Find a Gene Assignment – BGGN213

Jason Hsiao (A15871650)

1) Protein: Human RBP4 (Retinol Binding Protein 4 Isoform a Precursor)

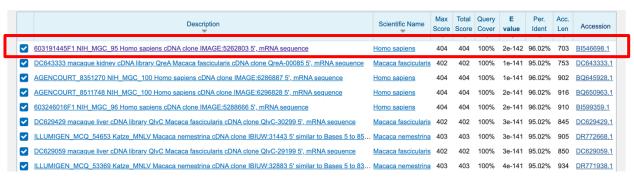
Accession Number: NP 001310446.1

Species: Human

Function: "This protein belongs to the lipocalin family and is the specific carrier for retinol (vitamin A alcohol) in the blood. It delivers retinol from the liver stores to the peripheral tissues. In plasma, the RBP-retinol complex interacts with transthyretin which prevents its loss by filtration through the kidney glomeruli. A deficiency of vitamin A blocks secretion of the binding protein post translationally and results in defective delivery and supply to the epidermal cells." [provided by RefSeq, Jul 2008].

2) BLAST Method Used: tblastn 2.14.1+ Database: Expressed sequence tags (est)





 <u>Land Company Company</u> 3) 603191445F1 NIH_MGC_95 Homo sapiens cDNA clone IMAGE:5262803 5', mRNA sequence Sequence ID: BI546698.1 Length: 703 Number of Matches: 1 Range 1: 52 to 654 GenBank Graphics ▼ Next Match ▲ Previous Match Method Identities Positives 404 bits(1037) 2e-142 Compositional matrix adjust. 201/201(100%) 201/201(100%) 0/201(0%) +1 MKWVWallllaalGSGRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGLFLQDNIV MKWVWALLLLAALGSGRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGLFLODNIV Sbjct 52 MKWVWALLLLAALGSGRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGLFLQDNIV Query 61 AEFSVDETGOMSATAKGRVRLLNNWDVCADMVGTFTDTEDPAKFKMKYWGVASFLOKGND AEFSVDETGOMSATAKGRVRI.LNNWDVCADMVGTFTDTEDPAKFKMKYWGVASFI.OKGND Query 121 DHWIVDTDYDTYAVOYSCRLLNLDGTCADSYSFVFSRDPNGLPPEAOKIVROROEELCLA Sbict 412 DHWIVDTDYDTYAVOYSCRLLNLDGTCADSYSFVFSRDPNGLPPEAOKIVROROEELCLA RQYRLIVHNGYCDGRSERNLL ROYRLIVHNGYCDGRSERNLL

Sbjct 592

ROYRLIVHNGYCDGRSERNLL

>B1546698.1 603191445F1 NIH_MGC_95 Homo sapiens cDNA clone IMAGE:5262803 5', mRNA sequence

Translated Protein Sequence via Emboss Transeq

>BI546698.1_1 603191445F1 NIH_MGC_95 Homo sapiens cDNA clone IMAGE:5262803 5', mRNA sequence

SGGLRAVPLPVGGFLGKMKWVWALLLLAALGSGRAERDCRVSSFRVKENFDKARFSGTWY AMAKKDPEGLFLQDNIVAEFSVDETGQMSATAKGRVRLLNNWDVCADMVGTFTDTEDPAK FKMKYWGVASFLQKGNDDHWIVDTDYDTYAVQYSCRLLNLDGTCADSYSFVFSRDPNGLP PEAQKIVRQRQEELCLARQYRLIVHNGYCDGRSERNLL*QYQESSFI*ELLISSX

Longest sequence without stop * of the 6 reading frames.

4)

Descriptions	Graphic Summary	Alignments	Taxonomy										
Sequences producing significant alignments					Download Y Select columns Y Show 100 Y								
select all 10	00 sequences selected			<u>GenPept</u>	Graphics D	istanc	e tree (of resul	lts <u>M</u>	ultiple ali	<u>ignme</u>	ent MSA Viewe	
Description					Scientific Name	Max Score		Query Cover	E value	Per. Ident	Acc. Len	Accession	
▼ retinol-binding protein 4 isoform X1 [Gorilla gorilla gorilla]					Gorilla gorilla go	454	454	91%	4e-160	100.00%	265	XP_018890983.	
retinol-binding protein 4 isoform X1 [Hylobates moloch]					Hylobates moloch	438	438	89%	2e-154	99.05%	214	XP_031992464.	
✓ retinol-binding protein 4 precursor [Pan troglodytes]					Pan troglodytes	421	421	85%	3e-148	100.00%	201	NP_001038960	
✓ retinol-binding protein 4 [Papio anubis]					Papio anubis	421	421	85%	6e-148	99.50%	201	XP_003904062.	
retinol-binding protein 4 [Nomascus leucogenys]					Nomascus leuco	420	420	85%	1e-147	99.00%	201	XP 003255281.	

No 100% identity with Homo sapiens, top hit is from gorilla.

5) Trimmed Alignment:

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>Danio_rerio_RBP4_(NP_570995.1)

XM---LRLCIAVCVLATCWAQDCQVSNFAVQQDFNRTRYQGTWYAVAKKDPVGL

FXLLDNIVANFKVEEDGTMTATAIGRVIILNNWEMCANMFGTFEDTEDPAKFKMKYWGAA
AYXLQTGYDDHWIIDTDYDNYAIHYSCRELDEDGTCLDGYSFIFSRHPDGLRPEDQAIVT
QKKXQDICFLGKYRRVAHTGFCEAA
>Rattus_norvegicus_RBP4_(NP_037294.1)

XMEWVWALVLLAALGGGSAERDCRVSSFRVKENFDKARFSGLWYAIAKKDPEGL
FXLQDNIIAEFSVDEKGHMSATAKGRVRLLSNWEVCADMVGTFTDTEDPAKFKMKYWGVA
SFXLQRGNDDHWIIDTDYDTFALQYSCRLQNLDGTCADSYSFVFSRDPNGLTPETRRLVR
QRQXEELCLERQYRWIEHNGYCQSR
>Callithrix_jacchus_RBP4_(XP_054099088.1)

KMKWVWALLLLAVLGISRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGL
FXLQDNIIAEFSVDETGQMSATAKGRVRLLNNWDVCADMVGTFTDTEDPAKFKMKYWGVA
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SFXLQKGNDDHWIIDTDYDTYAVQYSCRLLNLDGTCADSYSFVFSRDPNGLPPEAQRIIR QRQXEELCLARQYRLIVHNGYCDGK

>Gorilla_RBP4_(XP_018890983.3)

KMKWVWALILLAAIGSGRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGL FXLQDNIVAEFSVDETGQMSATAKGRVRLLNNWDVCADMVGTFTDTEDPAKFKMKYWGVA SFXLQKGNDDHWIVDTDYDTYAVQYSCRLLNLDGTCADSYSFVFSRDPNGLPPEAQKIVR QRQXEELCLARQYRLIVHNGYCDGR

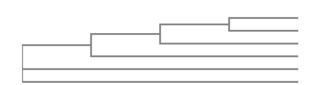
>Unknown protein (BI546698.1 1)

KMKWVWALLLLAALGSGRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGL FXLQDNIVAEFSVDETGQMSATAKGRVRLLNNWDVCADMVGTFTDTEDPAKFKMKYWGVA SFXLQKGNDDHWIVDTDYDTYAVQYSCRLLNLDGTCADSYSFVFSRDPNGLPPEAQKIVR QRQXEELCLARQYRLIVHNGYCDGR

>Homo sapiens RBP4 (NP 001310446.1)

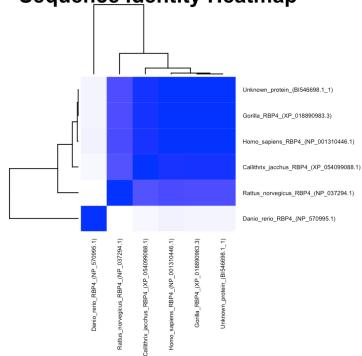
XMKWVWALLLLAALGSGRAERDCRVSSFRVKENFDKARFSGTWYAMAKKDPEGL FXLQDNIVAEFSVDETGQMSATAKGRVRLLNNWDVCADMVGTFTDTEDPAKFKMKYWGVA SFXLQKGNDDHWIVDTDYDTYAVQYSCRLLNLDGTCADSYSFVFSRDPNGLPPEAQKIVR QRQXEELCLARQYRLIVHNGYCDGR



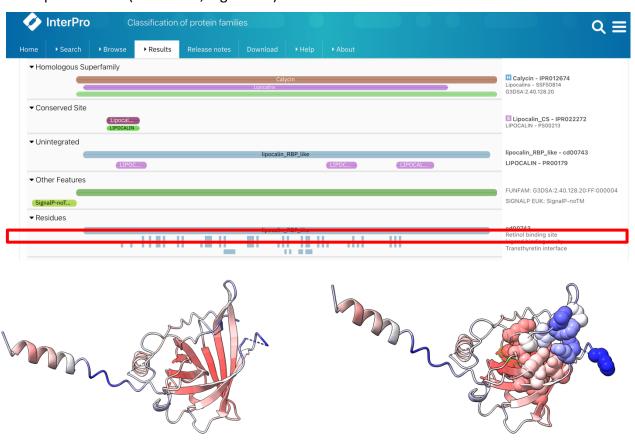


Danio_rerio_RBP4__NP_570995.1__0.36073
Rattus_norvegicus_RBP4__NP_037294.1__0.07804
Callithrix_jacchus_RBP4__XP_054099088.1__0.02431
Homo_sapiens_RBP4__NP_001310446.1__0.0025
Gorilla_RBP4__XP_018890983.3__0
Unknown_protein__BI546698.1_1__0

7) Sequen<u>ce Id</u>entity Heatmap

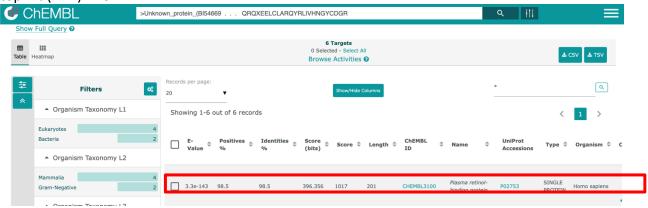


- 8) Ran trimmed alignment of 'unknown protein' from (5) on NCBI's protein blast against pdb database. Top 3 non-RBP4 hits and information are shown as follows:
- Sander vitreus Sandercyanin Fluorescent Protein (SFP), Chain A
 E-Value = 4e-08, Sequence Identity = 29.41%, PDB Identifier = 5EZ2, Method = X-ray diffraction, Resolution = 1.85 angstroms, Source Organism = Sander vitreus
- 2. Human Glycodelin, Chain A E-Value = 4e-06, Sequence Identity = 24.84%, PDB Identifier = 4R0B, Method = X-ray diffraction, Resolution = 2.45 angstroms, Source Organism = Homo sapiens
- 3. Human Apolipoprotein D, Chain A E-Value = 8e-06, Sequence Identity = 27.85%, PDB Identifier = 2HZQ, Method = X-ray diffraction, Resolution = 1.8 angstroms, Source Organism = Homo sapiens
- 9) One of 5 of the resulting pdb files were chosen for illustration. Using InterPro database for conserved domains and function, I found predicted retinol binding sites of my unknown protein (conserved residues between my MSA, boxed in red) and spacefilled with spheres (bottom right figure), and colored the sequence by pLDDT score (low = blue, high = red):



The spacefill clearly shows the retinol binding site within the beta barrel of the protein structure!

10) Yes, there are target associated assays and ligand efficiency data, as shown below for the top hit (RBP) in ChEMBL:



Associated Bioactivities

Associated Assays

