOURCE     Homo sapiens (clone: EST HFV091226) age normalized retinal foveae
           cDNA to mRNA.
ORGANISM   Homo sapiens
           Eukaryotae; mitochondrial eukaryotes; Metazoa; Chordata;
           Vertebrata; Eutheria; Primates; Catarrhini; Hominidae; Homo.
REFERENCE  1 (sites)
AUTHORS   Adams,M.D., Kerlavage,A.R., Fields,C. and Venter,J.C.
TITLE     3,400 new expressed sequence tags identify diversity of transcripts
           in human brain
JOURNAL   Nature Genet. 4 (3), 256-267 (1993)
MEDLINE   93364420
REFERENCE 2 (sites)
AUTHORS   Liew,C.C., Hwang,D.M., Fung,Y.W., Laurensen,C., Cukerman,E.,
           Tsui,S. and Lee,C.Y.
TITLE     A catalogue of genes in the cardiovascular system as identified by
           expressed sequence tags
JOURNAL   Proc. Natl. Acad. Sci. U.S.A. 91 (22), 10645-10649 (1994)
MEDLINE   95024171
REFERENCE 3 (bases 1 to 152)
AUTHORS   Bernstein,S.L., Borst,D.E., Neuder,M.E. and Wong,P.
TITLE     Characterization of a human fovea cDNA library and regional
           differential gene expression in the human retina
JOURNAL   Genomics 32 (3), 301-308 (1996)
```

FEATURES Location/Qualifiers

source 1. .152  
/organism="Homo sapiens"  
/note="Expressed sequence tags (first pass sequencing)  
from randomly selected bacteriophage clones (mRNA-cDNA)  
from human retinal fovea. The library is age normalized  
from ten sets of donor foveae 2-79 years old.  
/db\_xref="taxon:9606"  
/clone="EST HFV091226"  
/dev\_stage="age normalized"  
/tissue\_type="retinal foveae"

mRNA <1. .>152  
/standard\_name="EST HFV091226"

BASE COUNT 31 a 42 c 41 g 36 t 2 others

ORIGIN

```
analyze% typedata -ref sts:humswx153
! !NA_SEQUENCE 1.0
LOCUS          HUMSWX153      192 bp       DNA          STS        24-MAY-
1993
DEFINITION    Human chromosome X STS sWXD153; single read.
ACCESSION     L15212
NID           g292645
KEYWORDS      STS; primer; sequence tagged site.
SOURCE         Homo sapiens DNA.
ORGANISM       Homo sapiens
               Eukaryotae; mitochondrial eukaryotes; Metazoa; Chordata;
               Vertebrata; Eutheria; Primates; Catarrhini; Hominidae; Homo.
REFERENCE     1 (bases 1 to 192)
AUTHORS       Kere,J., Nagaraja,R., Mumm,S.R., Ciccodicola,A., D'Urso,M.
and
               Schlessinger,D.
TITLE          Mapping human chromosomes by walking with sequence-tagged
sites
               from end fragments of yeast artificial chromosome inserts
JOURNAL        Genomics 14, 241-248 (1992)
MEDLINE        93052321
```

COMMENT Submitted by: David Schlessinger,  
Center for Genetics in Medicine,  
Washington University School of Medicine, Box 8232 4566 Scott  
Avenue, St. Louis, MO 63110, USA  
e-mail: davids@wugenmail.wustl.edu  
Primer A: TAAAGGGATGCCAAGGAC  
Primer B: CTTACTCATTGCTGGATTCTC  
STS size: 85bp  
Template: 600 ng/100ul  
Primer: 40 pmoles/100ul  
dNTPs: 100 uM  
MgCl<sub>2</sub>: 1.5 mM  
KCl: 100 mM  
TrisHCl: 10 mM  
Taq Polymerase: 0.125 U  
NH<sub>4</sub>Cl: 5 mM  
pH: 8.6  
Total Vol: 5 ul  
PCR Profile:  
Denaturation: 94 degrees C for 1.00 minute(s)  
Annealing: 55 degrees C for 2.00 minute(s)  
Polymerization: 72 degrees C for 2.00 minute(s)  
PCR Cycles: 35  
Thermal Cycler: P-E.

FEATURES Location/Qualifiers

source 1. .192  
/organism="Homo sapiens"  
/db\_xref="taxon:9606"  
/map="Xq13-q24"

STS 60. .144  
/standard\_name="sWXD153"

primer\_bind 60. .78

primer\_bind complement(123. .144)

BASE COUNT 72 a 26 c 60 g 29 t 5 others

ORIGIN

analyze%



# Entrez Searching

- <http://www.ncbi.nlm.nih.gov/entrez/>
- Search via text patterns
- Access to all of NCBI's databases
  - Sequence
  - PubMed
  - OMIM
  - Linkage information



# Swiss-Prot

- <http://www.expasy.ch/sprot/>
- Protein Database
- University of Geneva
- Arranged by protein function
- Release 40.44
- February 22, 2003
- 44,864,044 amino acids 122,214 entries
- Provides annotated protein records



# Swiss-Prot TrEMBL

- Translation of all EMBL Nucleic Acid coding sequences not yet present in Swiss-Prot
- Allows rapid availability without immediate annotation
- Release 21.13
- February 14, 2003
- 725,373 entries



# National Biomedical Research Foundation

- Database begun over twenty years ago by Margaret O. Dayhoff
- Originally published sequences in book form
- Started with sequences derived from direct amino acid sequencing



# PIR

- <http://pir.georgetown.edu/>
- Protein Identification Resource
  - PIR-International Protein Sequence Database (PSD)
- National Biomedical Research Foundation
- Georgetown University
- Release 75.04, March 03, 2003
- 283,290 Entries



# PIR-NREF

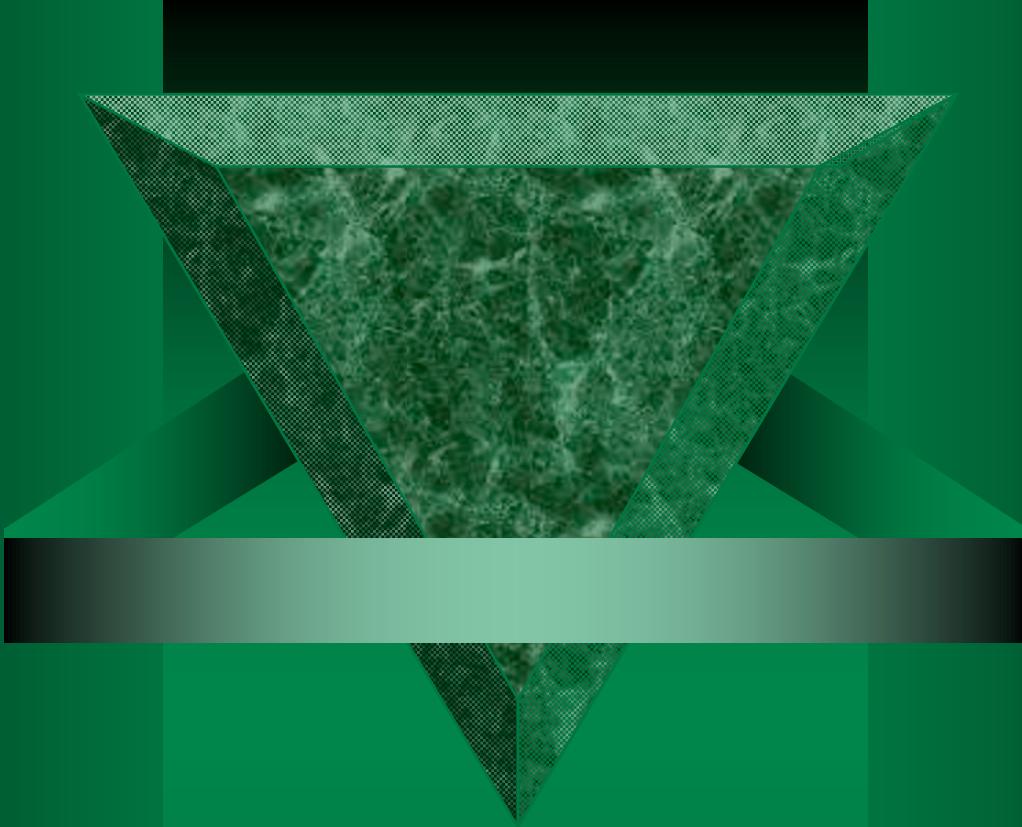
- Non-redundant REFERENCE protein database
- Current Release 1.17
- March 3, 2003
- 1,159,203 Entries



# iProClass Database - PIR

<http://pir.georgetown.edu/iproclass/>

- Comprehensive family relationships and structural/functional classifications and features of proteins
  - Superfamilies
  - Families
  - Domains



So I have a protein. What next?



# NCBI Sequence Services

- Obtain sequences directly from NCBI using Entrez
  - Sequence Searches
  - Sequence Retrieval
- Other services
  - BLAST Searches
  - Sequence Submission
  - PubMed Searches
- <http://www.ncbi.nlm.nih.gov>



# Sequence Similarities

- What other sequences have some primary sequence similarity to my query sequence?
- Time and cost of the search is dependent on the size of the database
  - Restrict the size of the database



# BLAST

- ✓ Search a sequence database for primary sequence similarities to some query sequence
- ✓ Provides a measure of the significance of the similarity
- ✓ Does not necessarily imply common evolutionary origin



# BLAST

- All search combinations possible
- nt vs. nt database
  - blastn
- protein vs. protein database
  - blastp
- translated nt vs. protein database
  - blastx
- protein vs. translated nt database
  - tblastn
- translated nt vs. translated nt database
  - tblastx



# A Few Genome Resources

## ➤ NCBI Genome Resources

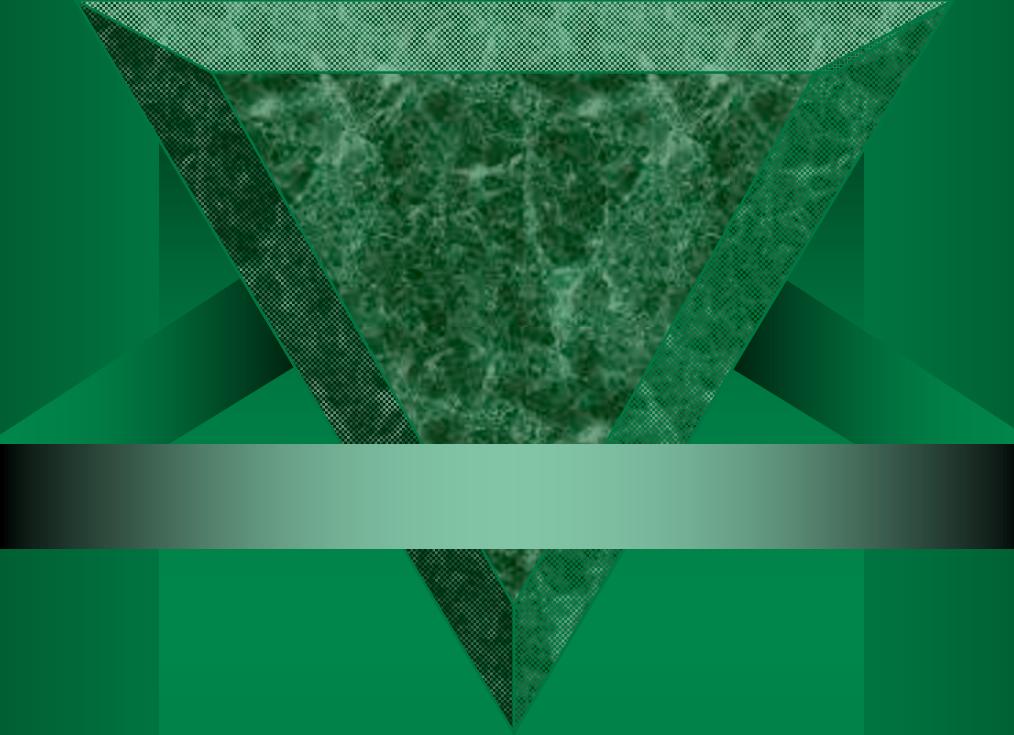
- National Center for Biotechnology Information
- [www.ncbi.nih.gov](http://www.ncbi.nih.gov)

## ➤ Ensembl Human Genome Server

- [www.ensembl.org](http://www.ensembl.org)

## ➤ UCSC Human Genome Browser

- [genome.ucsc.edu](http://genome.ucsc.edu)



# Analysis of PnP

Human Purine nucleoside phosphorylase

NCBI Sequence Viewer - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back Forward Stop Home Search Favorites Media Print Copy Paste Find Links

Address http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list\_uids=130377&dopt=GenPept Go Links »

NCBI Entrez Protein

PubMed Nucleotide Protein Genome Structure PMC Taxonomy OMIM Books

Search **Protein** for  Limits Preview/Index History Clipboard Details

Display default Show: 20 File

**□ 1: P00491. Purine nucleoside...[gi:130377]**

LOCUS P00491 289 aa linear PRI 15-JUN-2002  
DEFINITION Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP).  
ACCESSION P00491  
VERSION P00491 GI:130377  
DBSOURCE swissprot: locus PNPH\_HUMAN, accession P00491;  
class: standard.  
extra accessions:Q15160,created: Jul 21, 1986.  
sequence updated: Jul 21, 1986.  
annotation updated: Jun 15, 2002.  
xrefs: gi: [35564](#), gi: [35565](#), gi: [190150](#), gi: [387033](#), gi: [190147](#),  
gi: [190148](#), gi: [190149](#), gi: [66583](#), gi: [230387](#), gi: [230388](#)  
xrefs (non-sequence databases): Aarhus/Ghent-2DPAGE2108, MIM  
[164050](#), InterProIPR001369, PfamPF00896, PROSITEPS01240  
KEYWORDS Transferase; Glycosyltransferase; Polymorphism; Disease mutation;  
3D-structure.  
SOURCE Homo sapiens (human)  
ORGANISM [Homo sapiens](#)  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.  
REFERENCE 1 (residues 1 to 289)  
AUTHORS Williams,S.R., Goddard,J.M. and Martin,D.W. Jr.  
TITLE Human purine nucleoside phosphorylase cDNA sequence and genomic  
clone characterization  
JOURNAL Nucleic Acids Res. 12 (14), 5779-5787 (1984)  
MEDLINE [84272252](#)  
PUBMED [6087295](#)  
REMARK SEQUENCE FROM N.A.  
REFERENCE 2 (residues 1 to 289)  
AUTHORS Williams,S.R., Gekeler,V., McIvor,R.S. and Martin,D.W. Jr.  
TITLE A human purine nucleoside phosphorylase deficiency caused by a

Done

NCBI Sequence Viewer - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Address http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list\_uids Go Links »

FEATURES	Location/Qualifiers
source	1..289 /organism="Homo sapiens" /db_xref="taxon:9606"
<u>gene</u>	1..289 /gene="NP" /note="synonym: PNP"
<u>Protein</u>	1..289 /gene="NP" /product="Purine nucleoside phosphorylase" /EC_number=" <a href="#">2.4.2.1</a> "
<u>Region</u>	7 /gene="NP"
<u>Region</u>	8..17 /gene="NP" /region_name="Hydrogen bonded turn"
<u>Region</u>	18..20 /gene="NP" /region_name="Helical region"
<u>Region</u>	27..31 /gene="NP" /region_name="Hydrogen bonded turn"
<u>Region</u>	36..41 /gene="NP" /region_name="Beta-strand region"
<u>Region</u>	46..48 /gene="NP" /region_name="Helical region"
<u>Region</u>	50..52 /gene="NP" /region_name="Beta-strand region"
<u>Region</u>	51 /gene="NP" /region_name="Helical region"
<u>Region</u>	54..55 /note="S -> G (IN DBSNP:1049564). /FTId=VAR_002243." /gene="NP"

NCBI Sequence Viewer - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Address [http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list\\_uids=](http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list_uids=) Go Links »

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/gene="NP"
/region_name="Helical region"
Region 231..232
/gene="NP"
/region_name="Hydrogen bonded turn"
Region 234..244
/gene="NP"
/region_name="Beta-strand region"
Region 234
/gene="NP"
/region_name="Variant"
/note="R -> P (IN NP DEFICIENCY). /FTId=VAR_002247."
Region 263..278
/gene="NP"
/region_name="Helical region"
Region 279..281
/gene="NP"
/region_name="Helical region"

ORIGIN
1 mengytyedy kntaeawlsh tkhrpqvaii cgsgllg ltd kltqaqifdy seipnfprst
61 vpghagrlvf gflngracvm mqgrfhmyeg yplwkvtfpv rvfhllgvdt lvvtnaagg1
121 npkfevgd im lirdhinlpg fsgqnplrgp nderfgdrfp amsdaydrtm rqralstwkq
181 mgeqrelqeg tyvmwagsf etvaecrvliq klgadavgm s tvpeviv arh cglrvfgfs1
241 itnkvimdy e slekanheev laagkqaaqk leqfv silma siplpdkas
//
```

Revised: August 5, 2002.

NCBI Sequence Viewer - Microsoft Internet Explorer

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Back Search Favorites Media Structure PMC Taxonomy OMIM Books

Address http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list\_uids=130377&dopt=GenPept Go Links

**NCBI** Entrez Protein

PubMed Nucleotide Protein Genome Structure PMC Taxonomy OMIM Books

Search Protein for

Limits Preview/Index History Clipboard Details

Display FASTA Show: 20 Send to File

**1: P004** ASN.1 Fasta...[gi:130377] [BLINK](#), [Domains](#), [Links](#)

LOCUS 289 aa linear PRI 15-JUN-2002  
DEFINITION cleoside phosphorylase (Inosine phosphorylase) (PNP).  
ACCESSION I:130377  
VERSION : locus PNPH\_HUMAN, accession P00491;  
DBSOURCE standard.  
EXTRA acccessions:Q15160,created: Jul 21, 1986.  
sequence updated: Jul 21, 1986.  
annotation updated: Jun 15, 2002.  
xrefs: gi: [35564](#), gi: [35565](#), gi: [190150](#), gi: [387033](#), gi: [190147](#),  
gi: [190148](#), gi: [190149](#), gi: [66583](#), gi: [230387](#), gi: [230388](#)  
xrefs (non-sequence databases): Aarhus/Ghent-2DPAGE2108, MIM  
[164050](#), InterProIPR001369, PfamPF00896, PROSITEPS01240  
KEYWORDS Transferase; Glycosyltransferase; Polymorphism; Disease mutation;  
3D-structure.  
SOURCE Homo sapiens (human)  
ORGANISM [Homo sapiens](#)  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.  
REFERENCE 1 (residues 1 to 289)  
AUTHORS Williams, S.R., Goddard, J.M. and Martin, D.W. Jr.

NCBI Sequence Viewer - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back Search Favorites Media Go Links

Address http://www.ncbi.nlm.nih.gov/entrez/viewer.fcgi?cmd=&txt=&save=&cfm=&query\_key=60&db=protein&extrafeat=-1&view=1 Go

NCBI Entrez Protein

PubMed Nucleotide Protein Genome Structure PMC Taxonomy OMIM Books

Search Protein for  Go Clear

Limits Preview/Index History Clipboard Details

Display FASTA Show: 20 Send to File Get Subsequence

1: P00491. Purine nucleoside...[gi:130377] [BLink](#), [Domains](#), [Links](#)

>gi|130377|sp|P00491|PNPH\_HUMAN Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP)  
MENGYTYEDYKNTAEWLLSHTKHRPQVAAIICGSGLGGLTDKLTQAQIFDYSEIPNFPRSTVPGHAGRLVF  
GFLNGRACVMMQGRFHMYEGYPLWKVTFPVRVFHLLGVDTLVVTNAAGGLNPKFEVGDIMLIRDHINLPG  
FSGQNPLRGPNDERFGDRFPAMSDAYDRTMRQRALSTWKQMGEQRELQEGTYVMVAGPSFETVAECRVLQ  
KLGADAVGMSTVPEVIVARHCGLRVFGFSLITNKVIMDYESLEKANHEEVLAAGKQAAQKLEQFVSILMA  
SIPLPDKAS

Revised: August 5, 2002.

[Disclaimer](#) | [Write to the Help Desk](#)  
[NCBI](#) | [NLM](#) | [NIH](#)

Feb 19 2003 14:49:48

Done Internet



# Searching Genomic Sequences

- Where is my sequence located in the human genome?
  - Chromosome; band; mapping data
  - Genetic linkage relationships
- What is the genomic context of my sequence?
  - Alternative splicing
  - Regulation
- Are there any paralogs?
- Are there any pseudogenes?
- Comparative analysis with the same gene in other genomes

BLAST the Human Genome - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back Search Favorites Media

Address http://www.ncbi.nlm.nih.gov/genome/seq/page.cgi?F=HsBlast.html&&ORG=Hs

NCBI | Genomic Biology | Human Genome Guide | Human Sequence

Search LocusLink for  Go

**BLAST the Human genome**

Compare your query sequence to the working draft sequence of the human genome or its mRNA and protein products.

Database:  Program:   
 use MegabLAST

Enter an accession, gi, or a sequence in FASTA format:

```
>PNP [Homo sapiens] gi|35565|emb|CAA25320.1|
MENGYTYEDYKNTAEWLLSHTKHRPQVAAICGSGLGGLTDKLTQAFIDYSEIPNFP
RSTVPGHAGRLVF
GFLNGRACVMMQGRFHMYEGYPLWKVTFPVRFHLLGVDTLVVTNAAGGLNPKEVGDIM
LIRDHINLPG
FSGQNPLRGPNDERFGDRFPAMS DAYDRTMRQRALSTWKQMGEQRELQEGTYVMVAGPSF
```

**Optional parameters**

**Expect** **Filter** **Descriptions** **Alignments**

0.01  default  100  100

Advanced options:

Internet

BLAST the Human Genome - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back Search Favorites Media

Address http://www.ncbi.nlm.nih.gov/genome/seq/page.cgi?F=HsBlast.html&&ORG=Hs Go Links

NCBI | Genomic Biology | Human Genome Guide | Human Sequence

Search LocusLink for  Go

**BLAST the Human genome**

Compare your query sequence to the working draft sequence of the human genome or its mRNA and protein products.

**Database:** genome **Program:** tblastn

use Me genome  
RNA  
PROTEIN  
gscan\_mrna  
gscan\_protein  
Enter an accession number or sequence...  
>PNP [Homo sapiens]  
MENGYTYE HTGS  
VPGHAGRL ESTs  
GFLNGRAC EST Traces  
LIRDHINI Other Traces

ence in FASTA format:

```
65 |emb|CAA25320.1|  
VAIICGSGLGGLTDKLTQAQIFDYSEIPNFPQRST  
TFPVRFHLLGVDTLVVTNAAGGLNPKEVGDIM  
FSGQNPLRGPNDERFGDRFPAMSDAYDRTMRQRALSTWKQMGEQRELQEGTYVMVAGPSF
```

**Optional parameters**

**Expect** **Filter** **Descriptions** **Alignments**

0.01 default 100 100

Advanced options:

Begin Search Clear Input

Internet

BLAST the Human Genome - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back Search Favorites Media

Address http://www.ncbi.nlm.nih.gov/genome/seq/page.cgi?F=HsBlast.html&&ORG=Hs

NCBI | Genomic Biology | Human Genome Guide | Human Sequence

Search LocusLink for  Go

**BLAST the Human genome**

Compare your query sequence to the working draft sequence of the human genome or its mRNA and protein products.

Database:  Program:

use MegabLAST

Enter an accession, gi, or a sequence in FASTA format:

```
>PNP [Homo sapiens] gi|35565|emb|CAA25320.1|  
MENGYTYEDYKNTAEWLLSHTKHRPQVAIICGSGLGGLTDKLTQAFIDYSEIPNFP  
RSTVPGHAGRLVF  
GFLNGRACVMMQGRFHMYEGYPLWKVTFPVRFHLLGVDTLVVTNAAGGLNP  
KFEVGDIM  
LIRDHINLPG  
FSGQNPLRGPNDERFGDRFPAMS DAYDRTMRQRALSTWKQMGEQRELQEGTYVM  
VAGPSF
```

**Optional parameters**

Expect  Filter  Descriptions  Alignments

Advanced options:

Internet

NCBI Blast - Microsoft Internet Explorer

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Address http://www.ncbi.nlm.nih.gov/blast/Blast.cgi Go

**formatting BLAST**

Nucleotide Protein Translations Retrieve results for an RID

Your request has been successfully submitted and put into the Blast Queue.

Query = PNP [Homo sapiens] gi|35565|emb|CAA25320.1| (289 letters)

The request ID is 1046634425-024719-9809

**Format!** or **Reset all**

The results are estimated to be ready in 6 minutes but may be done sooner.

Please press "FORMAT!" when you wish to check your results. You may change the formatting options for your result via the form below and press "FORMAT!" again. You may also request results of a different search by entering any other valid request ID to see other recent jobs.

**Format**

Show  Graphical Overview  Linkout  Sequence Retrieval  NCBI-gi Alignment in HTML format

Number of: Descriptions 100 Alignments 100

Alignment view Pairwise

Limit results by entrez query  or select from: (none)

Expect value range:

Internet

 NCBI

## **results of BLAST**



TBLASTN 2.2.5 [Nov-16-2002]

### **Reference :**

Altschul, Stephen F., Thomas L. Madden, Alejandro A. Schäffer, Jinghui Zhang, Zheng Zhang, Webb Miller, and David J. Lipman (1997), "Gapped BLAST and PSI-BLAST: a new generation of protein database search programs", Nucleic Acids Res. 25:3389-3402.

RID: 1046634425-024719-9809

**Query=** PNP [Homo sapiens] gi|35565|emb|CAA25320.1|  
(289 letters)

**Database:** Homo sapiens genomic contig sequences  
953 sequences; 2,861,589,808 total letters

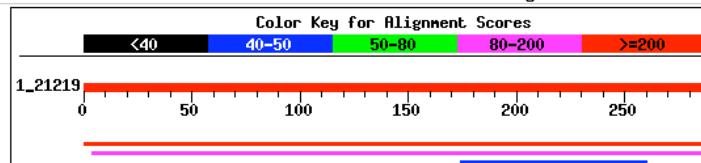
If you have any problems or questions with the results of this search please refer to the [BLAST FAQS](#)

Genome View

Show positions of the BLAST hits in the human genome using the Entrez Genomes MapViewer

## Distribution of 7 Blast Hits on the Query Sequence

Mouse-over to show defline and scores. Click to show alignments



RID=1046634425-024719-9809, PNP [Homo sapiens] gi|35565|emb|CAA25320.1 - Microsoft Internet Explorer

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Address http://www.ncbi.nlm.nih.gov/blast/Blast.cgi Go Links

Sequences producing significant alignments:

	Score	E
	(bits)	Value
ref NT_022184.10 Hs2_22340 Homo sapiens chromosome 2 genomic contig	327	5e-88
ref NT_037845.1 Hs14_37849 Homo sapiens chromosome 14 genomic contig	141	9e-62
ref NT_037734.1 Hs9_37738 Homo sapiens chromosome 9 genomic contig	47	0.001

**Alignments**

>ref|NT\_022184.10|Hs2\_22340 Homo sapiens chromosome 2 genomic contig  
Length = 13913408

Score = 327 bits (838), Expect = 5e-88  
Identities = 182/298 (61%), Positives = 211/298 (70%), Gaps = 9/298 (3%)  
Frame = -2

Query: 1 MENGTYEDYKNTAEWLLSHTKHRPQVAIICGSGLGGLTDKLTQAQIFDYSEIPNFPNST 60  
MENGTYEDY++TAEWLL HTKH QV +ICGS LG LTDKL QAQIF+ SE+ NF +ST  
Sbjct: 6971389 MENGTYEDYQSTAEWLLFHTKH\*TQVTVICGSELGDLTDKLIQAQIFNNSEMLNFFQST 6971210

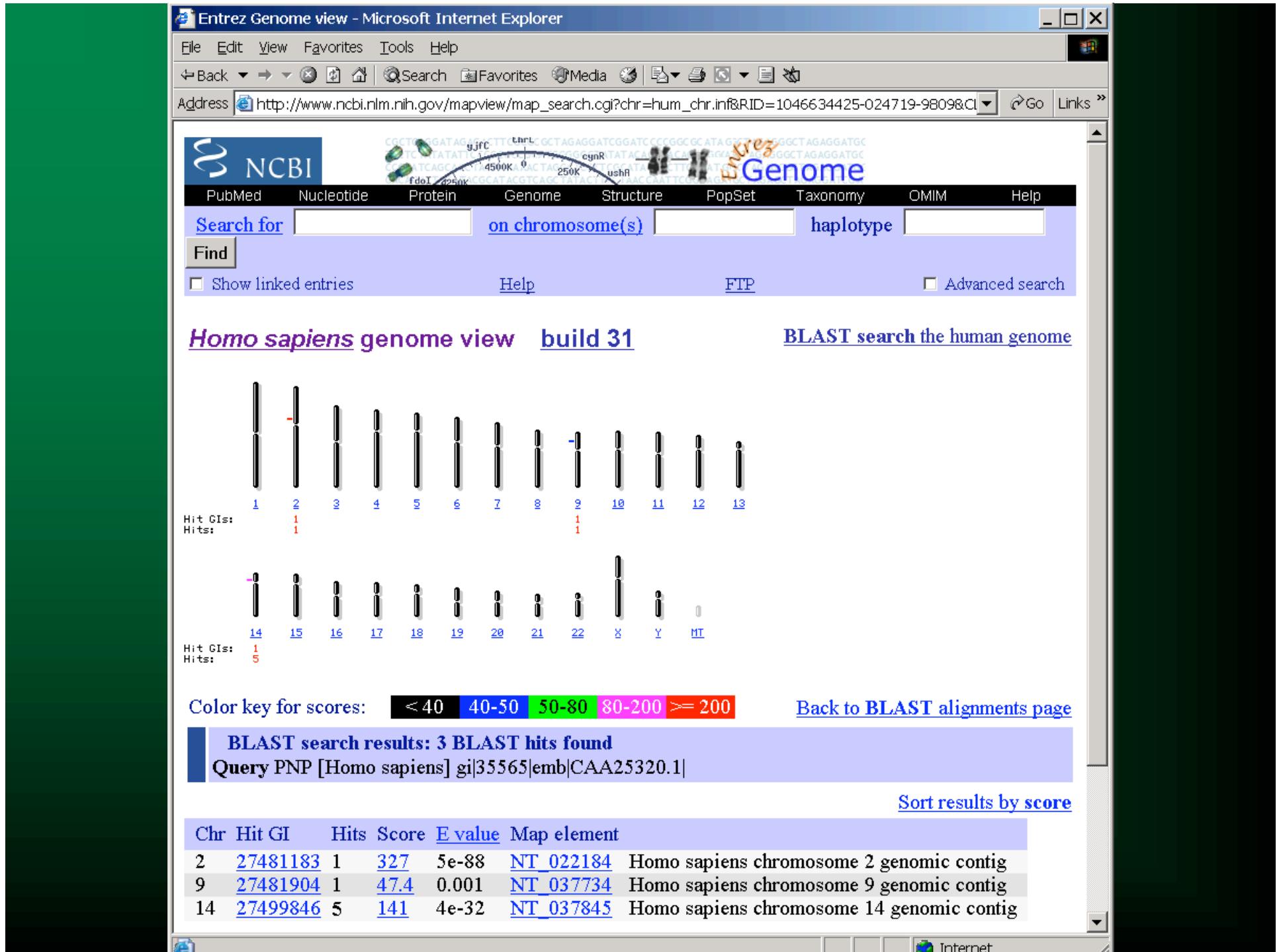
Query: 61 VPGHAGRLVFGFLNNGRACVMMQGRFHMYEGYPLWKVTFPVRFHLLGVDTLVVTNAAGGL 120  
VPGHA LVFGFLNG CVMMQGRF++Y+GY LW + F VF LLG + LV T+AAGGL  
Sbjct: 6971209 VPGHAV\*LVFGFLNGBTVCVMMQGRFYLYDGYLLWNMIFLHEVFQLLGGNILVATDAAGGL 6971030

Query: 121 NPKFEVGDIIMLIRDHINLPGFSQNPLRGPNDERFGDRFPAMSDAYDRTMRQRALSTWKQ 180  
NPK EVG IML+ DHI L GF QN +GPNDERFG FPA SDAY+ TM+Q+AL++ Q  
Sbjct: 6971029 NPKSEVGRIMLLCDHIKLLGFCDQNSPKGPNDERFGVHFATSDAYNWTMKQKALNS\*NQ 6970850

Query: 181 MGEQRELQEGTYVMVAGPSFETVAECRVLQKLGADAVGMSTVPEVIV--ARHCGLRVFG- 237  
MG+Q+E+Q+ TYVM +FET G D+ + A+H  
Sbjct: 6970849 MGKQQEVQKDTYVMAVNCNFET-----GRDSSDAEAGDGCCLA\*AQHQS\*SCMAL 6970700

Query: 238 -----FSLITNKVIMDYESLEKANHEEVLAAGKQAAQKLEQFVSILMASIPLPDKAS 289  
FSLITNKVIMDYESL+KANHE V A KQAAQKLEQFVSIL ASIPLPD A+  
Sbjct: 6970699 WTWSLCFSLITNKVIMDYESLKKANHE\*V\*EAVKQAAQKLEQFVSILKASIPLPDNAN 6970526

Done Internet



Ensembl Human Genome Browser (BlastView) - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back ▶ Search Favorites Media Links

Address [http://www.ensembl.org/Homo\\_sapiens/blastview?id=hs\\_s3a69Ev66Yg3&format=karyo\\_format](http://www.ensembl.org/Homo_sapiens/blastview?id=hs_s3a69Ev66Yg3&format=karyo_format) Go Links

Google Search Web Search Site News PageRank Page Info Up Highlight

**e!** Ensembl Human BLASTView The Wellcome Trust Sanger Institute EBI

Home ▶ Human ▶ What's New ▶ BLAST ▶ SSAHA ▶ EnsMart ▶ Export Data ▶ Download ▶ Disease Browser ▶ Docs ▶

Find All  [e.g. AP000462, RH9632, cancer]

Blast score ranges for this search: [ The highest scoring hit(s) are boxed ]

42 - 308    309 - 575    576 - 842

Location of Blast hits

The image shows a karyogram of the human genome. The chromosomes are arranged in two rows: the top row contains chromosomes 1 through 12, and the bottom row contains chromosomes 13 through 22, along with the X and Y chromosomes. Each chromosome is represented by a vertical bar with horizontal bands of different shades of gray and black, indicating genetic variation. A red rectangular box highlights a specific region on chromosome 2. Blue arrows point to specific bands on chromosomes 3, 9, 14, and 15. The chromosomes are numbered below them.

Human BLAT Search - Microsoft Internet Explorer

File Edit View Favorites Tools Help

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Address http://genome.ucsc.edu/cgi-bin/hgBlat?command=start&org=human

Google Search Web Search Site News PageRank Page Info Up Highlight

Home - Genome Browser - Blat Search - Table Browser - FAQ - User Guide

## Human BLAT Search

# BLAT Search Genome

Genome: Human Assembly: Human Nov. 2002 Query type: BLAT's guess Sort output: query,score Output type: hyperlink Submit

Please paste in a query sequence to search the genome. Multiple sequences can be searched at once if separated by new lines. If you enter a name.

BLAT's guess  
DNA  
protein  
translated RNA  
translated DNA

```
>PNP [Homo sapiens] gi|35565|emb|CAA25320.1|  
MENGYTYEDYKNTAEWLLSHTKHRPQVAIICGSGLGGLTDKLTQAQIFDYSEIPNFPRSTVPGHAGRLVF  
GFLNGRACVMMQGRFHMYEGYPLWKVTFPVRFHLLGVDTLVTNAAGGLNPFEVGDIIMLIRDHINLPG  
FSGQNPLRGPNDERFGDRFPAMSDAYDRTMRQRALSTWKQMGQRELQEGTYVMVAGPSFETVAECRVLQ  
KLGADAVGMSTVPEVIVARHGLRVFGSLITNKVIMDYESLEKANHEEVLAAGKQAAQKLEQFVSILMA  
SIPLPKAS
```

Rather than pasting a sequence, you can choose to upload a text file containing the sequence.

Upload sequence:  Browse... Submit File

Only DNA sequences of 25,000 or less bases and protein or translated sequence of 5000 or less letters will

Human BLAT Results - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Back Forward Stop Home Search Favorites Media | PageRank Page Info Up Highlight

Address http://genome.ucsc.edu/cgi-bin/hgBlat

Google Search Web Search Site News Page Rank Page Info Up Highlight

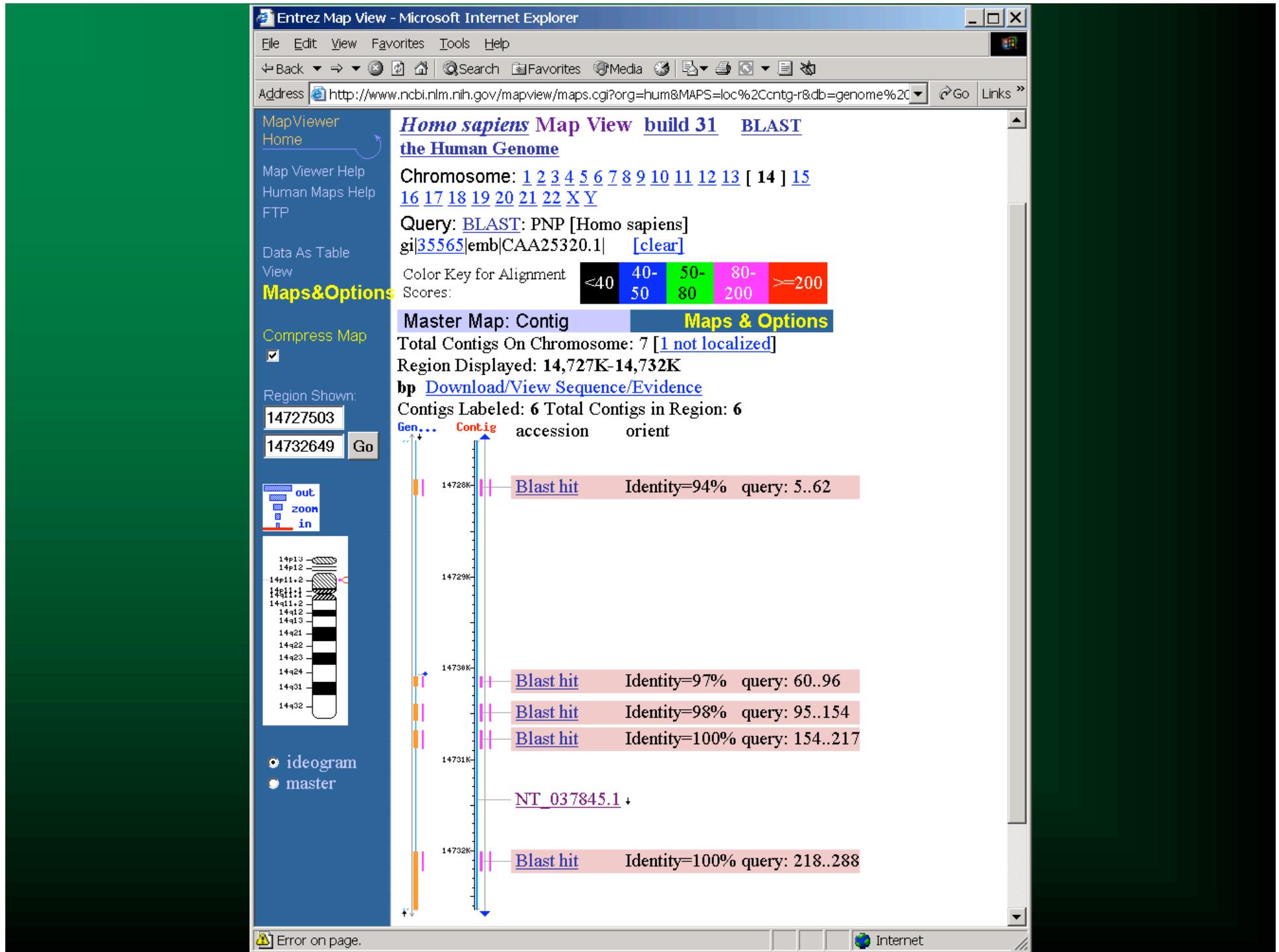
Home - Genome Browser - Blat Search - Table Browser - FAQ - User Guide

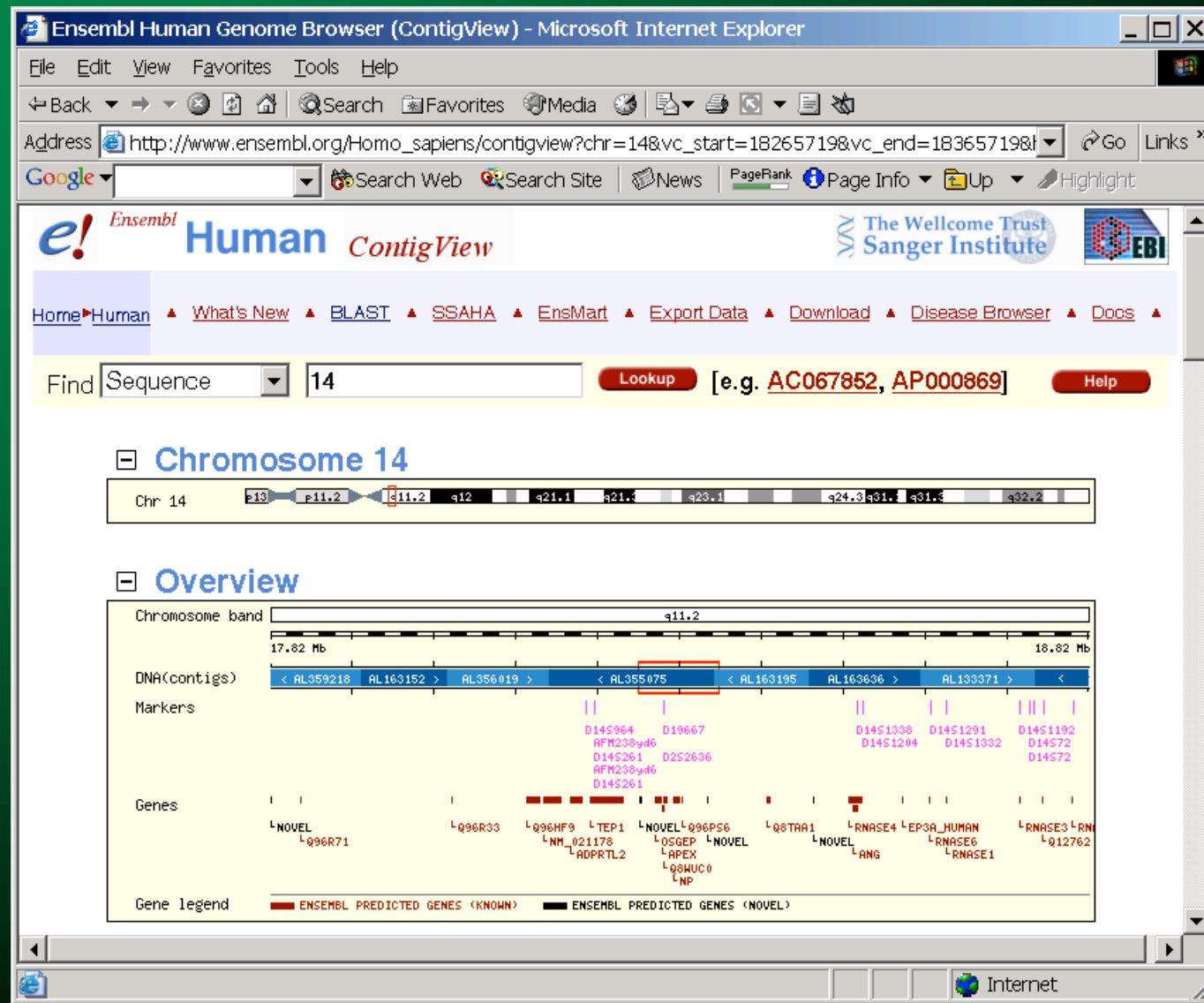
## Human BLAT Results

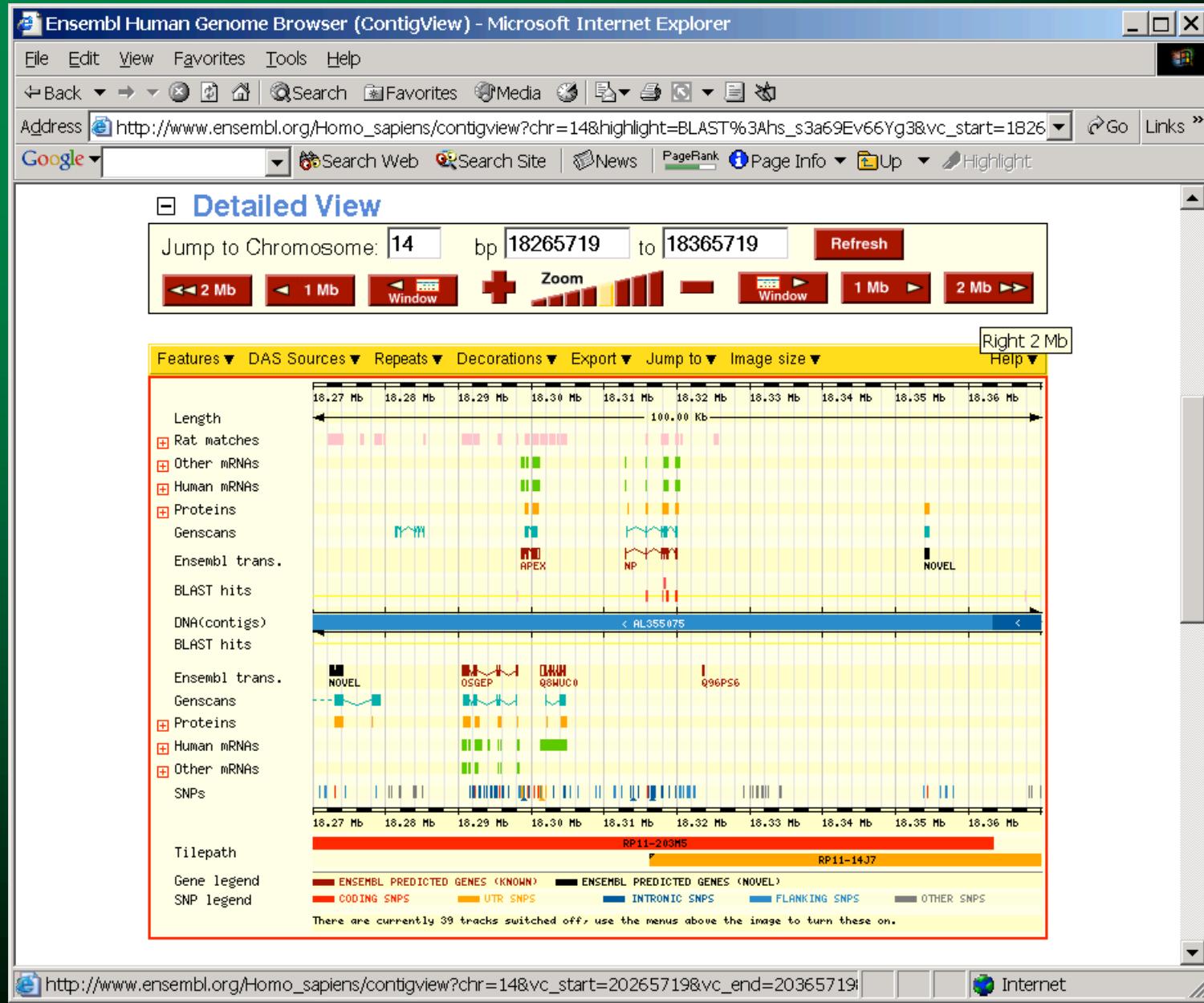
### BLAT Search Results

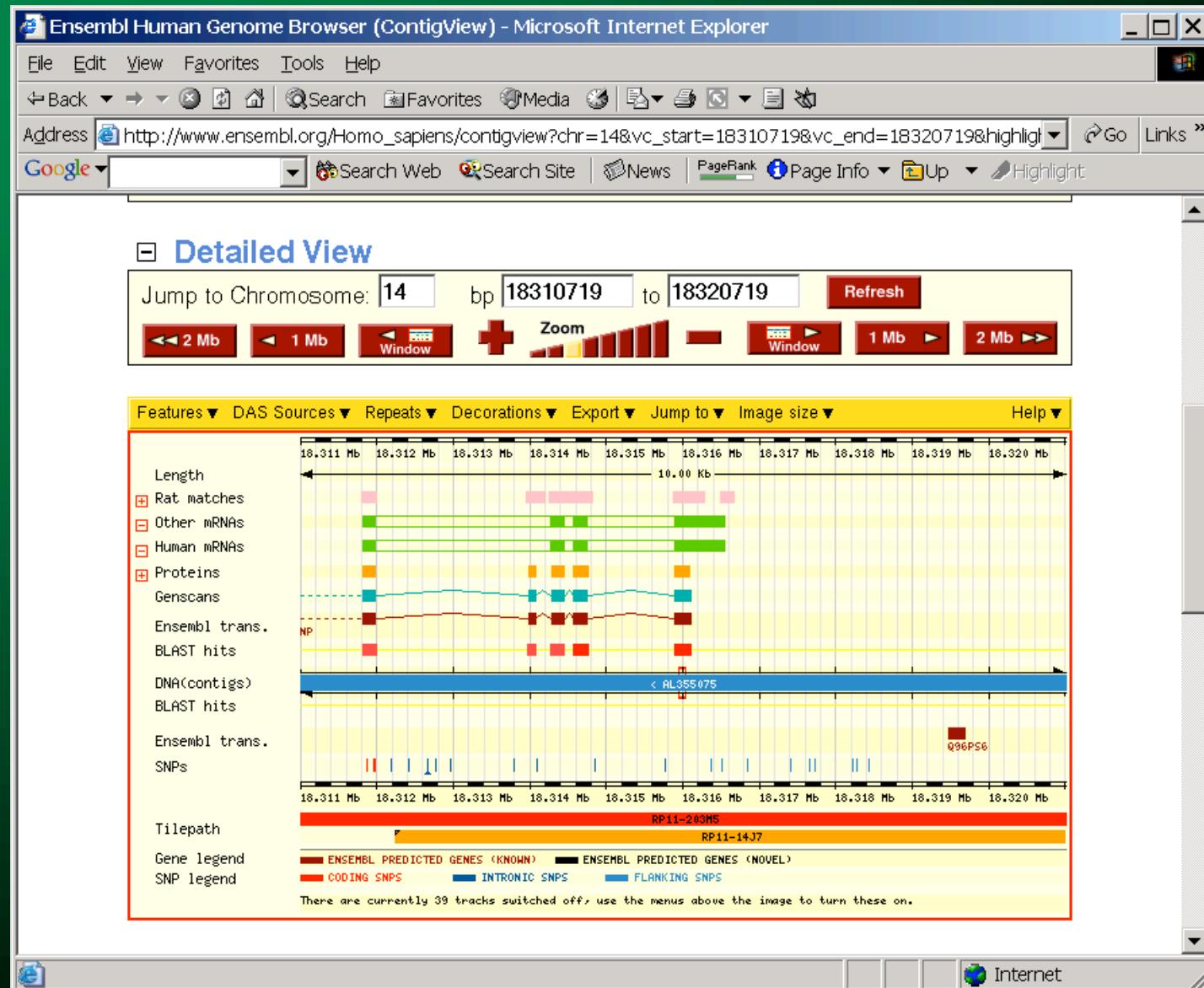
ACTIONS	QUERY	SCORE	START	END	QSIZE	IDENTITY	CHRO	STRAND	START	END
<a href="#">browser details</a>	PNP	279	4	289	289	99.7%	14	++	14727930	14732220
<a href="#">browser details</a>	PNP	101	0	288	289	70.0%	2	+-	76697947	76698808

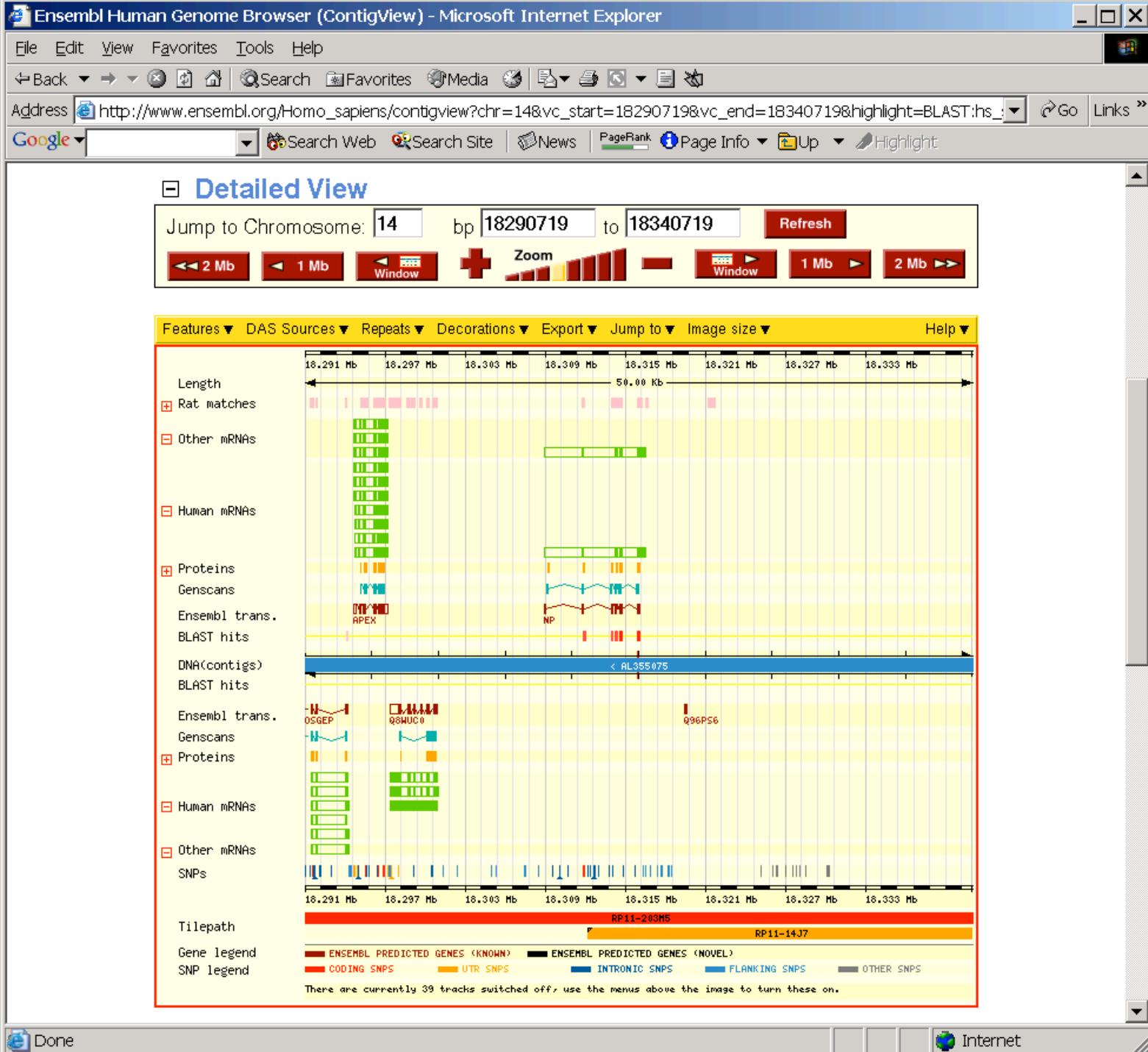
Done Internet











Ensembl Human Genome Browser (SyntenyView) - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Address http://www.ensembl.org/Homo\_sapiens/syntenyview?species=Mus\_musculus&chr=14&loc=18290719 Go Links

Google Search Web Search Site News PageRank Page Info Up Highlight

e! Ensembl Human SyntenyView The Wellcome Trust Sanger Institute EBI

Home ▶ Human ▶ What's New ▶ BLAST ▶ SSAHA ▶ EnsMart ▶ Export Data ▶ Download ▶ Disease Browser ▶ Docs ▶

Find All Lookup [e.g. AP000462, RH9632, cancer] Help

**Homo sapiens chromosome 14**

**Human Chromosome 14**

Jump to chromosome 14 ▾ Lookup  
Jump to mapview for chromosome statistics.

**Homology Matches**

Homo_sapiens Genes	Mus_musculus Homologues
OSGEP (18.29 Mb)	-> <a href="#">Q99LN8</a> (chr 14 : 42.78 Mb)
APEX (18.29 Mb)	-> <a href="#">Apex1</a> (chr 14 : 42.79 Mb)
Q8WUC0 (18.30 Mb)	-> <a href="#">ENSMUSG00000035953</a> (chr 14 : 42.79 Mb)
NP (18.31 Mb)	-> <a href="#">Pnp</a> (chr 14 : 42.81 Mb)
Q96PS6 (18.32 Mb)	
<a href="#">ENSG00000165787</a> (18.35 Mb)	-> <a href="#">4930474F22Rik</a> (chr 14 : 42.87 Mb)
Q8TAA1 (18.42 Mb)	
<a href="#">ENSG00000169431</a> (18.48 Mb)	-> <a href="#">ENSMUSG00000035932</a> (chr 14 : 42.93 Mb)
RNASE4 (18.52 Mb)	-> <a href="#">Rnase4</a> (chr 14 : 42.96 Mb)
ANG (18.53 Mb)	-> <a href="#">Angrp</a> (chr 14 : 43.06 Mb)
EP3A_HUMAN (18.59 Mb)	-> <a href="#">Ang</a> (chr 14 : 42.97 Mb)
	-> <a href="#">ENSMUSG00000021878</a> (chr 14 : 42.98 Mb)

Ensembl Human Genome Browser (BlastView) - Microsoft Internet Explorer

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Back Search Favorites Media PageRank Page Info Up Highlight

Address http://www.ensembl.org/Homo\_sapiens/blastview?id=hs\_s3a69983966YU3&format=karyo\_format Go Links

Google Search Web Search Site News PageRank Page Info Up Highlight

e! Ensembl Human BLASTView The Wellcome Trust Sanger Institute EBI

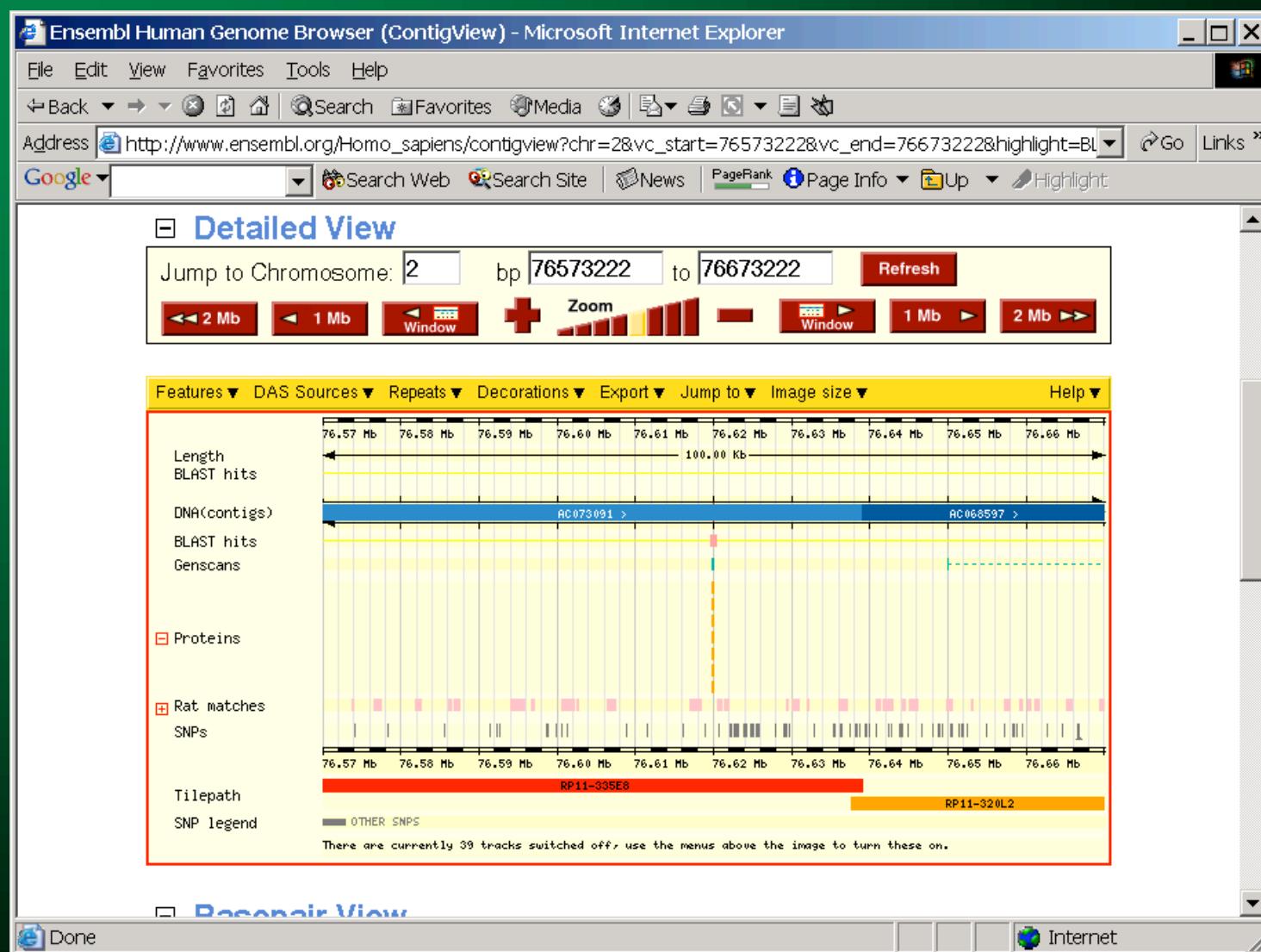
Home ▶ Human ▶ What's New ▶ BLAST ▶ SSAHA ▶ EnsMart ▶ Export Data ▶ Download ▶ Disease Browser ▶ Docs ▶

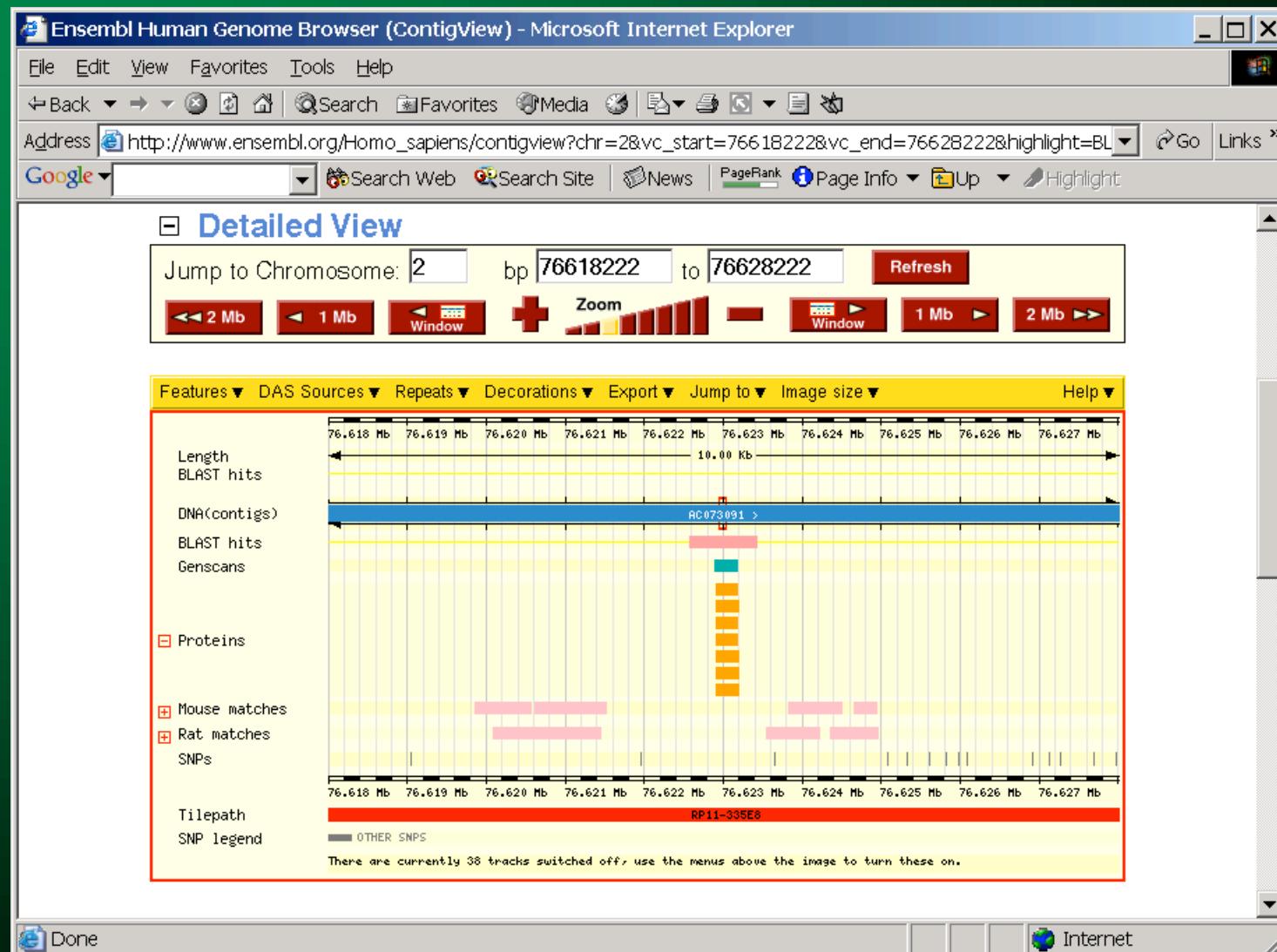
Find [All] Lookup [e.g. AP000462, RH9632, cancer] Help

1 2 3 4 5 6 7 8 9 10 11 12

13 14 15 16 17 18 19 20 21 22 X Y

Done Internet





Ensembl Human Genome Browser (BlastView) - Microsoft Internet Explorer

File Edit View Favorites Tools Help

Address http://www.ensembl.org/Homo\_sapiens/blastview?format=hit\_format&id=hs\_s3a69983966YU3&hit=AC073091.5.1.185174 Go Links

Google Search Web Search Site News PageRank Page Info Up Highlight

**e! Ensembl Human BLASTView**

Home ▶ Human ▶ What's New ▶ BLAST ▶ SSAHA ▶ EnsMart ▶ Export Data ▶ Download ▶ Disease Browser ▶ Docs ▶ Find All ▶ [e.g. AP000462, RH9632, cancer] ▶ Lookup ▶ Help

TBLASTN 2.0a13MP-WashU [10-Jun-1997] [Build 23:08:22 Jun 10 1997]  
Query= PNP  
(289 letters)  
Database: ensembl/Homo\_sapiens.latestgp.fa  
44521 sequences; 3200338544 total letters  
>AC073091.5.1.185174 Length: 185,174

Minus Strand HSPs:

Score = 842 (296.4 bits), Expect = 2.2e-81, P = 2.2e-81  
Identities = 181/868 (63%), Positives = 213/868 (74%), Frame = -1

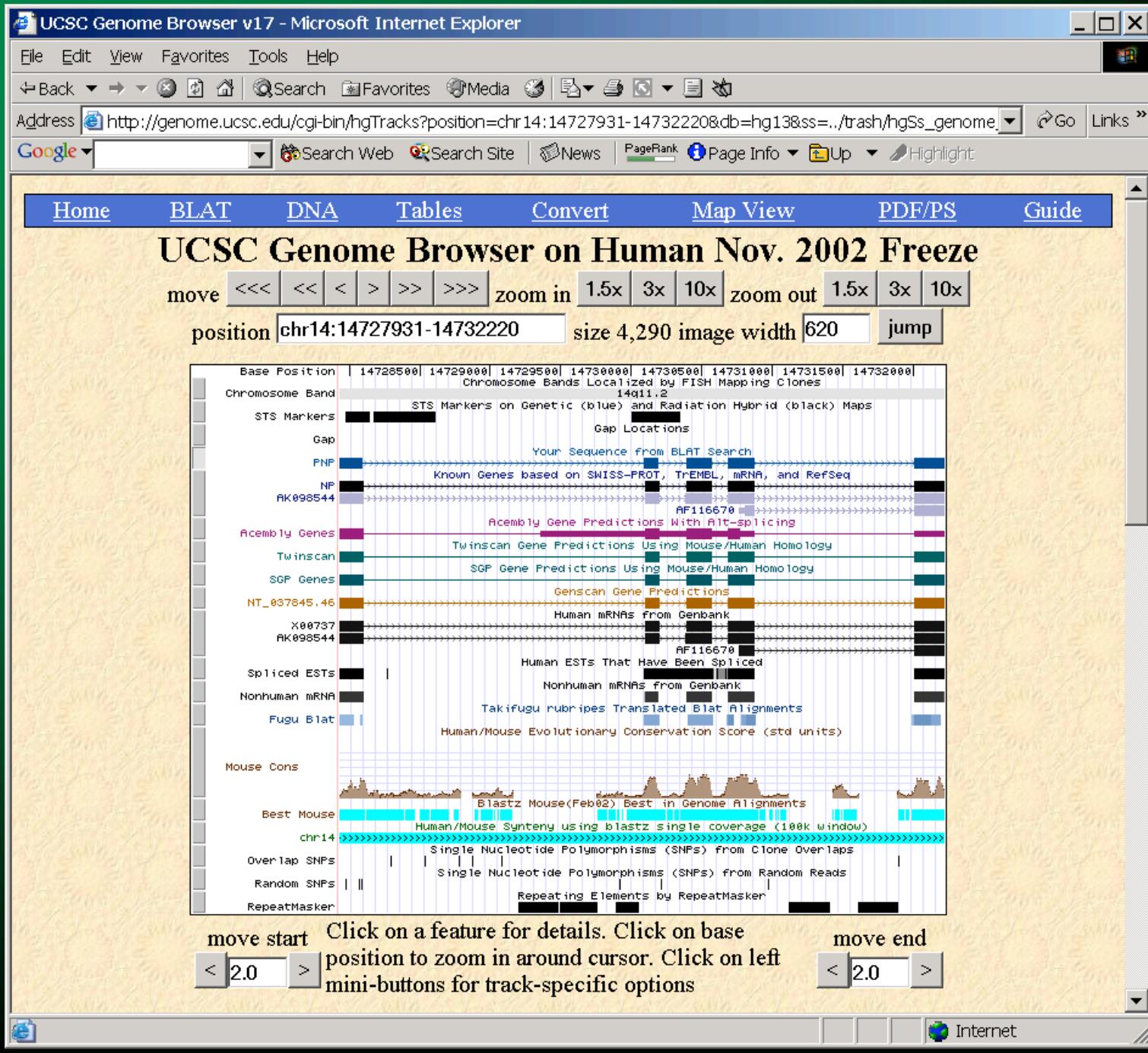
Query: 1 MENGYTYEDYKNTAEWLLSHTKHRPQVAIICGSGLGGLTDKLTQAQIFDYSEIPNFPRT 60  
MENGYTYEDY++TAEWLL HTKH QV +ICGS LG LTDKL QAQIF+ SE+ NF +ST  
Sbjct: 166478 MENGYTYEDYQSTAELLFHTKH\*TQVTVICGSELGDLTDKLIQAQIFNNSEMLNNFFQST 166299

Query: 61 VPGHAGRLVFGFLNGRACVMMQGRFHMYEGYPLWKVTFPVRVFHLLGVDTLVVTNAAGGL 120  
VPGHA LVFGFLNG CVMMQGRF++Y+GY LW + F VF LLG + LV T+AAGGL  
Sbjct: 166298 VPGHAV\*LVFGFLNGTVCVMMQGRFYLYDGYLLWNMIFLHEVFQQLLGGNILVATDAAGGL 166119

Query: 121 NPKFEVGIDMLIRDHINLPGFSGQNPLRGPNDERFGDRFPAMSDAYDRTMRQRALSTWKQ 180  
NPK EVG IML+ DHI L GF QN +GPNDERFG FPA SDAY+ TM+Q+AL++ Q  
Sbjct: 166118 NPKSEVGRIMLLCDHIKLLGFCDQNSPKGPNDERFGVHFPATSDAYNWTMKQKALNS\*NQ 165939

Query: 181 MGEQRELQEGTYVMVAGPSFETVAECRVLQKLGADAVGMSTVPEVIVARHC-GLRVFG-- 237  
MG+Q+E+Q+ TYVM +FET + + D ++ + C L +  
Sbjct: 165938 MGKQQEVQKDTYVMAVNCNFETGRDSSDAE--AGDGCLLA\*AQHQ--S\*SCMALWTWSLC 165771

Query: 238 FSLITNKVIMDYESLEKANHEEVLAAGKQAAQKLEQFVFSILMASIPLPDKAS 289  
FSLITNKVIMDYESL+KANHE V A KQAAQKLEQFVFSIL ASIPLPD A+  
Sbjct: 165770 FSLITNKVIMDYESLKKANHE\*V\*EAVKQAAQKLEQFVFSILKASIPLPDNAN 165615



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Address http://genome.ucsc.edu/cgi-bin/hgTracks?position=chr14:14727931-147322208&db=hg13&ss=..&trash/hgSs\_genom Go Links

Google Search Web Search Site News PageRank Page Info Up Highlight

**Mapping and Sequencing Tracks**

Base Position	<a href="#">Chromosome Band</a>	<a href="#">STS Markers</a>	<a href="#">FISH Clones</a>	<a href="#">GenMapDB Clones</a>
	<input type="button" value="dense"/>	<input type="button" value="dense"/>	<input type="button" value="hide"/>	<input type="button" value="hide"/>
Recomb Rate	<a href="#">Map Contigs</a>	<a href="#">Assembly</a>	<a href="#">Gap</a>	<a href="#">Coverage</a>
	<input type="button" value="hide"/>	<input type="button" value="hide"/>	<input type="button" value="dense"/>	<input type="button" value="hide"/>
BAC End Pairs	<a href="#">Fosmid End Pairs</a>	<a href="#">GC Percent</a>	<a href="#">BLAT Sequence</a>	
	<input type="button" value="hide"/>	<input type="button" value="hide"/>	<input type="button" value="hide"/>	<input type="button" value="full"/>

**Genes and Gene Prediction Tracks**

<a href="#">Known Genes</a>	<a href="#">RefSeq Genes</a>	<a href="#">Acembly Genes</a>	<a href="#">Twinscan</a>	<a href="#">SGP Genes</a>
<input type="button" value="pack"/>	<input type="button" value="hide"/>	<input type="button" value="dense"/>	<input type="button" value="dense"/>	<input type="button" value="dense"/>
<a href="#">Egenesh++ Genes</a>	<a href="#">Geneid Genes</a>	<a href="#">Genscan Genes</a>		
<input type="button" value="hide"/>	<input type="button" value="hide"/>	<input type="button" value="pack"/>		

**mRNA and EST Tracks**

<a href="#">Human mRNAs</a>	<a href="#">Spliced ESTs</a>	<a href="#">Human ESTs</a>	<a href="#">Nonhuman mRNA</a>	<a href="#">Nonhuman EST</a>
<input type="button" value="pack"/>	<input type="button" value="dense"/>	<input type="button" value="hide"/>	<input type="button" value="dense"/>	<input type="button" value="hide"/>
<a href="#">UniGene</a>	<a href="#">Gene Bounds</a>			
<input type="button" value="hide"/>	<input type="button" value="hide"/>			

**Expression and Regulation**

<a href="#">CpG Islands</a>				
<input type="button" value="hide"/>				

**Comparative Genomics**

<a href="#">Fugu Blat</a>	<a href="#">Mouse Cons</a>	<a href="#">Tight Mouse</a>	<a href="#">Best Mouse</a>	<a href="#">Blastz Mouse</a>
<input type="button" value="dense"/>	<input type="button" value="full"/>	<input type="button" value="hide"/>	<input type="button" value="dense"/>	<input type="button" value="hide"/>
<a href="#">Mouse Synteny</a>	<a href="#">Rat Synteny</a>			
<input type="button" value="full"/>	<input type="button" value="hide"/>			

**Variation and Repeats**

<a href="#">Overlap SNPs</a>	<a href="#">Random SNPs</a>	<a href="#">RepeatMasker</a>	<a href="#">Simple Repeats</a>
<input type="button" value="dense"/>	<input type="button" value="dense"/>	<input type="button" value="dense"/>	<input type="button" value="hide"/>

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Address http://genome.ucsc.edu/cgi-bin/hgTracks?position=chr2:76697948-76698808&db=hg13&ss=../trash/hgs; Go Links

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Home BLAT DNA Tables Convert Map View PDF/PS Guide

## UCSC Genome Browser on Human Nov. 2002 Freeze

move <<< << < > >> zoom in 1.5x 3x 10x zoom out 1.5x 3x 10x

position chr2:76697948-76698808 size 861 image width 620 jump

Base Position 76698100 76698200 76698300 76698400 76698500 76698600 76698700 76698800 Chromosome Bands Localized by FISH Mapping Clones

Chromosome Band

STS Markers STS Markers on Genetic (blue) and Radiation Hybrid (black) Maps

Gap Gap Locations

PNP Your Sequence from BLAT Search Known Genes based on SWISS-PROT, TREMBL, mRNA, and RefSeq

Assembly Genes Assembly Gene Predictions With Alt-splicing

Twinscan Twinscan Gene Predictions Using Mouse/Human Homology

SGP Genes SGP Gene Predictions Using Mouse/Human Homology

Genscan Genscan Gene Predictions

NT\_022184.153 Human mRNAs from Genbank

Spliced ESTs Human ESTs That Have Been Spliced

Nonhuman mRNA Nonhuman mRNAs from Genbank

Fugu Blat Takifugu rubripes Translated Blat Alignments

Human/Mouse Evolutionary Conservation Score (std units)

Mouse Cons

Best Mouse Blastz Mouse(Feb02) Best in Genome Alignments

chr6 Human/Mouse Suntensity using blastz single coverage (100K Window)

Overlap SNPs Single Nucleotide Polymorphisms (SNPs) from Clone Overlaps

Random SNPs Single Nucleotide Polymorphisms (SNPs) from Random Reads

RepeatMasker Repeating Elements by RepeatMasker

move start Click on a feature for details. Click on base move end  
position to zoom in around cursor. Click on left < 2.0 > mini-buttons for track-specific options < 2.0 >

< 2.0 > Internet



# Gene Information

## ✓ BLink

- BLAST Hits

## ✓ Domains

- Protein domains

## ✓ Links

- Varies with available information

## ✓ LinkOut

- “Custom” links to other relevant databases

NCBI Sequence Viewer - Microsoft Internet Explorer

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Back Forward Stop Home Search Favorites Media Print Copy Paste Find Links

Address http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list\_uids=130377&dopt=GenPept Go Links »

NCBI Protein

PubMed Nucleotide Protein Genome Structure PMC Taxonomy OMIM Books

Search **Protein** for  Go Clear

Limits Preview/Index History Clipboard Details

Display default Show: 20 Send to File Get Subsequence

**□ 1: P00491. Purine nucleoside...[gi:130377]**

[BLink](#), [Domains](#), [Links](#)

LOCUS P00491 289 aa linear PRI 15-JUN-2002

DEFINITION Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP).

ACCESSION P00491

VERSION P00491 GI:130377

DBSOURCE swissprot: locus PNPH\_HUMAN, accession P00491;  
class: standard.  
extra accessions:Q15160, created: Jul 21, 1986.  
sequence updated: Jul 21, 1986.  
annotation updated: Jun 15, 2002.  
xrefs: gi: [35564](#), gi: [35565](#), gi: [190150](#), gi: [387033](#), gi: [190147](#),  
gi: [190148](#), gi: [190149](#), gi: [66583](#), gi: [230387](#), gi: [230388](#)  
xrefs (non-sequence databases): Aarhus/Ghent-2DPAGE2108, MIM  
[164050](#), InterProIPR001369, PfamPF00896, PROSITEPS01240

KEYWORDS Transferase; Glycosyltransferase; Polymorphism; Disease mutation;  
3D-structure.

SOURCE Homo sapiens (human)

ORGANISM [Homo sapiens](#)  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.

REFERENCE 1 (residues 1 to 289)

AUTHORS Williams,S.R., Goddard,J.M. and Martin,D.W. Jr.

TITLE Human purine nucleoside phosphorylase cDNA sequence and genomic  
clone characterization

JOURNAL Nucleic Acids Res. 12 (14), 5779-5787 (1984)

MEDLINE [84272252](#)

PUBMED [6087295](#)

REMARK SEQUENCE FROM N.A.

REFERENCE 2 (residues 1 to 289)

AUTHORS Williams,S.R., Gekeler,V., McIvor,R.S. and Martin,D.W. Jr.

TITLE A human purine nucleoside phosphorylase deficiency caused by a

Done Internet



NCBI

BLAST

Protein

Structure

PubMed

Taxonomy

Genome

Nucleotide

3D-Domains

Books

Help

Query: gi|130377 purine-nucleoside phosphorylase (EC 2.4.2.1) [validated] - human

Matching gi: [35565](#), [4557801](#), [66583](#), [230387](#), [230388](#)

## COG0005 assigned by Cognitor (35 best hits)

Best hits

Common Tree

Taxonomy Report

3D structures

CDD-Search

GI list

148 BLAST hits to 98 unique species [Sort by taxonomy proximity](#)

22

Archaea

79

Bacteria

42

Metazoa

2

Fungi

0

Plants

0

Viruses

3

Other Eukaryotae

Keep only



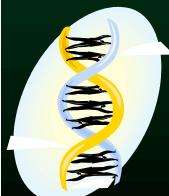
Cut-Off 100

Select

Reset

289 aa

	SCORE	P	ACCESSION	GI	PROTEIN DESCRIPTION
	1515	27	<a href="#">AAA36460</a>	<a href="#">387033</a>	purine nucleoside phosphorylase [Homo sapiens]
	1501	27	<a href="#">BAC05327</a>	<a href="#">21758578</a>	unnamed protein product [Homo sapiens]
	1341	21	<a href="#">1B8NA</a>	<a href="#">4558113</a>	Chain A, Purine Nucleoside Phosphorylase
	1341	21	<a href="#">P55859</a>	<a href="#">3287982</a>	Purine nucleoside phosphorylase (Inosine phosphorylase)
	1335	21	<a href="#">1FXUA</a>	<a href="#">11514560</a>	Chain A, Purine Nucleoside Phosphorylase From Canis lupus familiaris
	1334	21	<a href="#">AAB34886</a>	<a href="#">1042206</a>	purine nucleoside phosphorylase, PNP, purine nucleoside phosphorylase
	1332	21	<a href="#">1A9O</a>	<a href="#">3402089</a>	Chain , Bovine Purine Nucleoside Phosphorylase (bovine)
	1331	21	<a href="#">1VFN</a>	<a href="#">2624420</a>	Chain , Purine Nucleoside Phosphorylase
	1329	21	<a href="#">1A9T</a>	<a href="#">3318947</a>	Chain , Bovine Purine Nucleoside Phosphorylase (bovine)
	1329	21	<a href="#">1PBN</a>	<a href="#">1311143</a>	Chain , Purine Nucleoside Phosphorylase
	1324	21	<a href="#">1A9Q</a>	<a href="#">3402091</a>	Chain , Bovine Purine Nucleoside Phosphorylase (bovine)
	1290	21	<a href="#">CAA39888</a>	<a href="#">53750</a>	purine-nucleoside phosphorylase [Mus musculus]
	1290	21	<a href="#">AAC37635</a>	<a href="#">388921</a>	purine nucleoside phosphorylase
	1287	21	<a href="#">AAA39835</a>	<a href="#">200098</a>	purine nucleoside phosphorylase
	1282	21	<a href="#">AAC37706</a>	<a href="#">388923</a>	purine nucleoside phosphorylase
	1267	21	<a href="#">BAB25491</a>	<a href="#">12842148</a>	unnamed protein product [Mus musculus]
	1001	21	<a href="#">XP_214155</a>	<a href="#">27674996</a>	similar to purine-nucleoside phosphorylase [Mus musculus]
	814	8	<a href="#">EAA11700</a>	<a href="#">21299555</a>	agCP6049 [Anopheles gambiae str. PEST]
	760	8	<a href="#">AAF47654</a>	<a href="#">7292245</a>	CG16758-PB [Drosophila melanogaster]



NCBI CD Summary - Microsoft Internet Explorer

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Address [http://www.ncbi.nlm.nih.gov/Structure/cdd/wrpsb.cgi?INPUT\\_TYPE=precalc&SEQUENCE=130377](http://www.ncbi.nlm.nih.gov/Structure/cdd/wrpsb.cgi?INPUT_TYPE=precalc&SEQUENCE=130377) Go Links

**NCBI Conserved Domain Summary**

New Search PubMed Nucleotide Protein Structure CDD Taxonomy Help?

**Query:** [gi|130377|sp|P00491|PNPH\\_HUMAN](#) Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP)  
(289 letters)

**Database:** cdd.v.1.60

[gnl|CDD|4371 pfam00896, Mtap\_PNP, Phosphorylase... S= 295 E=3e-]

1 50 100 150 200 250 289  
**Mtap\_PNP**

Show Domain Relatives [gnl|CDD|4371 pfam00896, Mtap\_PNP, Phosphorylase family 2] Details

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Address http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=protein&list\_uids=130377&dopt=GenPept Go Links »

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PubMed Nucleotide Protein Genome Structure PMC Taxonomy OMIM Books

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Display FASTA Show: 20 Send to File Get Subsequence

□ 1: P00491. Purine nucleoside...[gi:130377]

LOCUS P00491 289 aa linear PRI 15-J

DEFINITION Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP)

ACCESSION P00491

VERSION P00491 GI:130377

DBSOURCE swissprot: locus PNPH\_HUMAN, accession P00491;  
class: standard.  
extra accessions:Q15160, created: Jul 21, 1986.  
sequence updated: Jul 21, 1986.  
annotation updated: Jun 15, 2002.  
xrefs: gi: [35564](#), gi: [35565](#), gi: [190150](#), gi: [387033](#), gi: [190147](#),  
gi: [190148](#), gi: [190149](#), gi: [66583](#), gi: [230387](#), gi: [230388](#)  
xrefs (non-sequence databases): Aarhus/Ghent-2DPAGE2108, MIM  
[164050](#), InterProIPR001369, PfamPF00896, PROSITEPS01240

KEYWORDS Transferase; Glycosyltransferase; Polymorphism; Disease mutation;  
3D-structure.

SOURCE Homo sapiens (human)

ORGANISM [Homo sapiens](#)  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.

REFERENCE 1 (residues 1 to 289)

AUTHORS Williams, S.R., Goddard, J.M. and Martin, D.W. Jr.

TITLE Human purine nucleoside phosphorylase cDNA sequence and genomic clone characterization

Links

- Related Sequences
- Domain Relatives
- OMIM
- PubMed
- Taxonomy
- LinkOut



OMIM - NUCLEOSIDE PHOSPHORYLASE; NP - Microsoft Internet Explorer

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Address <http://www.ncbi.nlm.nih.gov/entrez/dispmim.cgi?id=164050> Go Links

**NCBI**

MIM \*164050  
Text  
Allelic Variants  
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References  
Contributors  
Creation Date  
Edit History  
Clinical Synopsis  
Gene map

LocusLink  
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[GenBank](#)  
[Protein](#)  
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LinkOut  
[...CCR](#)  
[...HGMD](#)

**OMIM**  
OnLine Mendelian Inheritance in Man

 Johns Hopkins University

PubMed Nucleotide Protein Genome Structure PMC Taxonomy OMIM

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**\*164050** Links  
**NUCLEOSIDE PHOSPHORYLASE; NP**

**Alternative titles; symbols**

**PURINE-NUCLEOSIDE:ORTHOPHOSPHATE RIBOSYLTRANSFERASE; PNP**  
**NUCLEOSIDE PHOSPHORYLASE DEFICIENCY, INCLUDED**  
**ATAXIA WITH DEFICIENT CELLULAR IMMUNITY, INCLUDED**

Gene map locus [14q13.1](#)

**TEXT**

[Edwards et al. \(1971\)](#) described electrophoretic variants of nucleoside phosphorylase ([EC 2.4.2.1](#)), the enzyme that catalyzes the phosphorolytic cleavage of inosine to hypoxanthine. The enzyme appeared to be a trimer. Family studies indicated autosomal codominant inheritance of the variants. [Zannis et al. \(1978\)](#) and [Williams et al. \(1984\)](#) demonstrated that human PNP is a symmetric trimer composed of 3 identical 32,153-Da subunits, each with a substrate-binding site. PNP reversibly catalyzes the phosphorolysis of the purine nucleosides, (deoxy)inosine and (deoxy)guanosine, to their respective purine bases and the corresponding ribose-1-phosphate. 

OMIM - NUCLEOSIDE PHOSPHORYLASE; NP - Microsoft Internet Explorer

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Address <http://www.ncbi.nlm.nih.gov/entrez/dispmim.cgi?id=164050> Go Links



MIM \*164050

Text

Allelic Variants

• View List

See Also

References

Contributors

Creation Date

Edit History

• Clinical

Synopsis

• Gene map

LocusLink

N Nomenclature

R RefSeq

G GenBank

P Protein

U UniGene

LinkOut

...CCR

...HGMD

Deficiency of nucleoside phosphorylase results in defective T-cell immunity ([Giblett et al., 1975](#)). This may not be surprising since deficiency of adenosine deaminase, the next enzyme in the pathway, results in combined immune deficiency disease ([102700](#)). Absence of red cell NP was observed in a child with severe T-cell immunodeficiency. The parents were consanguineous and showed less than half the normal activity of the enzyme in their red cells ([Berglund et al., 1975](#)). In a patient with deficiency of nucleoside phosphorylase, [Cohen et al. \(1976\)](#) found severe hypouricemia and hypouricosuria, but excessive amounts of purines (mainly inosine and guanosine) in the urine. The immune defect was thought to be related to inhibition of adenosine deaminase by inosine. [Mitchell et al. \(1978\)](#) found that deoxyadenosine and deoxyguanosine are particularly toxic to T cells but not to B cells. Addition of deoxycytidine or dipyridamole prevented deoxyribonucleoside toxicity. [Stoop et al. \(1977\)](#) studied a 15-month-old girl, 2 sisters of whom had died of immunodeficiency. NP was lacking from red cells and lymphocytes. The parents and a normal brother had intermediate levels. Both T cells and B cells were normal at birth, but thereafter a gradual decrease in T-cell immunity occurred. The patient showed high inosine and guanosine levels in the blood, as well as hypouricemia and hypouricosuria. Spastic tetraparesis was present. In one patient with severely defective T-cell function and normal B-cell function, [Osborne et al. \(1977\)](#) found no detectable red cell NP and no detectable immunologically reactive material. The parents, second cousins, had less than half the normal enzyme activity. Two patients in a second family had 0.5% residual enzyme activity and about half-normal immunologically reactive material. The parents, who were not related, showed electrophoretically different mutant enzymes that were also different from those in the first family. Thus the affected children in the second family were genetic compounds, not true homozygotes. In T cells, the absence of PNP activity is thought to lead to an accumulation of deoxyguanosine triphosphate, which inhibits the enzyme ribonucleotide reductase ([Mitchell et al., 1978](#); [Ullman et al., 1979](#)). This inhibition blocks DNA synthesis, thereby preventing the cellular proliferation required for an immune response. ☺

The immune defect from NP deficiency is often accompanied by a neurologic disorder. [Watson et al. \(1981\)](#) reported the case of a 2.5-year-old boy who died of malignant lymphoma of the B-immunoblastic type. He had spastic tetraplegia also. [Rijken et al. \(1987\)](#) described a case in a 3-year-old boy who was admitted for investigation of a behavior disorder and spastic diplegia. Severe lymphopenia was found; however, clinical symptoms of immune deficiency did not become apparent until the age of 4 years. [Stephenson and Tolmie \(1990\)](#) informed me that the family reported by [Graham-Pole et al. \(1975\)](#) as having 'familial dysequilibrium-diplegia with T-lymphocyte deficiency' ([209000](#)) turned out to have PNP deficiency. The condition was diagnosed retrospectively from stored fibroblasts from an affected child and from demonstration that both parents had half-normal activity of PNP. [Stephenson and Tolmie \(1990\)](#) were prompted to restudy this family after diagnosing PNP deficiency in a young girl who presented with dysequilibrium syndrome with pyramidal signs (extensor plantar responses and exaggerated reflexes but not prominent spasticity) very similar to the neurologic picture in the family reported by [Graham-Pole et al. \(1975\)](#). The child had defective cell-mediated immunity and died of lymphoma shortly after her third birthday. ☺

Although early studies suggested that B-cell function is normal or even increased in PNP deficiency, later studies showed that B-cell function can be disrupted as well ([Markert, 1991](#)). This was the case in a patient in whom the nature of the molecular defects was demonstrated by [Aust et al. \(1992\)](#): she had normal B-cell counts but significantly depressed immunoglobulin levels. ☺

OMIM - NUCLEOSIDE PHOSPHORYLASE; NP - Microsoft Internet Explorer

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## ALLELIC VARIANTS (selected examples)

### .0001 NUCLEOSIDE PHOSPHORYLASE DEFICIENCY [NP, GLU89LYS]

[Williams et al. \(1987\)](#) cloned the mutant gene from an NP-deficient patient who was the offspring of a consanguineous mating. A single base difference was found in the coding region of the mutant gene, a G-to-A transition in the third exon. This single base mutation altered the codon at position 89 from glu-to-lys, a result consistent with previously published peptide mapping data. The patient was demonstrated to be homozygous for the single base mutation on the basis of hybridization of synthetic oligomers to genomic DNA digests. 

### .0002 NUCLEOSIDE PHOSPHORYLASE DEFICIENCY [NP, ALA174PRO]

[Markert and Barrett \(1989\)](#) demonstrated a G-to-C change of nucleotide 520, resulting in a substitution of proline for alanine as amino acid 174. The other allele carried the mutation described by [Williams et al. \(1987\)](#), namely, a G-to-A change of nucleotide 265, resulting in a glu-to-lys change in amino acid 89 ([164050.0001](#)). [Markert \(1992\)](#) indicated that when site-directed mutagenesis was used to create this mutation and the mutant allele was expressed in COS cells, it was found to have normal function. The possibility remains, however, that the mutation was the cause of the patient's clinical disorder, with an abnormality in protein stability or other posttranscriptional stages. 

### .0003 NUCLEOSIDE PHOSPHORYLASE DEFICIENCY [NP, ASP128GLY]

In a patient with nucleoside phosphorylase deficiency, [Aust et al. \(1992\)](#) found an asp128-to-gly substitution in the maternal allele and an arg234-to-pro mutation ([164050.0004](#)) in the paternal allele. In addition, the patient was homozygous for a ser51-to-gly substitution ([164050.0005](#)), which is a polymorphism. In order to prove that the 2 mutations were responsible for the disease state, each of the 3 mutations was constructed separately by site-

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**REFERENCES**

1. Aitken, D. A.; Ferguson-Smith, M. A. :  
**Regional assignment of nucleoside phosphorylase by exclusion to 14q13.**  
*Cytogenet. Cell Genet.* 22: 490-492, 1978.  
PubMed ID : [110525](#)

2. Allderdice, P. W.; Miller, O. J.; Miller, D. A.; Klinger, H. P. :  
**Spreading of inactivation in an (X;14) translocation.**  
*Am. J. Med. Genet.* 2: 233-240, 1978.  
PubMed ID : [263441](#)

3. Aust, M. R.; Andrews, L. G.; Barrett, M. J.; Norby-Slycord, C. J.; Markert, M. L. :  
**Molecular analysis of mutations in a patient with purine nucleoside phosphorylase deficiency.**  
*Am. J. Hum. Genet.* 51: 763-772, 1992.  
PubMed ID : [1384322](#)

4. Berglund, C.; Ammann, A. J.; Giblett, E. R. :  
**Characteristics of nucleoside phosphorylase in the parents of a child with deficiency of the enzyme. (Abstract)**  
*Am. J. Hum. Genet.* 27: 17A only, 1975.

5. Carapella De Luca, E.; Stegagno, M.; Dionisi Vici, C.; Paesano, R.; Fairbanks, L. D.; Morris, G. S.; Simmonds, H. A. :  
**Prenatal exclusion of purine nucleoside phosphorylase deficiency.**  
*Europ. J. Pediat.* 145: 51-53, 1986.  
PubMed ID : [3089796](#)

6. Cohen, A.; Doyle, D.; Martin, D. W., Jr.; Ammann, A. J. :  
**Abnormal purine metabolism and purine overproduction in a patient deficient in purine nucleoside phosphorylase.**  
*New Eng. J. Med.* 295: 1449-1454, 1976.  
PubMed ID : [975775](#)

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Query: gi|[130377](#) purine-nucleoside phosphorylase (EC 2.4.2.1) [validated] - human  
Matching gi: [35565](#), [4557801](#), [66583](#), [230387](#), [230388](#)

COG0005 assigned by Cognitor (35 best hits)

Best hits Common Tree Taxonomy Report 3D structures CDD-Search GI list

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22 Archaea 79 Bacteria 42 Metazoa 2 Fungi 0 Plants 0 Viruses 3 Other Eukaryotae

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289 aa

SCORE	P	ACCESSION	GI	PROTEIN DESCRIPTION
1515	27	<a href="#">AAA36460</a>	<a href="#">387033</a>	purine nucleoside phosphorylase [Homo sapiens]
1501	27	<a href="#">PAC05327</a>	<a href="#">21758579</a>	unnamed protein product [Homo sapiens]
1341	21	<a href="#">1B8NA</a>	<a href="#">4558113</a>	Chain A, Purine Nucleoside Phosphorylase
1341	21	<a href="#">P33639</a>	<a href="#">3287982</a>	Purine nucleoside phosphorylase (inosine phosphorylase)
1335	21	<a href="#">1FXUA</a>	<a href="#">11514560</a>	Chain A, Purine Nucleoside Phosphorylase From Calf Sple
1334	21	<a href="#">AAB34886</a>	<a href="#">1042206</a>	purine nucleoside phosphorylase, PNP, purine nucleoside
1332	21	<a href="#">1A9Q</a>	<a href="#">3402089</a>	Chain , Bovine Purine Nucleoside Phosphorylase Complex
1331	21	<a href="#">1VFN</a>	<a href="#">2624420</a>	Chain , Purine Nucleoside Phosphorylase
1329	21	<a href="#">1A9T</a>	<a href="#">3318947</a>	Chain , Bovine Purine Nucleoside Phosphorylase Complex
1329	21	<a href="#">1PBN</a>	<a href="#">1311143</a>	Chain , Purine Nucleoside Phosphorylase
1324	21	<a href="#">1A9Q</a>	<a href="#">3402091</a>	Chain , Bovine Purine Nucleoside Phosphorylase Complex
1290	21	<a href="#">CAA39888</a>	<a href="#">53750</a>	purine-nucleoside phosphorylase [Mus musculus]
1290	21	<a href="#">AAC37635</a>	<a href="#">388921</a>	purine nucleoside phosphorylase
1287	21	<a href="#">AAA39835</a>	<a href="#">200098</a>	purine nucleoside phosphorylase
1282	21	<a href="#">AAC37706</a>	<a href="#">388923</a>	purine nucleoside phosphorylase
1267	21	<a href="#">BAB25491</a>	<a href="#">12842148</a>	unnamed protein product [Mus musculus]

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1: 1B8NA. Chain A, Purine N...[gi:4558113]

LOCUS 1B8N\_A 284 aa linear MAM 02

DEFINITION Chain A, Purine Nucleoside Phosphorylase.

ACCESSION 1B8N\_A

VERSION 1B8N\_A GI:4558113

DBSOURCE pdb: molecule 1B8N, chain 65, release Feb 2, 1999;  
deposition: Feb 2, 1999;  
class: Transferase;  
source: Mol\_id: 1; Organism\_scientific: Bos Taurus;  
Organism\_common: Bovine; Organ: Spleen;  
Exp. method: X-Ray Diffraction.

KEYWORDS .

SOURCE Bos taurus (cow)

ORGANISM [Bos taurus](#)  
Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;  
Mammalia; Eutheria; Cetartiodactyla; Ruminantia; Pecora; Bovoidea;  
Bovidae; Bovinae; Bos.

REFERENCE 1 (residues 1 to 284)

AUTHORS Mao,C., Cook,W.J., Zhou,M., Federov,A.A., Almo,S.C. and Ealick,S.E.

TITLE Calf spleen purine nucleoside phosphorylase complexed with  
substrates and substrate analogues

JOURNAL Biochemistry 37 (20), 7135-7146 (1998)

MEDLINE [98254498](#)

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**Description:** Purine Nucleoside Phosphorylase.

**Deposition:** A.A.Fedorov, G.A.Kicska, E.V.Fedorov, B.V.Strokopytov, P.C.Tyler, R.H.Furneaux, V.L.Schramm & S.C.Almo, 2-Feb-99

**Taxonomy:** [Bos taurus](#)

**Reference:** [PubMed](#)   **MMDB:** [13072](#)   **PDB:** [1B8N](#)

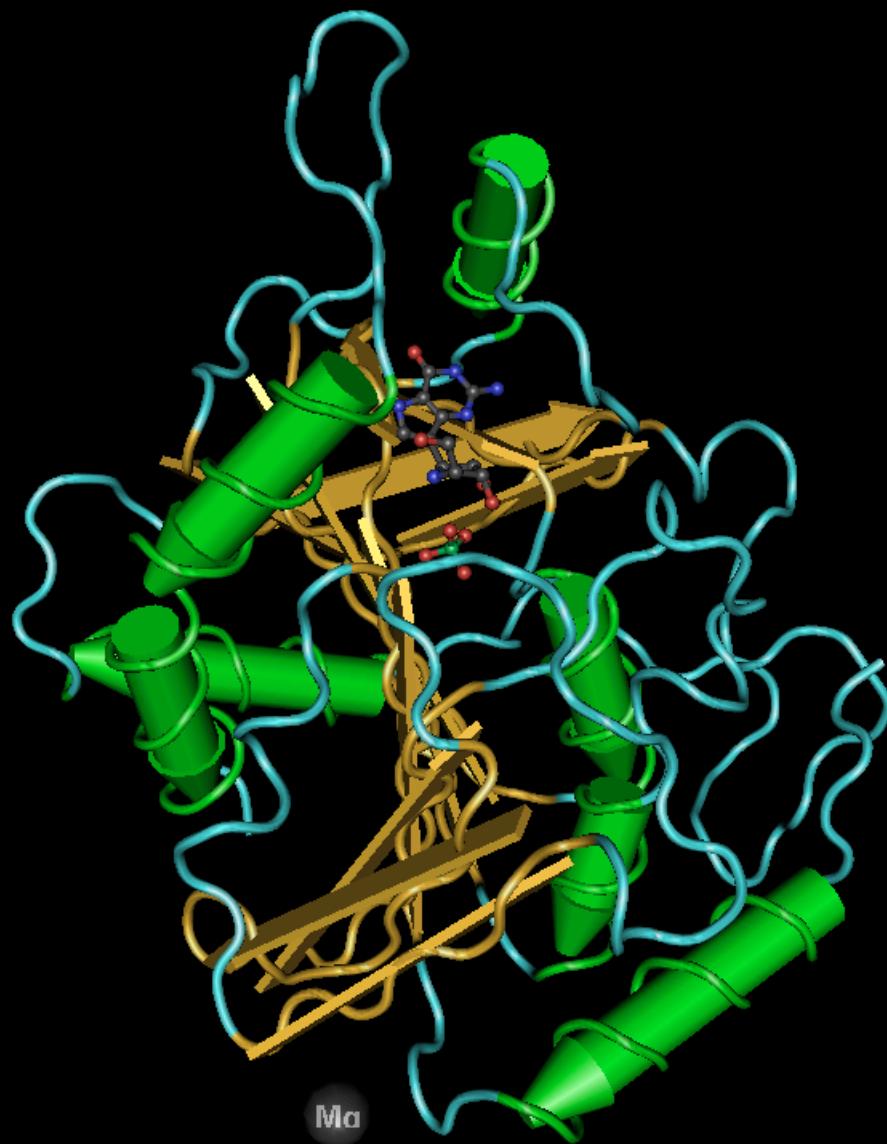
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Protein 1 50 100 150 200 250 284  
Chain A

CDs Mtap\_PNP

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LocusID Org Symbol Description Position Links

4860 Hs NP nucleoside phosphorylase 14q13.1 P O R G P H U V

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- ▶ ESTs, Highly similar to PNPH\_HUMAN Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP) [H.sapiens]-Bt.3800
- ▶ ESTs, Weakly similar to PHHUPN purine-nucleoside phosphorylase (EC 2.4.2.1) [validated] - human [H.sapiens]-Dr.3216
- ▶ purine-nucleoside phosphorylase-Pnp
- ▶ nucleoside phosphorylase-NP
- ▶ SPARC-like 1-Sparcl1

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<b>Rattus norvegicus</b>	Sparcl1	SPARC-like 1 <a href="#">UniGene</a>
<b>Drosophila melanogaster</b>	CG16758	Drosophila melanogaster CG16758 gene <a href="#">FlyBase</a>   <a href="#">UniGene</a>
<b>Danio rerio</b>	Dr.3216	ESTs, Weakly similar to PHHUPN purine-nucleoside phosphorylase (EC 2.4.2.1) [validated] - human [H.sapiens] <a href="#">UniGene</a>
<b>Xenopus laevis</b>	XI.16206	ESTs, Weakly similar to PNPH_HUMAN Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP) [H.sapiens] <a href="#">UniGene</a>
<b>Homo sapiens</b>	NP	nucleoside phosphorylase <a href="#">MapViewer</a>   <a href="#">LocusLink</a>
<b>Bos taurus</b>	Bt.3800	ESTs, Highly similar to PNPH_HUMAN Purine nucleoside phosphorylase (Inosine phosphorylase) (PNP) [H.sapiens] <a href="#">UniGene</a>

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Organism-Gene	Sequence	Percent ID	Sequence	Organism-Gene
► M.musculus -Pnp	<a href="#">NM_013632.1</a>	92.9	<a href="#">BF543947.1</a>	R.norvegicus - Sparc11
► M.musculus -Pnp	<a href="#">NM_013632.1</a>	86.5	<a href="#">BF230403.1</a>	B.taurus - Bt.3800
► M.musculus -Pnp	<a href="#">NM_013632.1</a>	85.0	<a href="#">NM_000270.1</a>	H.sapiens - NP
► M.musculus -Pnp	<a href="#">NM_013632.1</a>	73.1	<a href="#">BJ060275.1</a>	X.laevis - Xl.16206
► M.musculus -Pnp	<a href="#">NM_013632.1</a>			D.melanogaster - CG16758
► M.musculus -Pnp	<a href="#">NM_013632.1</a>	68.7	<a href="#">BF717799.1</a>	D.rerio - Dr.3216

### ADDITIONAL CALCULATED ORTHOLOGS

► B.taurus -Bt.3800	<a href="#">BF230403.1</a>	90.6	<a href="#">NM_000270.1</a>	H.sapiens - NP
► R.norvegicus -Sparc11	<a href="#">BF543947.1</a>	87.6	<a href="#">NM_000270.1</a>	H.sapiens - NP
► B.taurus -Bt.3800	<a href="#">AV606522.1</a>	84.9	<a href="#">BF543947.1</a>	R.norvegicus - Sparc11
► H.sapiens -NP	<a href="#">NM_000270.1</a>			D.melanogaster - CG16758
► R.norvegicus -Sparc11	<a href="#">BF543947.1</a>			D.melanogaster - CG16758
► X.laevis -Xl.16206	<a href="#">BI676077.1</a>	73.3	<a href="#">BF543947.1</a>	R.norvegicus - Sparc11
► B.taurus -Bt.3800	<a href="#">AV603243.1</a>			D.melanogaster - CG16758
► B.taurus -Bt.3800	<a href="#">AV667617.1</a>	71.3	<a href="#">BF717799.1</a>	D.rerio - Dr.3216
► D.rerio -Dr.3216	<a href="#">BF717799.1</a>	71.2	<a href="#">NM_000270.1</a>	H.sapiens - NP
► X.laevis -Xl.16206	<a href="#">BJ060275.1</a>	70.5	<a href="#">NM_000270.1</a>	H.sapiens - NP
► B.taurus -Bt.3800	<a href="#">BE481281.1</a>	70.3	<a href="#">BJ060275.1</a>	X.laevis - Xl.16206
► X.laevis -Xl.16206	<a href="#">BI676077.1</a>			D.melanogaster - CG16758

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for sequences similar to one or more query sequences of any
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finds.

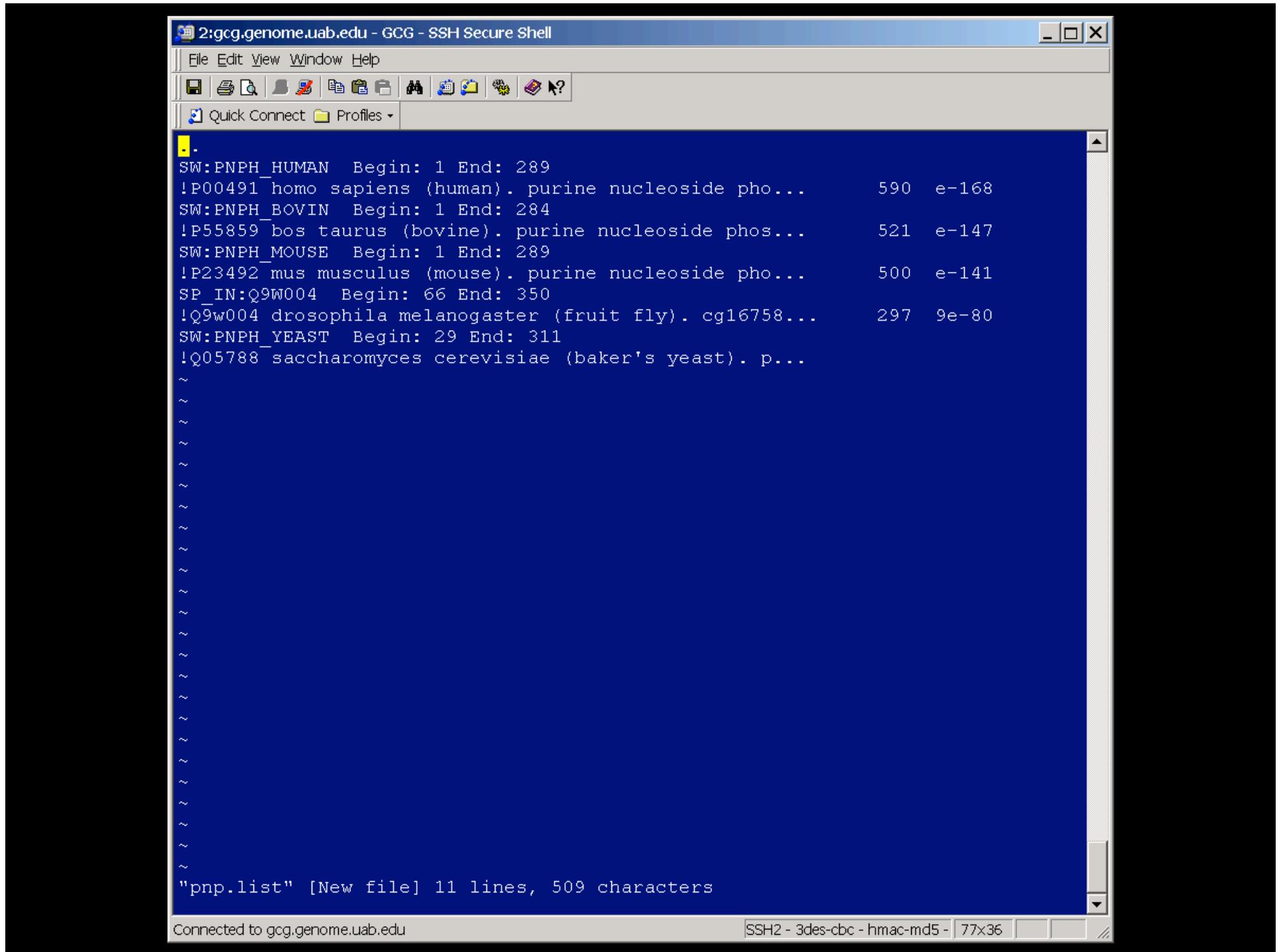
      Begin (* 1 *) ?
      End (*    289 *) ?

Search for query in what sequence database:

 1) pir          p Protein Information Resource
 2) nrl_3d       p NRL_3D Protein Sequence-Structure Database
 3) swplus       p SWISS-PROT + SP-TREMBL
 4) genembl     n GenBank + EMBL (HTGs Removed)
 5) htg          n High Throughput Genomes (HTG from GenBank and EMBL)
 6) est_human    n Human Expressed Sequence Tags (GenBank and EMBL)
 7) est_mouse    n Mouse Expressed Sequence Tags (GenBank and EMBL)
 8) est_other    n All Other Expressed Sequence Tags (GenBank and EMBL)
 9) htc          n High Throughput cDNA (GenBank and EMBL)
10) gss          n Genome Survey Sequences (GSS from GenBank and EMBL)
11) genpept     p GenPept (Translated GenBank)

Please choose one (* 1 *): 3
```

Sequences producing significant alignments:			
	Score (bits)	E Value	..
SW:PNPH_HUMAN Begin: 1 End: 289			
!P00491 homo sapiens (human). purine nucleoside pho...	590	e-168	
SP_HUM:Q8N7G1 Begin: 1 End: 293			
!Q8n7g1 homo sapiens (human). hypothetical protein ...	582	e-165	
SW:PNPH_BOVIN Begin: 1 End: 284			
!P55859 bos taurus (bovine). purine nucleoside phos...	521	e-147	
SW:PNPH_MOUSE Begin: 1 End: 289			
!P23492 mus musculus (mouse). purine nucleoside pho...	500	e-141	
SP_RO:Q9D8C9 Begin: 16 End: 301			
!Q9d8c9 mus musculus (mouse). adult male small intes...	492	e-138	
SP_IN:Q9W004 Begin: 66 End: 350			
!Q9w004 drosophila melanogaster (fruit fly). cg16758...	297	9e-80	
SP_IN:Q95TC1 Begin: 72 End: 353			
!Q95tc1 drosophila melanogaster (fruit fly). gh03781...	295	3e-79	
SP_IN:Q9BMI9 Begin: 10 End: 285			
!Q9bmi9 schistosoma mansoni (blood fluke). purine-nu...	279	2e-74	
SW:PNPH_YEAST Begin: 29 End: 311			
!Q05788 saccharomyces cerevisiae (baker's yeast). p...	265	3e-70	
SP_BAP:Q8XHI8 Begin: 8 End: 272			
!Q8xhi8 clostridium perfringens. purine-nucleoside ...	257	8e-68	
SP_FUN:Q9UTG1 Begin: 40 End: 312			
!Q9utg1 schizosaccharomyces pombe (fission yeast). ...	256	2e-67	
SP_BAP:Q9X259 Begin: 29 End: 292			
!Q9x259 thermotoga maritima. purine nucleoside phos...	251	6e-66	
SP_BAP:Q92A55 Begin: 21 End: 271			
!Q92a55 listeria innocua. pnp protein. 10/2002	248	5e-65	
SW:PUNA_BACST Begin: 7 End: 271			
!P77834 bacillus stearothermophilus. purine nucleos...	248	5e-65	
SP_BAP:Q9KCN8 Begin: 21 End: 269			
!Q9kcn8 bacillus halodurans. purine nucleoside phos...	246	2e-64	
SP_BAP:Q8Y5V2 Begin: 21 End: 271			
!Q8y5v2 listeria monocytogenes. pnp protein. 6/2002	245	3e-64	



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```
> pileup @pnp.list

PileUp creates a multiple sequence alignment from a group of related
sequences using progressive, pairwise alignments. It can also plot a
tree showing the clustering relationships used to create the alignment.

      1      PNPH_HUMAN    289 aa
      2      PNPH_BOVIN     284 aa
      3      PNPH_MOUSE     289 aa
      4      Q9W004        285 aa
      5      PNPH_YEAST     283 aa

What is the gap creation penalty (* 8 *) ?  5

What is the gap extension penalty (* 2 *) ?  1

This program can display the clustering relationships graphically.
Do you want to:

A) Plot to a FIGURE file called "pileup.figure"
B) Plot graphics on HP7550 attached to /dev/tty15
C) Suppress the plot

Please choose one (* A *):  c

What should I call the output file name (* pnp.msf *) ?

Determining pairwise similarity scores...

      1   x     2       4.72
      1   x     3       4.45
      1   x     4       2.69
      1   x     5       2.52
      2   x     3       4.45
      2   x     4       2.70
```

1	50
PNPH_HUMAN	MENGYTYEDY KNTAEWLLSH TKHRPQVAAI CGSGLGGLTD KLTQ....AQ
PNPH_BOVIN	MQNGYTYEDY QDTAKWLLSH TEQRPVAVI CGSGLGGLVN KLTQ....AQ
PNPH_MOUSE	MENEFTYEDY ETTAKWLLQH TEYRPQVAVI CGSGLGGLTA HLKE....AQ
Q9W004	~EDTYPYEV~ EEIADFITKG SGMRPKIGII CGSGLGSLAD MIQD....PK
PNPH YEAST	~~~~~F KNTTNF.... .EPPRTLII CGSGLGGIST KLSRDNPPPV
51	100
PNPH_HUMAN	IFDYSEIPNF PRSTVPGHAG RLVFGFLNGR ACVM.MQGRF HMYEGYPLWK
PNPH_BOVIN	TFDYSEIPNF PESTVPGHAG RLVFGILNGR ACVM.MQGRF HMYEGYPFWK
PNPH_MOUSE	IFDYNEIPNF PQSTVQGHAG RLVFGFLNGR CCVM.MQGRF HMYEGYSLSK
Q9W004	IFEYEKIPNF PVSTVEGHAG RLVVGTLEG. ATVMAMQGRF HFYEGYPLAK
PNPH YEAST	TVPYQDIPGF KKSTVPGHSG TLMFGSMNGS PVVL.MNGRL HGYEGNTLFE
101	150
PNPH_HUMAN	VTFPVVRVFHL LG.VDTLVVT NAAGGLNPKF EVGDIMLIIRD HINLPGFSQ
PNPH_BOVIN	VTFPVVRVFR LG.VETLVVT NAAGGLNPNF EVGDIMLIIRD HINLPGFSGE
PNPH_MOUSE	VTFPVVRVFHL LG.VETLVVT NAAGGLNPNF EVGDIMLIIRD HINLPGFCGQ
Q9W004	CSMPVRVMKL CG.VEYLFA~ NAAGGINPRF AVGDIMLMHD HVMLGFAGN
PNPH YEAST	TTFFPIRVLNH MGHVRNLIVT NAAGGINAKY QACDLMCIYD HLNIPGLAQ
151	200
PNPH_HUMAN	NPLRGPNDER FGDRFPAMSD AYDRTMRQRA LSTWKQMGEQ RELQEGETYVM
PNPH_BOVIN	NPLRGPNNEER FGVRFPAMSD AYDRDMRQKA HSTWKQMGEQ RELQEGETYVM
PNPH_MOUSE	NPLRGPNDER FGVRFPAMSD AYDRDMRQKA FTAWKQMGEQ RKLQEGETYVM
Q9W004	SPLQGPNDPR FGPRFPALVN SYNKDLINKA IEIAKAMGIE SNIHGVYSC
PNPH YEAST	HPLRGPNLDE DGPRFLALSD AYDLELRKLL FKKWKEKIQ RPLHEGETYTF
201	250
PNPH_HUMAN	VAGPSFETVA ECRVLQKLGA DAVGMSTVPE VIVARHCGLR VFGFSLITNK
PNPH_BOVIN	LGGPNFETVA ECRLLRNLGA DAVGMSTVPE VIVARHCGLR VFGFSLITNK
PNPH_MOUSE	LAGPNFETVA ESRLLKMLGA DAVGMSTVPE VIVARHCGLR VFGFSLITNK
Q9W004	LGGPNEYETIA ELKALRMMGV DAVGMSTVHE VITARHCDMK VFAFSLITNK
PNPH YEAST	VSGPTFETRA ESKMIRMLGG DAVGMSTVPE VIVARHCGWR VLALSLITNT
251	300
PNPH_HUMAN	VIMDY.....ES...L..E KANHEEVLA~ GKQAAQKLEQ FVSILMASIP
PNPH_BOVIN	VIMDY.....ES...Q..G KANHEEVLEA GKQAAQKLEQ FVSILMASIP
PNPH_MOUSE	VVMDY.....EN...L..E KANHMEVLD~ GKAAAQTLER FVSILMESIP
Q9W004	CATEY.....SD...KKDD EAHDEVMAV AKNRQKACCE LVSLIREIH
PNPH YEAST	CVVDSPASAL DESPVPLEKG KATHAEVLEN GKIASNDVQN LIAAVM <b>GEL</b> ~
301	
PNPH_HUMAN	LPDKAS
PNPH_BOVIN	V~~~~~
PNPH_MOUSE	LPDRGS
Q9W004	L~~~~~
PNPH YEAST	~~~~~

**Adobe Acrobat - [pnp.pdf]**

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Bookmarks

File: C:pnp.ps Date: Sun Mar 02 16:52:31 2003

Page 1 of 1

CLUSTAL G (1.3) MULTIPLE SEQUENCE ALIGNMENT

PH\_HUMAN MENGYYEDVYINIAEWLLRHTHRPGVAIICSGGLGGIADRLW...  
 PH\_BOVIN MNGTYTYEDVYIIRAKWLLQHIEJRPJVAVICSGGLGGIIVNKLW...  
 PH\_MOUSE MENEFYTYEDVYIIRAKWLLQHIEJRPJVAVICSGGLGGIWAHLKE...  
 Q9W004 -EDTIPYEVIEERIADPPIKGSGMRPEIGIICSGGLGSLADMIDQ...  
 PH\_YEAST FPKWLNPF-EPPPTLIIICSGGLGGITLRLDPPVTVPYQI...  
 1.....10.....20.....30.....40.....50.....60.....70.....80.....90.....100.....110.....120.....130.....140.....150.....160

154  
154  
154  
153  
144

PH\_HUMAN PGDPFPAMDADETMQGALALTWKQMGEBRELCEGYVMVAGPSPETVAECRVV...  
 PH\_BOVIN PGVRPPAMDAIDRDMRKAFAWLMGEBORKLCEGYIVMLGGPWFETV...  
 PH\_MOUSE PGVRPPAMDAIDRDMRKAFAWLMGEBORKLCEGYIVMLGGPWFETV...  
 Q9W004 PGPRPEPALVNSWHEDLINKAIEIAKAMGIESNIHVGVNSCLGGPWEET...  
 PH\_YEAST DGPFPFLALDALELPFLFLPKWYELKIQORPLBEGTYFVSGPTPE...  
 .....170.....180.....190.....200.....210.....220.....230.....240.....250.....260.....270.....280.....290.....300

289  
284  
289  
285  
283

Comments Thumbnails Signatures

1 of 1 11 x 8.5 in

NCBI CDD pfam00896 with Query Sequence added - Microsoft Internet Explorer

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Address http://www.ncbi.nlm.nih.gov/StructureG/cdd/cddsrvcgi?ascbin=2&maxln=10&seltype=3&uid=pfam00896&queryg Go Links

# Conserved Domain Database

**CDD: pfam00896.6, Mtap PNP, Query added PSSM-Id: 4371**

**Source:** Pfam[US], Pfam[UK]

**Description:** Phosphorylase family 2.

**Taxa:** cellular organisms

**Status:** Alignment from source

**Aligned:** 14 rows

**Proteins:** [Click here for CDART summary of Proteins containing pfam00896]

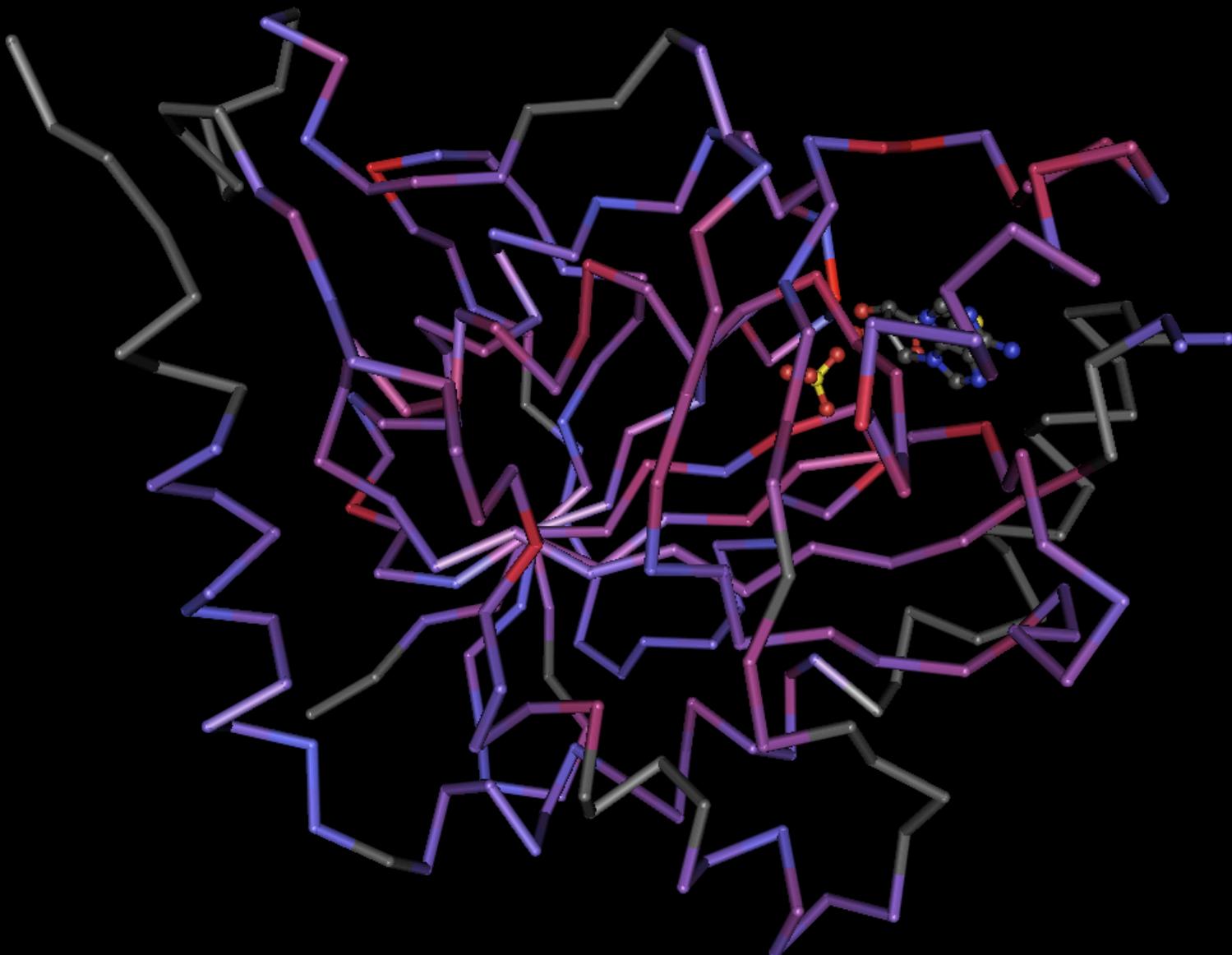
View 3D Structure with Cn3D using Virtual Bonds (To display structure, download Cn3D)

View Alignment as Hypertext width 60 color at 2.0 bits

Subset Rows up to 10 sequences most similar to the query

	10	20	30	40	50	60
consensus	.....*	.....*	.....*	.....*	.....*	.....*
query	1 PPMIGIIGGSGLGDLADLLEREV	-----V-PYS	DTPYGKPSTVIGHAGTLVGG	--GVAV	52	
<u>1CG6_A</u>	24 RPQVAIIICGSGLGGLTDKITQAQ	-----IfDYSEIPNFPRSTVPGHAGRLVFGflNGRAC	78			
<u>1B80_A</u>	9 AVKIGIIGGTGLDDPEILEGRTE	-----K--YVDTPFGKPSDALILG-KIKN	--VDCVL	57		
<u>1I80_A</u>	24 RPQAVVICGSGLGGLVNKLTQAQt	-----Fd-YSEIPNFPESTVPGHAgRLVFGgilNGRAC	78			
qi_2113997	27 EHDVAVVLGSGWLPAVAALGSPPT	-----vLp-QAELPGFPPTAACGHAgELLsvpiGAHRV	82			
qi_2622720	5 GRMLGVIGGSGFYTFFGSDRTV	-----NSDTPYGQPSAPITIG-TIGv-HDVAF	52			
qi_2506345	1 --MIGIIGGTGIYEMAEYGRLER	-----Rg-SLITPYGKTPEISVFK-LHG	--RRVAF	47		
qi_2494051	25 TPRVAFILGSGLGLADQIENAV	-----Is-YEKLPGFPVSTVHGHAgELVLghlQGVPV	79			
qi_1169270	37 PPRTLIIICGSGLGGISTKLSRDNpppvTvP-YQDIPGFKKSTVPGHSgtIMFgsmNGSPV	95				
	19 SPKIGLILGSGLGLADEIENPV	k----Lk-YEDIPEFPVSTVEGHAgQLVLgtlEGVSV	73			
	.....*	.....*	.....*	.....*	.....*	.....*
consensus	53 LARHGRGHDYEPHKVNRYAN-IRALKALG-VERLIVTNAAGSLREELEPGDLVV	PDDHIN	110			
query	79 VMMQGRFHMYEGYPLWKVTFpVRVFHLLG-VDTLWVTAAGGLNPKFEVGDIML	IRDHIN	137			
<u>1CG6_A</u>	58 LARHGRQHTIMPSKVNYQAN-IWALKEEG-CTHVIVTTACGSLREEIQPGDIVIIDQFID	115				
<u>1B80_A</u>	79 VMMQGRFHMYEGYPFWKVTpVRVFRILLG-VETLWVTAAGGLNPNFEVGDIML	IRDHIN	137			
<u>1I80_A</u>	83 IVLAGRIHAYEGHDLRYVVHpVRAARAAG-AQIMVLTNAAGGLRADLQVGQPVLISDHLN	141				
qi_2113997	53 LPRHGAHHQYSAHAVPYRAN-MWALRALG-VRRVFGPCAVGSDLPELEPGAVVV	PDQLVD	110			
qi_2622720	48 IPRHS PGHDKPPHMVNRYAN-IWALKELG-VRQIIATNAVGSLKRSIGPGDFVV	PHDFLD	105			
qi_2506345	80 VCMKGRGHFYEGRGMTIMTdaIRTFKLLG-CELLFCTNAAGSLRPEVGAGSILVAL	KDHIN	138			
qi_2494051	96 VLMNGRLHGYEGNTLFETTFpIRVLNHMhVRNLIVTNAAGGINAKYQACDI	MCYDHIN	155			
qi_1169270	74 IAMQGRFHYEYEGYSMEKVTFpVRVMKALG-VEALIVTNAAGGVNTEFRAGDI	MIITDHIN	132			

Done Internet





## Only One Final Word of Wisdom...

- “...although the computer is a wonderful helpmate for the sequence searcher and comparer, biochemists and molecular biologists must guard against the blind acceptance of any algorithmic output; given the choice, think like a biologist and not a statistician.”  
– - Russell F. Doolittle, 1990



**Farewell!**