

# **SLiM Analysis in Human SALMs and Linx**

A Major Qualifying Project Report

Submitted to the Faculty of

WORCESTER POLYTECHNIC INSTITUTE

In partial fulfillment of the requirements for the

Degree of Bachelor of Science

In

Biology and Biotechnology

By

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December 15, 2016

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# Abstract

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Leucine rich repeat and Immunoglobulin domain (LIG) containing proteins play a critical role in protein-protein and intercellular interactions. While the extracellular domains of many of these proteins have been studied in depth, little is known about their intracellular domains, which can add to our knowledge of LIG protein functionality. Here, within the intracellular domains of two subfamilies of LIG molecules, the SALM and Linx subfamilies, sequences conserved over evolutionary time in four species: human, mouse, chicken, and elephant shark and representing putative functionally relevant Short Linear Motifs, SLiMs have been identified.

# Acknowledgements

I would like to thank Duff for being my advisor and for being the first to introduce me to the world of research and all it entails. I want to thank Duff for all of the career and life advice that I have obtained along the way, that has helped shape who I am and has helped me grow beyond my comfort zone in the classroom and in my life. This project has added immensely to my knowledge of many LIGs structure- function relationships. I am grateful to have been a part of the continuous process of discovery that is characteristic of Biology and science in general. Thank you Duff, for being there for me from Day 1 and believing in my capabilities from the outset. Your support has been invaluable to me. I am also extremely grateful to all the Biology professors and courses I have taken which have enhanced my undergraduate career and assisted me in identifying my future career aspirations.

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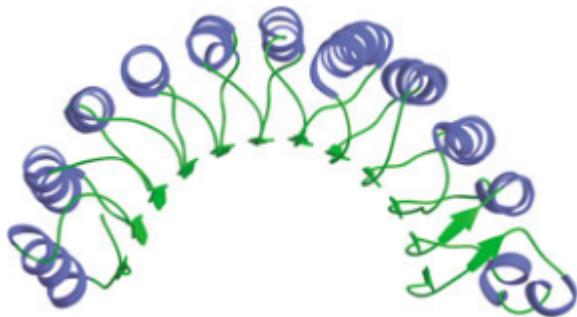
## **1. Introduction**

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Transmembrane proteins play an important role in intercellular and protein-protein signaling events that are essential to metazoan survival. Leucine rich repeats (LRR) and Immunoglobulin (Ig) domain containing proteins are thought to be one class of transmembrane proteins involved in these intercellular interactions. 36 human proteins with LRRs and Ig domains, called LIGS, have been identified and many have been shown to be involved in neuronal growth and synapse formation (Homma et al., 2008).

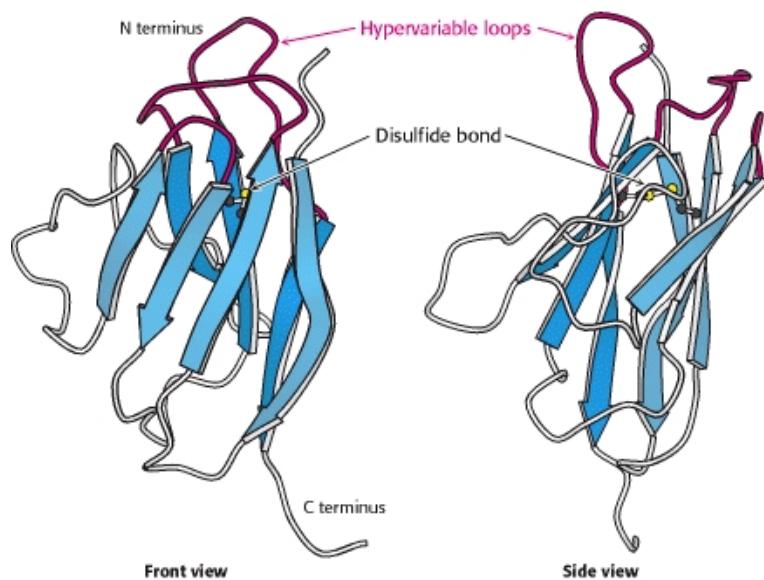
### ***Domain Architecture of LIGs***

Leucine Rich Repeats are between 20 to 30 amino acids and their N terminus has the conserved sequence LxxLxLxxN/CxL, with x representing any amino acid. These repeats are organized into loops, forming a horseshoe-like shape region. These LRR play a role as key binding sites for many protein-protein interactions to form. Figure 1 depicts the structure of these LRR (Hilling et al., 1999).



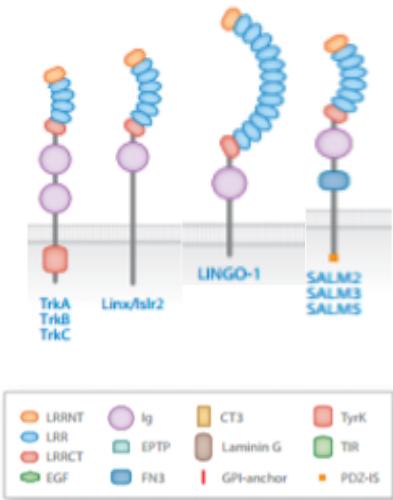
**Figure 1. LRR structure found in Rna1p**

The immunoglobulin domain is made up of a pair of beta sheets bonded by a disulfide bond where each beta sheet surrounds a hydrophobic core. At the N terminus, there are three loops called hypervariable loops that are present in antibodies and T cell receptors. This domain is one of the most prevalently encoded in the human genome with over 750 genes encoding proteins with at least one Ig domain (Berg et al, 2002) and is believed to play a role in protein-protein interactions. Figure 2 depicts the typical structure for Ig domains (Berg et al., 2002).



**Figure 2. Structure of Ig domain (adapted from Berg et al.,2002)**

All LIGs start with LRRs, presumably with their canonical horseshoe shape followed by an Ig domain(s), as can be seen below in Figure 3 (Wit et al., 2011). Different LIGs have variable numbers of each, but must contain both LRR and Ig domains to be considered a LIG. Some LIG proteins may also have additional domains as the SALMs contain a fibronectin domain as well.



**Figure 3. Domain Architecture of some LIG proteins (adapted from Wit et al., 2011)**

### **SALMs: LIG protein family serve as CAMs**

Cell adhesion molecules (CAM) play a major role in the formation of synapses and many are involved in the development and maintenance of these synapses (Missler et al., 2012). These transmembrane molecules can also act to hold neuronal membranes together. In addition, they often are necessary to allow for proper communication and contact between axons and dendrites and are involved in intercellular signaling (Missler et al., 2012). One family of LIG proteins that appear to function as adhesion molecules are known as Synaptic Adhesion Like Molecules, or SALMs. This family has 5 members, SALM1, SALM2, SALM3, SALM4, and SALM5 that are structurally similar to each other as they all contain extracellular Leucine rich repeats (LRR), Ig C2 type domains, and fibronectin type III domains (Wang et al., 2006). SALMs have been found to have both pre and post synaptic functions and play a role in neurite outgrowth and branching (Choi et al., 2016). These molecules, which are also called Lrfn because they contain the LRR and Fibronectin type III motifs, are transmembrane proteins that also contain a PDZ-domain

binding site (Wang et al., 2006). This site is about 4 amino acids in length with the sequence – X,-T,-X,-V at the carboxyl terminus with x being any amino acid, that binds PDZ domain proteins that play a critical role in protein-protein recognition, protein trafficking, localization, and cell signaling (Lee et al., 2004). These PDZ domain binding sites are found in SALM1-3, but not SALM4 and SALM5, suggesting a variety of functionalities between family members.

CAMs are important for proper neuronal and brain development and mutations in CAMs have been associated with neurological disorders and developmental damage (Seabold et al., 2012). Specifically, SALM1 and SALM5 have been associated with autism disorders and changes in neuronal morphology. Understanding the structural components of these proteins can give further insight into the functional properties of these molecules and how they contribute to the development of neurological disorders. While the extracellular domains of many of these proteins have been studied in detail and have been found to be highly conserved, little is known about the intracellular domains. Studying the intracellular domains of these molecules can add to our knowledge of their function. Specifically, identifying functionally important sequences or motifs in the intracellular domains, such as SLiMs for example, can direct further research efforts in understanding the function of many LIG proteins.

Short linear motifs or SLiMs are approximately 3-10 adjacent amino acid stretches in a protein's primary sequence that are believed to be functionally important for protein activity. SLiMs are thought to mediate between 15-40 % of protein-protein interactions and are therefore critical to understand their mechanistic contributions to cellular signaling events (Edwards et al., 2007). Unfortunately, these sequence elements are very difficult to identify due to their short length sequence. One potential method to identify SLiMs is to take a phylogenetic approach

through the identification of protein orthologs and analysis of conserved sequences among the orthologous molecules.

Identification of orthologs, which represent the same functional protein in different species, is a critical step towards understanding the function of a protein and can be used to identify functionally important sequences that are conserved over time. In this study, the objective was to take a comparative approach to identifying putative SLiMs within the intracellular domains of the SALM and Linx members of the LIG family to better understand their biological activity and mechanism of action. Sequence comparison of SALMs and Linx orthologs across a phylogenetically diverse set of vertebrates, including *Homo sapien* (human), *Mus musculus* (mouse), *Gallus gallus* (chicken), and *Callorhinichus milii* (elephant shark) species was performed. Orthologs were identified using BLAST, their intracellular domains identified using the online transmembrane prediction tool CCTOP, and their sequence conservation analyzed through protein alignments. Sequence conservation over time revealed intracellular motifs, or putative SLiMs, present within and among the SALMs and Linx orthologs, indicative of sequence units whose study is likely to reveal novel *in vivo* functions linked to the LIG family.

## 2. Materials and Methods

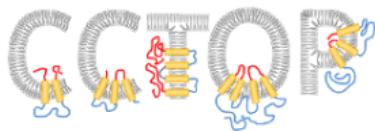
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### *Identification of protein sequences of LIGS*

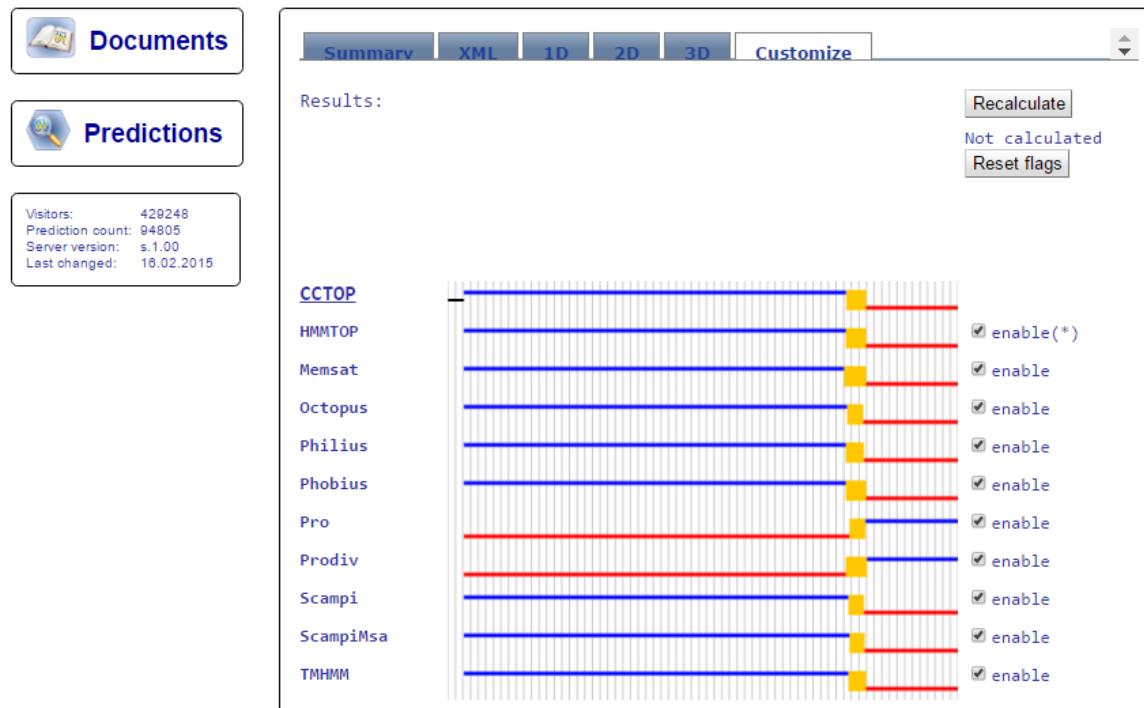
The NCBI protein database was utilized in order to identify the LIG protein sequences. Using their respective accession numbers, collected from the literature (Homma et al., 2008), the Homo sapien sequences were identified. The NCBI protein BLAST program was then used to identify orthologs of the SALM and Linx proteins in *Mus musculus*, *Gallus gallus*, and *Callorhinchus milii*, mouse (Tax ID10090), chicken (Tax ID 9031), and elephant shark (Tax ID7868), respectively. Putative SALM and Linx orthologs were defined as the match with the highest identity to the human protein query, which in a reciprocal BLAST also identified the initial human LIG query as the highest match.

### *Identification of the IC domain*

The program CCTOP was used to identify the transmembrane region for all orthologs, thereby allowing for the extracellular and intracellular regions of each protein to be defined. CCTOP combines output from 10 different programs to generate a consensus for the prediction of the transmembrane region of proteins. Below is a sample output of a CCTOP consensus and output (Figure 4).



## Constrained Consensus TOPology prediction server



**Figure 4. CCTOP based prediction and representation of transmembrane domains**

These programs all have the underlying assumption that the transmembrane domain is made up of stretch of hydrophobic amino acids and use amino acid physical properties, as well as structural information to generate a prediction of the position and length of the transmembrane domain and extracellular and intracellular regions by consequence.

### *Ortholog IC domain Sequence Alignment*

After obtaining the predictions of the extracellular, transmembrane, and intracellular regions of the SALM and Linx proteins, the IC domains were collected for each ortholog and sequence alignments built using Clustal Omega. Using only the IC domain, the orthologs were aligned and output in a fasta format. The program Boxshade was then used to create a highlighted protein

alignment of the sequence conservation between orthologs identified in each species. Using the CLUSTAL fasta output, the Boxshade program shades sequences of amino acids based on the level of agreement of the amino acid sequence between species when aligned. For the purpose of this study, conservation was set to shade residues at either 100% or 75 % identity between species as shown in Appendix D. The shading is black when the level of sequence identity is 100% between amino acids at a certain residue, while simple sequence conservation at a residue is shaded in gray. A consensus line is a part of the output that specifies which amino acids are conserved between the 4 species.

#### *Identification of Fingerprints*

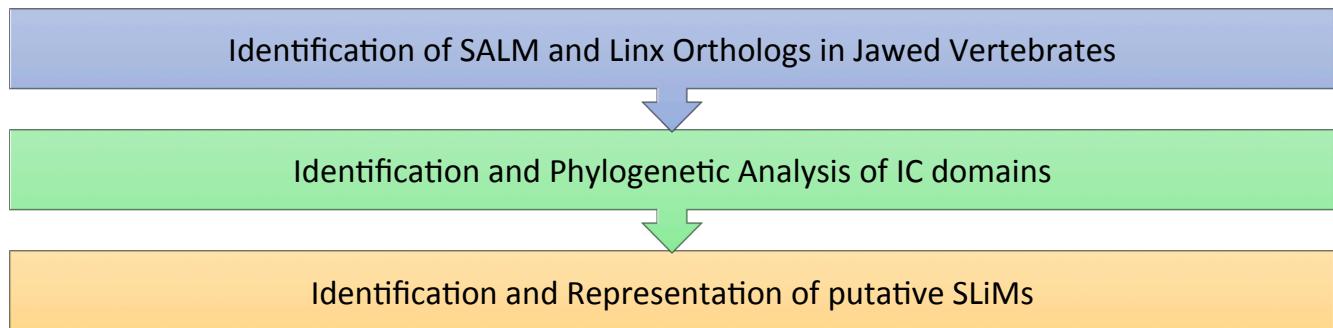
WebLogo was used to generate a graphical representation of putative IC domain SLiMs based on the overall sequence alignments built with CLUSTAL and Boxshade. The logo is built using a stack of symbols, with the height of the stack indicating sequence conservation at the given position, and with each stack representing a single position in the sequence. A custom color scheme was used in order to also provide information regarding conservation of biochemical properties of amino acids at each position (below).

**Chemistry (AA):** Color amino acids according to chemical properties.

Polar	G,S,T,Y,C	green
Neutral	Q,N	purple
Basic	K,R,H	blue
Acidic	D,E	red
Hydrophobic	A,V,L,I,P,W,F,M	black

### 3. Results

To better understand the biological activity and mechanism of action of human LIGs, a comparative approach to identifying putative SLiMs was undertaken. Of the 36 human LIGs, 17 were chosen for initial analyses of IC domain sequences with respect to location and length (see Appendix A). Of the 17, six proteins (SALMs 1-5 and Linx) were analyzed for SLiMs. To identify these short linear motifs, the human amino acid sequences for all proteins were found, followed by identifying orthologs in selected vertebrates (mouse, chicken, and elephant shark) that served as a basis for comparison of protein sequences to determine conservation. Conservation in the IC domain was the primary focus since little is known for these regions in LIGs as compared with their extracellular domains with the hope of gaining a deeper understanding of protein functionality. An overview of the experimental approach is presented in Figure 5.



**Figure 5. Phylogenetic analyses and SLiM identification in LIGs.**

#### 3.1 Identification of LIG family in Jawed Vertebrates

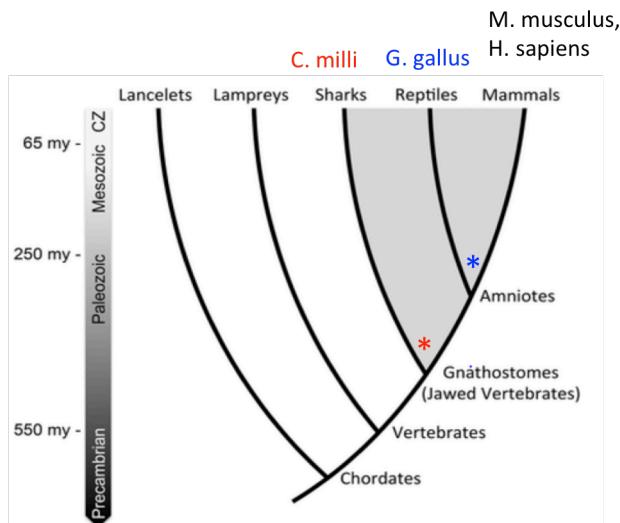
To understand and identify conserved sequences, the human amino acid sequences of 17 LIGs were obtained. Using the accession numbers for each LIG and the NCBI protein database, the

LIG protein sequence in homo sapiens were identified (Homma et al., 2008). Complete sequences for all 17 can be found in Appendix A.

In order to identify SLiMs in the human protein sequence, the idea of evolutionary conservation was utilized as a technique for motif discovery. Because SLiMs are often very short (4-10 amino acid residues) in length, they are difficult to identify (Edwards et. al., 2007). Therefore, a phylogenetic approach was undertaken to identify short sequences conserved in LIGs from species with different evolutionary relationships. With this methodology, short but highly identical amino acid sequences between species would suggest an important functionality of the protein that needed to be conserved over varying evolutionary distances/time.

Three species were identified on the basis of their evolutionary relationships to Homo sapiens, and each other, for this analysis, *Mus musculus*, *Gallus gallus*, and *Callorhinichus milii* or mouse, chicken, and elephant shark, respectively. Together the four species represent divergence times up to ~450 Mya and two branchpoints - divergences between cartilaginous fishes and bony vertebrates, and reptiles and mammals (Smith and Keinath, 2015; Schmutz and Grimwood, 2004; Venkatesh et. al., 2014; Waterston et. al., 2002). *Mus musculus* is the closest in genetic makeup to *Homo sapiens* as it has had the least amount of time to evolve as compared to the chicken and elephant shark genomes. The elephant shark has had the longest time to evolve and is the least similar to *Homo sapien* of the three species, while chicken falls in the middle of evolutionary distance relative to humans and sharks.

Figure 6 below illustrates the evolutionary relationships between these species in a phylogenetic tree.



**Figure 6. Phylogenetic tree for Homo sapien, Mus musculus, Gallus gallus, and Callorhinchus milii (modified from Smith and Keinath, 2015).** This phylogeneic tree depicts a timeline for evolution between chordates and vertebrates, including sharks, reptiles, and mammals. The timeline shows key events in the evolution of these major family groups.

After the LIG sequences were identified in *Homo sapiens*, NCBI BLAST and protein database was used to find the ortholog sequences of the SALM subfamily and LINX in the selected species - *Mus musculus*, *Gallus gallus*, and *Callorhinchus milii*. Using Protein Blast, the sequences of each SALM (1-5) and Linx were used to find their respective orthologs in the other species (see Materials and Methods for details). Percent similarity, high score and reciprocal BLASTing back to the *h. sapiens* database were used to determine the correct orthology. Table 1 below summarizes the presence of SALM and Linx orthologs in these species. While Linx was found in all species analyzed here, SALM3 was not found in chicken, and SALM4 was not found in either chicken or elephant shark.

Name	Human	Mouse	Chicken	Elephant Shark
LINX	Yes	Yes	Yes	Yes
SALM1	Yes	Yes	Yes	Yes
SALM2	Yes	Yes	Yes	Yes
SALM3	Yes	Yes	No	No
SALM4	Yes	Yes	No	Yes
SALM5	Yes	Yes	Yes	Yes

**Table 1.** Presence of Linx or SALMs in the human, mouse, chicken, and elephant shark genomes.

The sequence data for each LIG and the accession numbers can be found in Appendix B.

### 3.2 Identification of IC domains using CCTOP

After orthologs were identified, the protein sequences were analyzed using the prediction program CCTOP, which predicts the location of transmembrane domains, as well as the extracellular (EX) and IC domains. Green was used to highlight the EX domain, blue for the transmembrane domain, and red for the IC domain. A complete set of topology predictions for all SALM1-5 and Linx orthologs can be found in Appendix A and B. A compilation of the predicted IC domain length for each protein can be found in Table 2 below.

Name	Human	Mouse	Chicken	Elephant Shark
LINX	132	132	134	100
SALM1	234	233	79	226
SALM2	214	209	217	215
SALM3	96	97	NA	NA
SALM4	69	66	NA	121
SALM5	169	196	187	225

**Table 2.** Length of IC domains (amino acids) in SALM and Linx orthologs.

Within the SALM family, SALM1's IC domain was the largest across human, mouse, and elephant shark species, but not in chicken. As expected based on the relatively short evolutionary distance between human and mouse the length of the IC domains across orthologs was most similar between human and mouse species.

### **3.3 Alignment of ortholog sequences in Clustal Omega**

With the orthologs in hand and their IC domains defined, sequence alignments were performed to identify regions of conservation. The IC domains of each LIG and their respective orthologs were aligned using Clustal Omega to look for sequence conservation between species. The IC domain sequences for each LIG in each species that were used for the alignments can be found in Appendix C.

### **3.4 Identification of Motifs using Box shade**

To identify putative SLiMs, or conserved regions, the CLUSTAL alignments were analyzed with the program Boxshade. Parameters were varied to display regions of either 100% and 75% conservation and the consensus amino acid residue conserved between all represented sequences. The box shade outputs can be found in Appendix D. These alignments were then analyzed for putative SLiMs, conserved sequences across species, as well as between other members of the SALM family. Three motifs were identified in Linx based on visual analysis of sequences. Putative SLiMs were selected based on 100% conservation and were extended to include areas when only some conservation was found. Figure 7 displays the motifs for Linx. SALM1 was found to have six motifs, while SALM2 had four, SALM3 had one motif, SALM4 had two motifs, and SALM5 contained four motifs. For a list of all motifs identified and the sequence alignments refer to appendix E. In total, twenty motifs were identified.

**DA18**

KHPGKPYRLILRPQAPDPM  
KHPGKPYRLILRPQAPDPM  
KYQGKTYKLIMKAQNPDQM  
KYRGKTYKLIMKTQPPESL

**DA19**

FDPRASYLESEKSY  
FDPRASYLESEKSY  
FDPRASYLESEKNY  
FDPSASFQGSEKIY

**DA20**

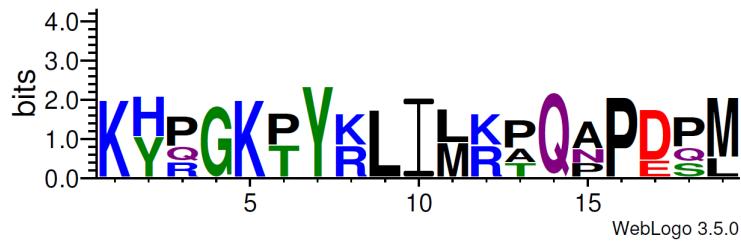
LAACSLVESQSKANQEEFEAGSEYSDRLPLGAEAVNIAQEINGNYRQ  
LAGCSLVESQSKANQEEFEAGSEYSDRLPLGAEAVNIAQEINGNYRQ  
VAASSMAESQSKANGEEFEVRSEYSDKLPLGAEAVTISQEINGNYRQ  
VVAESVPVSQTKANPEEFEACSEYSDRLPLGAEAVNISPEINGNYRQ

**Figure 7.** Putative Linx motifs.

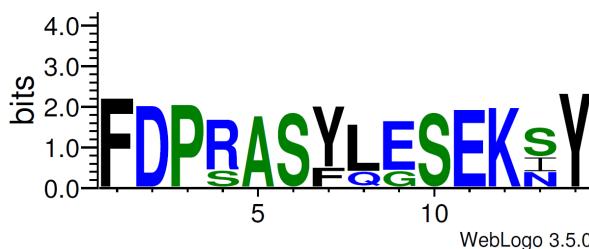
### 3.5 Identification and Representation of putative SLiMs

To provide a graphical representation of the degree and possible biochemical characteristics associated with regions conservation, the bioinformatics tool WebLogo was used to visualize all motifs. Figure 8 represents the graphical output for the three Linx motifs - DA18, DA19, and DA20.

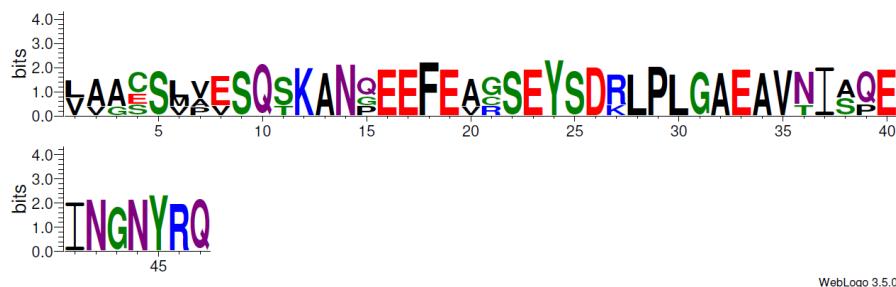
## DA18



## DA19



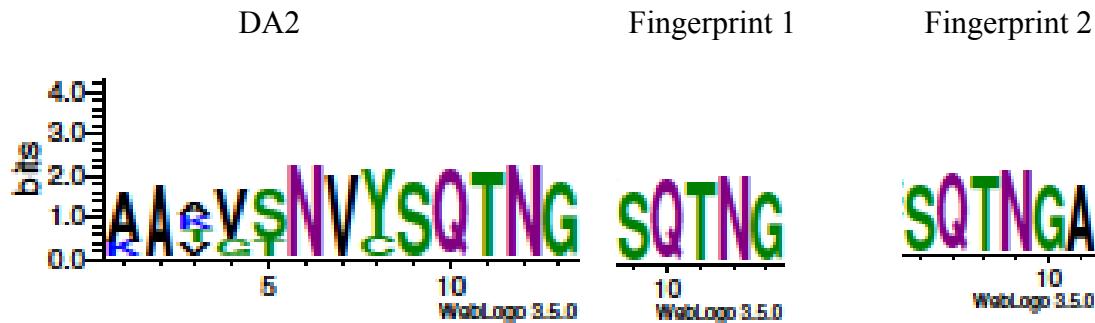
## DA20



**Figure 8. Putative SLiMs for Linx represented in WebLogo.**

The graphic shows the representation of specific amino acids found at every position in a given SLiM. Letter height represents the frequencies of specific residues at a certain position, while different colors are representative of specific chemical properties of the amino acids.

Conserved sequences were analyzed within motifs across the family and among different family members, specifically for the SALM family. In some cases submotifs present in a SLiM were defined as fingerprints. Two such fingerprints were identified from analysis of motifs in the SALM family and are shown in Figure 9.



**Figure 9. Identification of DA2 Fingerprints from SALM family**

The shortened amino acid sequence SQTNG in DA2 is found in all SALM homologs and was therefore identified as a fingerprint because its conservation across species, as well as across different SALM family members. Four additional fingerprints were identified through this analysis, including ESVV, ESTV, RYKV, and SFD. Motifs were identified that were also unique to their specific SALM and were not conserved between all family members. The sequence ESTV at the terminus of the IC domain was conserved in SALM1-3, but not SALM4-5. The sequence RYKV was found in SALMs1, 2, 4, and 5 but not SALM3. The fingerprints for each LIG can be found in Appendix E. Table 3-7 below summarize the fingerprints found in the SALM family in the 4 species.

Name	Human	Mouse	Chicken	Elephant Shark
SALM1	+	+	+	+
SALM2	+	+	+	+
SALM3	+	+	+	+
SALM4	+	+	+	+
SALM5	+	+	+	+

**Table 3. Prevalence of SQTNG Fingerprint in SALM1-5**

Name	Human	Mouse	Chicken	Elephant Shark
SALM1	+	+	+	+
SALM2	+	+	+	+
SALM3	-	-	-	-
SALM4	-	-	-	-
SALM5	-	-	-	+

**Table 4. Prevalence of ESTV Fingerprint in SALM1-5**

Name	Human	Mouse	Chicken	Elephant Shark
SALM1	-	-	-	-
SALM2	-	-	-	-
SALM3	+	+	-	-
SALM4	-	-	-	-
SALM5	-	-	-	-

**Table 5. Prevalence of ESVV Fingerprint in SALM1-5**

Name	Human	Mouse	Chicken	Elephant Shark
SALM1	+	+	+	+
SALM2	+	+	+	+
SALM3	-	-	-	-
SALM4	+	+	-	+
SALM5	+	+	+	+

**Table 6. Prevalence of RYKV Fingerprint in SALM1-5**

Name	Human	Mouse	Chicken	Elephant Shark
SALM1	+	+	+	+
SALM2	+	+	+	+
SALM3	-	-	-	-
SALM4	-	-	-	-
SALM5	-	-	-	-

**Table 7. Prevalence of SFD Fingerprint in SALM1-5**

As Table 3 portrays, the fingerprint SQTNG, is found in all SALM family members. The fingerprint ESTV is found in all species for SALM1 and SALM2, but not for the other family members. The fingerprint ESVV is found only in SALM3. The fingerprint RYKV is found in SALMs 1,2,4 (except for *Gallus gallus*), and 5 in all four species. The fingerprint SFD is only found in SALM1 and SALM2 for all species. The presence and distribution of unique fingerprints suggests both diversification and redundancy in functionality of the proteins over evolutionary time.

# Discussion

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Leucine rich repeat and immunoglobulin (LIG) containing proteins are thought to have a significant role in protein-protein and intercellular interactions. Like the SALM proteins, many are involved with cellular interactions during stages of neural development and function. While the extracellular domains of these proteins have been studied with some detail, little is known about the functionalities of their intracellular domains. Through the study of the IC domains of these LIG proteins, we can add to our knowledge of these protein's roles *in vivo* and the molecular mechanisms by which they act.

In this study, I identified evolutionarily conserved sequences that may represent functional SLiMs in the intracellular domains of two families of LIG molecules, SALM and Linx, that are conserved in four species: human, mouse, chicken, and elephant shark. These putative SLiMs are likely conserved between species because they provide specific essential or vital functions that an organism or species needs to survive. The sequences were found using the idea of evolutionary conservation because of the short length that would not be easily recognizable using standard search algorithms.

The fingerprints found were SQTNG, ESVV, ESTV, RYKV, and SFD. The SQTNG fingerprint's functionality can be emphasized in that it appears in every species and is conserved among all SALM family members. This prevalence suggests a functional importance and a clear target for further studies. The ESVV and ESTV fingerprints end the IC domain sequence in SALM3 and SALMs 1 and 2, respectively, across all species. They do not appear as motifs in

SALM4 and 5. These fingerprints are consensus PDZ-domain binding sites and suggest some degree of functional diversification between SALMs 1, 2, and 3 with SALMs 4 and 5.

The SLiMs and fingerprints founds can be used for further analysis and to further our understanding of SALM and Linx proteins *in vivo*. The SALM subfamily has been associated with autism disorders and changes in neuronal morphology. Understanding the relationship between these sequence elements and their contributions to protein function can give further insight into the role of these molecules and how they contribute to the development of neurological disorders.

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# Appendix A: Protein Sequence for Human LIGS

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## SALM1

Genbank:Q9P244.2

771aa

Leucine rich repeat and fibronectin type III domain containing 1 [Homo sapiens]

>gi|189028858|sp|Q9P244.2|LRFN1\_HUMAN RecName: Full=Leucine-rich repeat and fibronectin type III domain-containing protein 1; AltName: Full=Synaptic adhesion-like molecule 2; Flags: Precursor

MAPGPFSALLSPPPAALPFLLLWAGASRGQPCPGRCICQNVAPTLTMLCAKTGLLFVPPAIDRRVVEL  
RLTDNFIAAVRRRDFANMTSLVHHTLSRNTIGQVAAGAFADLRALRALHLDNSNRLAEVRGDQLRGLGNLR  
HLILGNNQIRRVESAAFDAFLSTVEDLDSYNNEALPWEAVGQMVNNTLDHNLIDHIAEGTFVQLH  
KLVRLDMTSNRLHKLPPDGLFLRSQGTGPKPPTPLTVSGGNPLHCNCCELLWLRRLTREDDLETCATPEH  
LTDRYFWSIPEEEFLCEPPLITRQAGGRALVVEGQAVSLRCRAVGDPPEPVHWVAPDGRLLGNSSRTRVR  
GDGTLDVTITLRDGTFTCIASNAAGEATAPVECVVPLPLMAPPPAAPPLTEPGSSDIATPGPGAN  
DSAERRLVAELTSNSVLIRWPAQRPVPGIRMYQVQYNSSVDDSLVYRMIPSTSQTFLVNDLAAGRAYD  
LCVLAVYDDGATALPATRVVGCVQFTTAGDPAPCRPLRAHFLG

GTMIIAIGGVIVASLVFIVLLMI

RYKVGDGDSRRVKGSRSLPRVSHVCSEQNGAGTGAAQAPALPAQDHYEALREVESQAAPAVAVEAKA  
MEAETASAEPVVVLGRSLGGSATSLCLLSEETSGEESRAAVGPRRSRSGALEPPTSAPPTLALVPGGAAAR  
PRPQQRYSFDGYGALFQSHSYPRRARRTKRHRSTPHLDGAGGGAAGEDGDLGLGSARACLAFTSTEWM  
LESTV

## SALM2

Leucine rich repeat and fibronectin type III domain containing 2 [Homo sapiens]

GenBank: AAI42617.1 (NP\_065788)

789 aa

[GenPept Identical Proteins Graphics](#)

>gi|148745628|gb|AAI42617.1| Leucine rich repeat and fibronectin type III domain containing 2 [Homo sapiens]

METLLGGLLAFGMAFAVVDACPKYCVCQNLSESLGTLCPSKGLFVPPDIDRRTVELRLGGNFIHISRQ  
DFANMTGLVDLTLRNTISHIQPFSFLDLESRLSLHLDNRPLSLGEDTLRGLVNLQHLIVNNNQLGGIA  
DEAFEDFLTLEDLDSYNNLHGLPWDSVRRMVNLHQSLDHNLIDHIAEGTFADLQKLARLDLTSNRLQ  
KLPPDPFARSQASALTATPFAPPLSFSFGGNPLHCNCCELLWLRRLERDDDLETGSPGLKGRYFWHVR  
EEEVCEPPLITQHTHKLLVLEGQAATLKCKAIGDPSPLIHVVAPDDRLVGNSSRTAVYDNGTLDIFITT  
SQDGAFTCIAANAAGEATAMVEVSIVQLPHLSNSTSRTAPPKSRLSDITGSSKTSRGGGGGGEPPKS  
PPERAVLVSEVTTTSALVKWSVSKSAPRKMYQLQYNCSDDEVLIYRMIPASNKAFFVNNLVSGTGYDLC  
VLAMWDDTATTATNIVGCAQFFTAKADYPQCQSMHSQI

LGGTMILVIGGIIVATLLVFIVLMV

RYKVCNHEAPSKMAAAAVSVYQTNAGAQPPPSAAGAPPQGPKVVVRNELLDFTASLARASDSSSSSS  
LGSGEAAGLGRAPWRIPPSAPRPKPSLDRLMGAFASLDLKSRKEELLSRTPAGRAGTSARGHHSRDREP  
LLGPPAARARSLLPLPLEGKAKRSHSFDMGDFAAAAAGGVVPGGYSPRKVSNIWTKRSLVNGMLLPFEE  
SDLVGARGTFGSSEWVMESTV

#### SALM3

Leucine rich repeat and fibronectin type III domain containing 3 [Homo sapiens]

GenBank: AAH03578.1 (NP\_078785.1)

628 aa

>gi|13097762|gb|AAH03578.1| Leucine rich repeat and fibronectin type III domain containing 3 [Homo sapiens]  
MAILPLLCLLPLAPASSPPQSATPSPCPRRCRCQTQLPLSVLCPGAGLLFVPPSLDRRAELRLADNF  
IASVRRRDLANMTGLLHLSSLRNTIRHVAAGAFADLRALRALHLDGNRLTLSLGEGLRGLVNLRHLILSN  
NQLAALAAGALDDCAETLEDLDLSYNNEQLPWEALGRLGNVNTLGLDHNLASVPAGAFSRLHKLARD  
MTSNRLTTIPPDPLFSRLPLLARPRGSPASALVLAFFGNPLHCNCLEVWLRLAREDDLEACASPPALGG  
RYFWAVGEEEFVCEPPVVTHRSPPLAVPAGRPAALRCRAVGDPPEPRVRWVSPQGRLLGNSSRARAFPN  
LELLVTEPGDGGIFTCIAANAAGEATAAELTVGPPPPQLANSTCDPPRDGDPDALTPPSAASASAKV  
ADTGPPTDRGVQVTEHGATAALVQWPDQRPIPGRIMYQIQYNSSADDILVYRMIPAESRSFLTDLASGR  
TYDLCVLAVYEDSATGLTATRPGCARFSTEPALRPCGAPHAPF  
LGGTMIIALGGVIVASVLVFIFVLL  
MRYKVHGGQPPGKAKIPAPVSSVCSQTNGALGPTPTPAPPAPEPAALRAHTVVQLDCEPWGPHEPVGP

#### SALM4

Leucine rich repeat and fibronectin type III domain containing 4 [Homo sapiens]

GenBank: AAH15581.2 (NP\_076941)

635 aa

#### GenPept Identical Proteins Graphics

>gi|22800525|gb|AAH15581.2| Leucine rich repeat and fibronectin type III domain containing 4 [Homo sapiens]  
MAPPLLLLLASGAAACPLPCVCQNLSESLSTLCAHRGLLFVPPNVDRRTVELRLADNFIQALGPPDFRN  
MTGLVDLTLRNAITRIGARAFGDLESRLSHLDGNRLVELGTGSLRGPVNLQHLILSGNQLGRIAPGAF  
DDFLESLEDDLSYNNLRQVPWAGIGAMPALHTLNLDHNLIDALPPGAFAGQLGQLSRLDLTSNRLATLAP  
DPLFSGRDAEASPAPLVLFSGNPLHCNCCELLWLRRLARPDDLETASPPGLAGRYFWAVPEGEFSCEP  
PLIARHTQRLWVLEGQRATLRCRALGDPAPTMHWVGPDRLVGNSSRARAFPNGTLEIGVTGAGDAGGY  
T  
CIATNPAGEATARVELRVLALPHGGNSAEGGRPGPSDIAASARTAAEGETLESEPAVQVTEVTATSGL  
VSWGPGRPADPVWMFQIQQYNSSEDETLYRIVPASSHHFLKKHLVPGADYDLCLLALSPAAGPSLTATR  
LLGCAHFSTLPASPLCHALQAHV  
LGGTLTVAVGGVLVAALLVFTVALLV

RGRGAGNGLPLKLSHVQSQTNGGPSPTKAHPPRSPPPRPQRSCSLDLDAGCYGYARRLGGAWARRSH  
SVHGGLLGAGCRGVGGSAERLEESVV

### Linx

#### Immunoglobulin superfamily containing leucine-rich repeat 2 [Homo sapiens]

GenBank: ([NP\\_065902.1](#))AAI52430.1

745 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|156230954|gb|AAI52430.1| Immunoglobulin superfamily containing leucine-rich repeat 2 [Homo sapiens]  
MFPLRALWLVWALLGVAGSCPEPCACVDKYAHQFADCAYKELREVPEGLPANVTTLSANKITVLRRGA  
FADVTQVTSWLWLAHNEVRTVEPGALAVSQLKNLDSLHNFISFPWSDLRNLSALQLKMNHNRLGSLPR  
DALGALPDRLSRINNNRRTLAPGTFDALSAHLQLYHNPFCGCGLVWLQAWAASTRVSLPEPDSIA  
CASPPALQGVVPVYRLPALCAPPsvHLSAEPPLEAPGTPLRAGLAFVLHCIADGHPTPRLQWQLQIPGGT  
VVLEPPVLSGEDDGVGAAEEGEGEGDGDLTTQATPPTPAPAWPAPPATPRFLALANGSLLVPLLSAKEA  
GVYTCRAHNELGANSTSIRVAVAATGPPKHAPGAGGEPDGQAPTSERKSTAKGRGNSVPSKPEGKIKGQ  
GLAKVSILGETETEPEEDTSEGEAAEDQILADPAEEQRCNGDPSRYVSNHAFNQSAELKPHVFELGVIA  
LDVAEREARVQLTPLAARWGPGPAGGAPRPGRPLLLYLCPAGGGAAVQWSRVEEGVNAYWFRGL  
RP  
GTNYSVCLALAGEACHVQVFSTKKELPSL  
LVIVAVSVFLLVLATVPLLGAAC  
CHLLAKHPGKPYRLILRPQAPDPMEKRIAADFPRASYLESEKSYPAGGEAGGEEPEDVQGEGLDEDAEQG  
DPSGDLQREESLAACSLVESQSKANQEEFEAGSEYSDRLPLGAEAVNIAQEINGNYRQTAG

#### LRIT2 protein [Homo sapiens]

GenBank: AAI44476.1

560 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|219518288|gb|AAI44476.1| LRIT2 protein [Homo sapiens]  
MASVFHYFLLVLVFLDTAAQPFCLPGCTCSEESGRTLQCTSVLGKIPGNLSEEFKQVRIENSPLFEM  
PQGSFINMSTLEYLWLNFNNISVIHLGALEHLPRELRLGENKLCSVWTAFRATPLRVLDLKRNKID  
ALPELALQFLVSLTYLDLSSNRRTVVSKSFLNPAYQKCRQDCGAEILSSLVVALHDNPWVCDCLR  
LVQFVKSITLPVILVNSYLICQGPLSKAGQLFHETELSACMKPQISTPSANITIRAGQNVTLRCLAQASP  
SPSIAWTYPLSMWREFDGLGGKHLTPVLTSSGTEDTALSELAIPAAHLVDSGNYTCMASNSIGKSNLVI  
SLHVQPAQALHAPDSLSIPSEGNAIDLRRVKQTVHGILLEWLAADTSKEEWFTLYIASDEAFRKEVVH

IGPGINTYAVDDLPGTKYEACLSLEGQPPHQGQCVAFVTGRDAGGLEAREH  
LLHVTVVLVLCVLLAVPGAYAWAAQGPC  
SCSKWVLRGCLHRRKAPSCTAAPQSKEGIDTEGDKEKGTEDNS

LRIT3 protein [Homo sapiens]

GenBank: AAI04038.1

552 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|74355215|gb|AAI04038.1| LRIT3 protein [Homo sapiens]  
MPLLRTLDLHNNKITSVPNEALRYLKNLAYLDLSSNRLTTPDFLENWTHLVSTPSGVLDLSPSRIILG  
LQDNPWFCDCCHISKMIELSKVVDPAIVLLDPLMTCEPERLTGILFQRAELEHCLKPSVMTSATKIMSAL  
GSNVLLRCDATGFPTPQITWTRSDSSPVNYTVIQESPEEGVRWSIMSLTGIISSKDAGDYKCKAKNLAGMS  
EAVVTVTVLGITTPIPPDTERTGDHPEWDVQPGSGRSTSASSYLWSSSFPTSSFSASTLSPPST  
ASFSLSPFSSSTVSSTTLSTSISASTTMANKRSFQLHQGGKRLKVAKNGSKLPPASTSKKEELALLDQ  
TMLTETNATIENLRVVSETKESVTLMWNMINTTHNSAVTVLYSKYGGKDLLLNADSSKNQVTIDGLEPG  
GQYMACVCPKGVPPQKDQCITFSTERVEGDDSQWS  
LLLVTSTACVVILPLICFLL  
YKVCKLQCKSEPFW  
EDDLAKETYIQFETLFPRSQSVGELWTRSHRDDSEKLLCSRSSVESQVTFKSEGRPEYYC

Leucine rich repeat containing 24 [Homo sapiens]

GenBank: AAI11068.1

513 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|83405784|gb|AAI11068.1| Leucine rich repeat containing 24 [Homo sapiens]  
MALRAPALLPLLLLLPLRAAGCPAACRCYSATVECGALRLRVVPLGIPPGTQTLFLQDNNIARLEPGAL  
APLAALRRLYLHNNSLRALEAGAFRAQPRLLELALTSNRLRGLRSGAFVGLAQLRVLYLAGNQLARLLDF  
TFLHPRLQELHLQENSIELLEDQALAGLSSLALLDSRNQLGTISREALQPLASLQVRLTENPWRCDC  
ALHWLGAWIKEGGQRLLTSRDRKIMCAEPPRLALQSLLDVSHSSLICIPPSVHVQPLELTANLGEDLRVA  
CQASGYPQLVTWRKVPQPREGPRPRAQAQLEGGGLGGHSASDTGSGMLFLSNITLAHAGKYECEASNA  
GGAARVPFLLVNASRQQPQQPAQPPPAARPAGSEPRPEAGSMAFRALGVATQ  
TAIAAAIALLALTALLVAMI  
CRRRRRRKARGPPGEGALFVNDYLDGPCTFAQLEELRDERGHEMFVINRSKPLFAEGPAEAPAD  
CGPAQGAGPGLRVPPPVAEYIHC

LRRC4C protein [Homo sapiens]

GenBank: AAH41374.3

640 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|73909151|gb|AAH41374.3| LRRC4C protein [Homo sapiens]

MLNKMTLHPQQIMIGPRFNRALFDPLLVLLALQLLVAGLVRAQTCPSVCSCSNQFSKVICVRKNLREV  
PDGISTNTRLLNLHENQIQIICKVNSFKHLRHLEILQLSRNHIRTIEIGAFNGLANLNTLEFDNRLLTIP  
NGAFVYLSKLKELWLRNNPIESIPSYAFNRIPSLRRLDLGELKRLSYISEGAEGLSNRLYLNLCNL  
EIPNLTPLIKLDELDLSGNHLSAIRPGSFQGLMHLQKLWMIQSQIVIERNADNLQSLVEINLAHNNLT  
LLPHDLFTPLHHLERIHLHHNPWCNCIDLWSWWIKDMAPSNTACCACNTPPNLKGRYIGELDQNYFT  
CYAPVIVEPPADLNVTTEGMAAEELKCRASTSLTSVSWITPNGTVMTGAYKVRIAVLSDGTLNFTNVTVQD  
TGMYTCMVSNSVGNTTASATLNVTAATTPFSYFSTVTETMEPSQDEARTTDNNVGPTPVVDWETTNVT  
TSLTPQSTRSTEKTFTIPVTDINSGIPGIDEVMKTTK  
IIIGCFVAITLMAAVMLVIF  
**YKMRKQHHRQNHH**  
**APTRTVEIINVDDITGDTPMESHLPMPAIEHEHLNHYNSYKSPFNHTTVNTINSIHSSVHEPLLIRMN**  
**SKDNVQETQI**

Leucine rich repeat containing 4 [Homo sapiens]

GenBank: AAI11562.1

653 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|109730363|gb|AAI11562.1| Leucine rich repeat containing 4 [Homo sapiens]

MKLLWQVTVHHHTWNAILPFVYLTAQVWILCAAIAAAASAGPQNCPSCSCSNQFSKVVCTRRGLSEVP  
QGIPSNTRYNLMMENNIQMIQADTFRHLHHLEVLQLGRNSIRQIEVGAFNGLASLNTLEFDNWLTVPIS  
GAFEYLSKLRELWLRNNPIESIPSYAFNRVPSLMRRLDLGELKKLEYISEGAEGLFNLKYLNLCNIKD  
MPNLTPLVGLEELEMSGNHFPEIRPGSFHGLSSLKKLWVMNSQVSLIERNADGLASLVELNLAHNLLSS  
LPHDLFTPLRYLVELHLHHNPWCNCDCDILWLAWWLREYIPTNSTCCGRCHAPMHMRGRYLVEVDQASFQ  
C  
SAPFIMDAPRDLNISEGRMAELCRTPPMSSVKWLLPNGTVLSHASRPRISVLDGTNFHVLLSDTG  
VYTCMVTNVAGNSNASAYLNVSTAELNTSNYSFFTIVTETTEISPEDTRKYKPVPTTSTGYQPAYTTS  
TTVLIQTTTRVPKQVAVPATDTDKMQTSLDEVMKTTK  
IIIGCFVAVTLLAAAMLIVF  
**YKLRKRHQQRSTV**  
**TAARTVEIIQVDEDIPAATSAAATAAPSGVSGEGA VVLPTIHDHINYNTYKPAHGAHWTENSLGNSLHPT**  
**VTTISEPYIIQTHTKDKVQETQI**

Leucine rich repeat neuronal 1 [Homo sapiens]

GenBank: AAH34947.1

716 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|23273823|gb|AAH34947.1| Leucine rich repeat neuronal 1 [Homo sapiens]

MARMSFVIAACQLVLGLLMTSLTESSIQNSECPLCVCEIRPWFTPQSTYREATTVDCNDLRLTRIPSNL  
SSDTQVLLQSNNIAKTVDELQQLFNLTELDFSQNNFTNIKEVGLANLTQLTTLHLEENQITEMTDYCLQ  
DLSNLQELYINHNQISTISAHAFAGLKNNRLHLNSNKLKVIDSRWFDPNLEILMIGENPVIGILDMN  
FKPLANLRSVLVLAGMYLTDIPGNALVGLDSLESLSFYDNKLVKVPQLALQKVPSLKFLDLNKNPIHKIQE  
GDFKNMLRLKELGINNMGELVSVDRYALDNLPELTKEATNNPKLSYIHRLAFRSVPVAESLMLNNNALN  
AIYQKTVESLPNLREISIHSNPLRCDCVIHWINSNKTNIRFMEPLSMFCAMPPEYGHQVKEVLIQDSSE  
QCLPMISHDSFPNRLNVDIGTTVFLDCRAMAEPEPEIYWVTPIGNKITVETLSDKYKLSSEGTLIESNIQ  
IEDSGRYTCVAQNVQGADTRVATIKVNGETLDGTQVLKIYVKQTESHSILVSWKVNSNVMTSNLKWWSSAT  
MKIDNPHTYTARVPVDVHEYNLTHLQPSTDYEVCLTVSNIHQQTQKSCVNVTKNAAFAVDISDQETST

A

LAAVMGSMFAVISLASIAV

YFAKRFKRKNYHHSKKYMQKTSSIPLNELYPPLINLWEGDSEKDKGSA

DTKPTQVDTSRSYMW

Leucine rich repeat neuronal 2 [Homo sapiens]

GenBank: AAH68541.1

713 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|46249796|gb|AAH68541.1| Leucine rich repeat neuronal 2 [Homo sapiens]

MRLLVAPLLLAWVAGATAAVPVVPWHVPCPPQCACQIRPWYTPRSSYREATTVDCNDLFTAVPPALPAG  
TQTLQQNSIVRVDQSELGYLANLTELDSQNSFSDARDCDFHALPQLLSHLEENQLTRLEDHSFAGL  
ASLQELYLNHNQLYRIAPRAFSGLSNLLRLHLNSNLLRAIDSRRWFEMLPNLEILMIGGNKVDAILDNNFR  
PLANLRSVLVAGMNLREISDYALEGLQSLESLSFYDNQLARVPRRALEQVPGLKFLDLNKNPLQRVGPGD  
FANMLHLKELGLNNMEELVSIDKFALVNLPELTKLIDITNNPRLSFIHPRAFHLPQMELMLNNNALSAL  
HQQTAEQLNLQEVLGNPIRCDCVIRWANATGTRVRFIEPQSTLCAEPPDLQRLPVREVPFREMTDH  
LPLISPRSFPPSLQVASGESMVLHCRALAEPEPEIYWVTPAGLRLTPAHAGRRYRVYPEGTLELRRVTAE  
EAGLYTCVAQNLVGADTKTVVVGRALLQPGRDEGQGLELRVQETHPYHILLSWVTPPNTVSTNLTWSS  
ASSLRGQGATALARLPRGTHSYNITLLQATEYWACLQVAFADAHTQLACVWARTKEATSCHRALGDRP  
GLIAILALAVLLAAGLAHLG

TGQPRKGVGGRPLPPAWAFWGWSPPSVRVVSAPLVLWPWNPGRKLPSS

## EGETLLPPLSQNS

Leucine rich repeat neuronal 3 [Homo sapiens]

GenBank: AAH35133.1

708 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|23242678|gb|AAH35133.1| Leucine rich repeat neuronal 3 [Homo sapiens]

MKDMPLRIHVLLGLAITTLVQAVDKVDCPRLCTCEIRPWFTPRTSIYMEASTVDCNDLGLLTFPARLPAN  
TQILLQTNNIAKIEYSTDFPVNLTSLDLSQNNLSSVTNINVKKMPQLLSVYLEENKTELPEKCLSELS  
NLQELYINHNLLSTISPGAFIGLHNLLRHLNSNRQMINSKWFDALPNLEILMIGENPIIRIKDMNFKP  
LINLRSLSVIAGINLTEIPDNALVGLENLESISFYDNRLIKVPHVALQKVVLKFLDLNKNPINRIRRGDF  
SNMLHLKELGINNMPELISIDSLAVDNLPLRKIEATNNPRLSYIHPNAFFRLPKLESMLNSNALSALY  
HTGIESLPNLKEISIHSNPIRCDCVIRWMNNMKTNIRFMEPDSLFCVDPPFQGQNVRQVHFRDMMEICL  
PLIAPESFPSNLNVEAGSYVSFHCRATAEPQPEIYWITPSGQKLLPNTLTDKFYVHSEGTLINGVTPKE  
GGLYTCIATNLVGADLKSVMIKVDGSPQDNNGSLNIKIRDIHANSVLVSWKASSKILKSSVKWTAFVKT  
ENSHAAQSARIPSDVKVYNLTHLN PSTEYKICIDIPTIYQKNRKKCVNVT KGLHPDQKEYEKNNTTT  
LMACLGGLLGIIGVICLISCLS

PEMNCDGGHSYVRNYLQKPTFALGELYPPLINLWEAGKEKSTS LKV KATV

IGLPTNMS

Neurotrophic tyrosine kinase, receptor, type 1 [Homo sapiens]

GenBank: AAI44240.1

790 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|219841840|gb|AAI44240.1| Neurotrophic tyrosine kinase, receptor, type 1 [Homo sapiens]

MLRGGRGQLGWH SWAAGPGSLLAWLILASAGAAPCPDACC PHGSSLRCTR DGA LDSLHH LPGA ENLT  
E  
LYIENQQHLQHLELRDLRGLGELRNLTIVKSGLRFVAPDAFHTPRLSRLNLSFNALESLSWKT VQGLSL  
QELVLSGNPLHCSCALRWLQRWEEGLGGVPEQKLQCHGQGPLAHMPNASCGVPTLK VQVPNASVDVGD  
D  
VLLRCQVEGRGLEQAGWILTELEQSATVMKSGGLPSLGLTLANVTS DLRKNVTCWAENDVGRAEVSVQ  
V  
NVSFPASVQLHTAVEMHHWCIPFSVDGQPAPSLRWFNGSVLN NETSFIFTEFLEPAANETVRHGCLRLNQ  
PTHVNNGNYTLLAANPFGQASASIMA AFMDNPFEFN PEDPIPDTNSTSGDPVEKKDET PFG  
VSVAVGLAVFACFLSTLLLVL  
NKCGRRNKG INRPAVLAPEDGLAMSLHFMTLGGSSLSPTEGKGSGLQGHIIENPQY

FSDACVHHIKRRDIVLKWELEGAFGVFLAECNLLPEQDKMLVAVKALKEASESARQDFQREAELLTM  
LQHQHIVRFFGVCTEGRPLLMVFEMRHDNLRFRLSHGPDAKLLAGGEDVAPGPLGLGQLLAVASQVA  
A  
GMVYLAGLHFVHRDLATRNCLVGQGLVVKIGDFGMSRDIYSTDYYRVGGRTMLPIRWMPPEISLYRKFTT  
ESDVWSFGVVLWEIFTYGKQPWYQLSNTAIDCITQGRELERPRACPPEVYAIMRGCWQREPQQRHSIKD  
VHARLQALAQAPPVYLDVLG

Neurotrophic tyrosine kinase, receptor, type 2 [Homo sapiens]

GenBank: AAH31835.1

477 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|21594337|gb|AAH31835.1| Neurotrophic tyrosine kinase, receptor, type 2 [Homo sapiens]  
MSSWIRWHGPAMARLWGFCWLVVGFWRAAFACPTCSKCSASIWCSDPSPGIVAFRLEPNSVDOPENITE  
IFIANQKRLEIINEDDVEAYVGLRNLTVDSGLKFAHKAFKLNSNLQHINFTRNKLTSRKHFRHLDL  
SELILVGNPFTCSCDIMWIKTLQEAKSSPDTQDLYCLNESSKNIPLANLQIPNCGLPSANLAAPNLTVEE  
GKSITLSCSVAGDPVPNMYWDVGNLVSKHMNETSHTQGSLRITNISSDDSGKQISCVAENLVGEDQDSVN  
LTVHFAPTITFLESPTSDHHWCIPFTVKGNPKPALQWFYNGAILNESKYICTKIHTVNHTHEYHGCLQLDN  
PTHMNNGDYTLIAKNEYGKDEKQISAHFMGWPGRIDGANPNYPDVYEDYGTAAANDIGDTTNRSEIPST  
DVTDKTGREHLS  
**VYAVVVIASVVGFCLLVMLFLL**  
**KLARHSKFGMKGFVLFHKIPLDG**

Neurotrophic tyrosine kinase, receptor, type 3 [Homo sapiens]

GenBank: AAH13693.1

612 aa

[GenPept](#) [Identical Proteins](#) [Graphics](#)

>gi|15489168|gb|AAH13693.1| Neurotrophic tyrosine kinase, receptor, type 3 [Homo sapiens]  
MDVSLCPAKCSFWRIFLLGSVWLDYVGSQLACPANCVCSKTEINCRRPDDGNLFPLLEGQDSGNNSGNAS  
INITDISRNITSIHIENWRSLHTLNAVDMELYTGLQKLTIKNSGLRSIQPRAFAKNPHLRYINLSSNRLT  
TLSWQLFQTLSRELQLEQNFFNCSCDIRWMQLWQEAKLNSQNLYCINADGSQLPLFRMNISQCDLP  
EISVSHVNLTVREGDNAVITCNGSGSPLPDVDWIVTGLQSINTHQTNLNWNVHAINLTLVNVTSEDNGF  
TLTCIAENVVGMSNASVALTVYYPPRVSLEEPELRLEHCIEVVVRGNPPTLHWLHNGQPLRESKIHV  
EYYQEGERSEGCLFNKPHTHNNNGNYTLIAKNPLGTANQTINGHFLKEPFESTDNFILFDEVSPTPPIT  
**VTHKPEEDTGF**  
**VSIAVGLAAFACVLLVVLFVMI**

NKYGRRSKFGMKGPVAVISGEEDSASPLHHINHGITT  
PSSLDAGPDTVIGMTRIPVIENPQYFRQGHNCHKPDTWVFSNIDNHGILNLKDNRDHLVPSTHYIYEEP  
EVQSGEVSYPRSHGFREIMLNPISLPGHSKPLNHGIYVEDVNVYFSKGRHGF

## Appendix B: Protein Sequence for other Species LIGS

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### Linx

#### **immunoglobulin superfamily containing leucine-rich repeat protein 2 isoform a [Mus musculus]**

NCBI Reference Sequence: NP\_001155007.1

[GenPept Identical Proteins Graphics](#)

protein	1	fasta
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>gi|238859603|ref|NP\_001155007.1| immunoglobulin superfamily containing leucine-rich repeat protein 2 isoform a [Mus musculus]

MHSPFLPTATATDARSSLRLSPESGDRLAAPQHHTASQRAAGVTMGPFGALCLAWALLGVVRACPEPCAC  
VDKYAHQFADCAYKELREVPEGLPANVTTLSLSANKITVLRRGAFVNVTQVTSWLWLAHSEVRTVESGALA  
VLSQLKNLDSLHNLISNFPWSDLRNLSALQLLKMNHNRLGSLPRDALGALPDLRSLRINNNRLRTLEPGT  
FDALSALSHLQLYHNPFHCSCLGVWLQAWAASTRVSLPEPDSIACASPPPELQGVPVHRLPALPCAPPSSVR  
LSAEPPPEAPGTPLRAGLAGMLHCVAEGHPTPRLQWLQIPGGTVVLVPPVLSKEEDGGDKVEDGEKGDG  
EDLPTQTEAPTPTPAPAWPAPPATPRFLALANGSLLVPLLSAKEAGIYTCRAHNELGTNSTSLRVTVAAA  
GPPKHAPGTGEEPDAQVPTSERKATTKGRSNVLPFKPEGKTKGQGLARVSVLGEIEAELEETDEGEQME  
GQIPADPMGEKHCGHGDPSRVSNHAFNQSSDLKPHVFEGLVIALDVAEREARVQLTPLAARWGPGPDGA  
SGARRPGRRPLRLYLCPAGGGTAVQWSRVEEGVNAYWFRGLRPGTNYSVCLALAGEACHVQVFSTKK  
E

LPSL

LVIVTVSVFLLVLATVPLLGAAC

CHLLAKHPGKPYRLILRPQAPDPMEKRIAADFDPASYLESEK

SYPARGEAGGEEPEEVPEEGLDEDVEQGDPSGDLQREESLAGCSLVESQSKANQEEFEAGSEYSDRLPLG  
AEAVNIAQEINGNYRQTAG

#### **immunoglobulin superfamily containing leucine-rich repeat protein 2 precursor [Gallus gallus]**

NCBI Reference Sequence: NP\_001038132.1

[GenPept Identical Proteins Graphics](#)

>gi|113206126|ref|NP\_001038132.1| immunoglobulin superfamily containing leucine-rich repeat protein 2 precursor [Gallus gallus]

MAPALWLWALLGSARACPEPCACVDKYAHQFADCAYKDLQVVTGLPSNVTLSANKITALQRRSF  
VEVTQVTSWLAHNEIRAIEPGAFAILVQLKNLDISHNQIVDFPWQDLYNLSALQLKMNNHMAVVPQG  
AFHTLKDRLSLRINNNKFTTLAEGIFDSLSSLSHLQIYNNPFECSCKLQWLKKMDSTLISIPEKESITC  
SLPEQLRGVEVGKIPDTQCTSPVQLTYYPNLDTTEFDGFTLHCAVTGAPPVSWKIRTSSQTLEL  
SGSPSESAGKDPPRQDPERFLVFKNGTLVIPHSKREEGTYTCLATNEMGSNQTSVNVAVAGSQKYPLQP  
GRDPTGGKAQPGDKPGAKGAKNSVLTDPERSKPLSPTRQSQQPSAAGMEPTGDGKVPFQLPPFEKKCGS  
MPTSRVYISNHAFNQSGDFKQHTFDLGVIALDVSERDARVQLPTVQPDKVHLRMLYLCQESSRGHALVQ  
WSKIEGVNSYWFQGLKPGTNYSVCLTYLGEDCQVQVVFTTKKEIPS  
LIIIVVSIFLLLATLPLMGATWCPLL  
SKYQGKTYKLIMKAQNPDQMEKHMAADFDPRASTYESEKYNPSEVGEGEAEEEDEDEEDDDEGG  
RRRRRREAEETTELEREESVAASSMAESQSKANGEEFEVRSEYSDKLPLGAEAVTISQEINGNYRQRPR

**PREDICTED: immunoglobulin superfamily containing leucine-rich repeat protein 2-like [Callorhinchus milii]**

NCBI Reference Sequence: XP\_007906282.1

[GenPept Identical Proteins Graphics](#)

>gi|632938762|ref|XP\_007906282.1| PREDICTED: immunoglobulin superfamily containing leucine-rich repeat protein 2-like [Callorhinchus milii]

MLEKLLCVISGYVFCPWGVRCPEPCVCQDKYFNQFADCAYKNFQAVPVGLPSNVTLSANSKIKSLL  
RADFAEVTVQTSWLAHNEIRKIEKGSLTVLLQLKNLDISHNQIVDFPWEDLYNLTALQLKMNNYMVH  
LSRDAFSTLKERSLRINSNKFTIWEGTFDSSLSSLSHLQIYSNPFSCTCNLQWLKGWIDQALISIPEQK  
DIVCSAPEEFKGTVPVELPDMQCIAPLVHLTYQASNEKGELYEGYALTMHCNATGSPVPVIRWKIQTANK  
EIELNDANVEPERNELLLENRKEVRDRFVVLKNGTLVIPHLTKYEEGAYTCLATNEIGSNRSTLNVAVTA  
SPKREPTYIQERIPSQPGERKPGLKLPKNNAISWAKPGQKGQRISPATARSFPQGQTERNAVFLPPVAKN  
CSKSQGSHYITNHAFNRSSEMVKQHTFDYGIIALEVETETDAKVLQTPFQTAPDKISLEMLYLCAEQGGKAA  
TVVQWMSMIESGVNSYRFQGLNPGSNYTLCLTYTGQDCQVQVFSTR  
RKIPSLLIMIIIVSSFLLGLATIPLVAATCCHLM  
KYRGKTYKLIMKTQPPESLHQNAPCTFPSASFQGSEKIYNPSEVGEESVVAESVPVSQT  
KANPEEFEACSEYSDRLPLGAEAVNISPEINGNYRQPVR

SALM1

**leucine-rich repeat and fibronectin type III domain-containing protein 1 isoform 1 precursor [Mus musculus]**

NCBI Reference Sequence: NP\_001135393.1

[GenPept Identical Proteins Graphics](#)



>gi|213972562|ref|NP\_001135393.1| leucine-rich repeat and fibronectin type III domain-containing protein 1 isoform 1 precursor [Mus musculus]

MAPGPFSGLSPPPAALPFLLLWAGASRGQPCPGRCICQNVAPTLTMLCAKTGLLFVPPAIDRRVVEL  
RLTDNFIAAVRRDFANMTSLVHHTLSRNTIGQVAAGAFADLRALRALHLDNRRAEVRGDQLRGLGNLR  
HLILGNQNQIRKVESAAFDAFLSTVEDLDSYNNLEALPWEAVGQMVLNTLTDHNLIDHIAEGTFVQLH  
KLVRLDMTSNRLHKLPPDGLFLRSQGGPKPPTPLTSFGGNPLHCNCCELLWLRRLTREDDLETATPEH  
LTDRYFWSIPEEEFLCEPPLITRQAGGRALVVEGQAVSLRCRAVGDPPEPVHVWVAPDGRLLGNSRTRVR  
GDGTLDVTITLRSQGTFTCIASNAAGEATAPVEVCVPLPLMAPPPAAPPPLTEPGSSDIATPGRPGAN  
DSTSERRLVAAELTSSSVLIRWPAQRPVPGIRMYQVQYNSSADDLTVYRMIPSTSQTFLVNDLAAGRAYD  
LCVLAVYDDGATALPATRVVGVQFTTAGDPAPCRPLRAHFLG

GTMIIAIGGVIVASVLVFIVLLMI

RYK

VYGDGDSRIKGTSTPPRVSHVCQNTNGAGAQQAQQASAPPAPDRYEALREVAVPAAIEAKAMEAEATSTEL  
EVVLGRSLGGSATSLCLLSEETSGEESRAMTGPRRSRSGALGPPTSAPPTLALVPGGAPARPRPQQRYS  
FDGDYGAFFQSHSYPRARRTKHRSTPHLDGAGGGAAAGEDGDLGLGSARARLAFTSTEWMLESTV

**PREDICTED: leucine-rich repeat and fibronectin type III domain-containing protein 1-like protein isoform X1 [Gallus gallus]**

NCBI Reference Sequence: XP\_423347.4

[GenPept Identical Proteins Graphics](#)

>gi|513240280|ref|XP\_423347.4| PREDICTED: leucine-rich repeat and fibronectin type III domain-containing protein 1-like protein isoform X1 [Gallus gallus]

MMTVCPSPTMDRLVCLVVSAAVKAMLCPKRCMCQNLSPSFTILCTKGLLFVPPSIDRRTAELRLMDN  
FITLRRKDFAINTNLIHLTLSRNTISQIMPYAFFDLKGLHALHLDNRRLTYINEDHFKGHLINLRHLILS  
NNQLSYISPGSLDDFIETIEDLDSYNNLVNVPWETVAKLSNVNTVSLDHNLIEFVPEGIFSNLHKLARL  
DMTSNKLKKIPPPLFSRIPVYAKSKGSPLTSVLVLSFGGNPLHCNCCELVWLRLTREDDLETCASPELM  
GKYFWSIKEEFVCEPPMITHRTPKAVSEGQSVSLCKAVGDPDPYVRWIAPDGKLVNTSRTTSYENG  
TLDIAGTSLGDKGTFTCIASNAAGESTAPVELVVTPYPNLANSTNCEKEAENGPSDILISAKSSFPNETK  
GPQERAVVVGELETSSALIQWPSQQHLPGIRMFQIQYQNSSSDEILVYRMIPAASKSFFLTDLVAGREYDL  
CVLAVYDDGLTSLTATRVIGCVQFTTQEYKQCRLHAQF

LGGTMIIIIGGIIVASVLVFIVLLM

KYKVYNNHHKNKAAKVSNVCSQTNGSHGSMARSTS KLTEGSHQECSASSSKGKAVLDSDGDKVTPTTH  
TTFLT

## TDPLS

### PREDICTED: leucine-rich repeat and fibronectin type III domain-containing protein 1 [Callorhinus milii]

NCBI Reference Sequence: XP\_007907747.1

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>gi|632981721|ref|XP\_007907747.1| PREDICTED: leucine-rich repeat and fibronectin type III domain-containing protein 1 [Callorhinus milii]

MESLLCALVVLGVTVTAQLCPKRCVCQNLSPSIAILCAKTGLLFVPPFIDRRTVELRLTDNFITSVRK  
RDFANMTSLVHHTLSRNTISQIMPHSGFDLRLGLRALHLDNSNRLTKLVD AHLRGVLNLRHLILNNNQLNAI  
SDGSFDDFLGSLEDLDMSYNNLETFPWEAISKMVNLNTSLDHNLIDHIEEGTFSVLHKLSRLDMTSNRL  
HKLPPDPLFLRTQLVNTRGSHFSFLVLSFGGNPLHCNCCELLWLRRLMREDDLETASPPHLMGKYFWSI  
AEEEFICEPPLITRLQATKTFVMEGQGVTLKCKAVGDPDPSILWSLPEGKLVSNSTSRTIYDNGTLDILI  
TTLKDNGRFACIASNAAGESATNITIGIPLPHFVNLTQHIKVDPGSSDISTSSKGAPSNSSDTKSTQ  
DKKVTASELTTTSALVRWPSQRSIPGIRMYQIQYNSSSDNTLVYRMIPSTSQLFLVNDLAPGRDYELCVL  
AVYDDGMTTLTATRAVGCVRFTTEQEYTQCHSVHTQF

LGGTMIIIGGIIVASVLVFIIILMI

RYKVYSS

GLGDSKAVGTVNVYSQTNGNGSHNGALDRSCSKPEGPGESVPEALVELPDQSQTVVLSVMCEKAGGAHTT  
A  
SATASASASVTVPTEGALPQAQRRRVQPGATGQHQHQQQLEPQTSEEQHTEASTTDSSMSVCLISSRG  
TLPGRGKPAKLSNISLLPREISRTQHRHSFDGDYSLFQSHSYPRRARTKRSLTGSGQQLHCEDRRGTFSS  
TEWMLESTV

## SALM2

### leucine-rich repeat and fibronectin type-III domain-containing protein 2 precursor [Mus musculus]

NCBI Reference Sequence: NP\_081728.2

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protein	1	fasta
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>gi|226246673|ref|NP\_081728.2| leucine-rich repeat and fibronectin type-III domain-containing protein 2 precursor [Mus musculus]

METLLGGLLAFGMAFAVVDACPKYCVCQNLSESLGTLCPSKGLFVPPDIDRRTVELRLGGNFIIHIGRQ  
DFANMTGLVDLTLSRNTISHIQPFSFLDLESRLSHLDNSNRLPSLGEDTLRGLVNLQHLIVNNNQLGGIA  
DDAFEDFLLTLEDLDSYNNLHGLPWDSVRRMVNLHQLSLDHNLLDHIAEGTFADLQKLARLDLTSNRLQ

KLPPDPIFARSQASLLATPFAPPLSFSFGGNPLHCNCELLWLRLERDDDLETGSPGSLKGRYFWHIR  
EEEFVCEPPLITQHTHKLLVLEGQAATLKCKAIGDPSPLIHVVAPDDRLVGNSSRTAVYDNGTLDILITT  
SQDSGPFTCIAANAAGEATATVEVSIVQLPHLSNSTSRMAPKSRLSDITGSSKTSRGGGSGAGEPPKS  
TPERAFLVSDVTTTSALVKWSVSKSAPRVKMYQLQYNCSDDEVLIYRMIPASNKAFFVNNLVSGTGYDLC  
VLAMWDDTATTATNIVGCAQFFTAKADYPQCQSMHSQI

LGGTMILVIGGIIVATLLFIVILMV

RYKVC

NHDTPGKMAAATSVNVSQNTNGSQPPPLGGIPVGQLPQAPPKVVRNELMDFSTSLARACDSSSSSSLGS  
GEAAGLGRGPWRLPPPAPRKPSLDRLMGAFASLDLKSQRKEELDSRTPAAGRAGTSSRGHHSDREPLL  
GPPATRARSLLPLPLEGKAKRSHSFDMGDFAAAAAVPGGYSPPRRVSNIWTKRSLSVNGMLLPFEESDL  
VGARGTGFSEWVMESTV

#### PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 2 [Gallus gallus]

NCBI Reference Sequence: XP\_004935432.1

[GenPept Identical Proteins Graphics](#)

>gi|513175233|ref|XP\_004935432.1| PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 2 [Gallus gallus]

MEKLLCGILVFGMAVMVNACPKYCVQCNLSESLGTLCPKGLLFVPLIDRRTVELRLGGNFIINISRQD  
FANMSGLVDLTLSRNTISYIQPYSFDTLESRLSHLDNSRLPDIGEDILRGLINLQHLILNNNQLTSISD  
EAFEDFLLTLEDLDSYNNLR SIPWESIRKMINLHQSLDHNLIDYITEGTFADLQKLARLDDTSNRLQK  
LPPDPIFARSQVIPLAVTPFSPPLSLSFGGNPLHCNCELLWLRRRLDRDDMETCASPPGLKGRYFWYVRE  
EEFVCEPPLITQHTHKLLVLEGQTATLKCKAIGDPTPIIHVVAPDDRLIGNSSRTSVYDNGTLDILITTS  
KDYGFTFCIAANAAGESTATIELSIVQLPHLSNGTGRAAPPKSRLSDITSSSKSNRGETKGPPERAVLVS  
EVTTTSALVKWTVSKSAPRVKMYQLQYNCSDDEVLIYRMIPATNKAFVVNNLVSGTGYDLCVLAMWDDT  
A

TTLTATNIVGCAQFFTAKADYPQCQSMHSFLGGT

MILIIGGIIVATLLFIVILMV

RYKVCNNSQGKMSS

VSNVVSQNTNGAQPVQNGVLPQVNPKVVVRNELMEFNSGSVRSSISSSSSMNSRDCDNYSSEQGTLS  
KWRPPSRSKHNIDRLMGAFASLELKCKKEETDSRTSTAARHSDKEPLLGQPESKFRSLLMLPLEGKTK  
RSHSFMDGMDFATSQCCTYPKKITNIWTKRSLSVNGMLLQYDDNDLTGAKGTYGSSEWVMESTV

#### PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 2 [Callorhinchus milii]

NCBI Reference Sequence: XP\_007908260.1

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>gi|632982664|ref|XP\_007908260.1| PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 2 [Callorhinus milii]

MEKLLCNLLVIGMAVTYACPKYCVCQNLSESLGTLCPSKGLFVPPNIDRRTVELRLGGNFILSINRQD  
FGNMTGLVDLTLRSNTIDYIQPYSFADLESLRSLHLDNSNRLTRIGSNDRGLLNQHLILNNNQLNSILD  
EAFDDFLTLEDLDSYNNLVSLPWEALGKMINLHTLSLDHNLIDYIPEGTFTDLLKLARLDLVSNRLQK  
LPPDPFARSETFVLSTTPYFAPLSLSIGGNPLHCNCCELLWLRLRSREDDMETCASPSSLKGRYFWYVPE  
EEFVCEQPLITQHSHKVLVLEGQTATLRCKAIGDPKPVIHWVAPDDRILGNSSRTVIYDNGTLDILITTS  
KDYGTFCTIAANAAGESTASIELSIVQLPHLSNGTGRAVQPGSRLSDITSSSKTYRGETMSKPEKVVKVV  
DVTASTALVKWSVGRSAPKVEMYQFQYNSSTDDEVLYVRMIPASNKAFFVKNLVPSSNYDLCVLAIWDDT  
L  
TTLTATNVVGCVRFTTSEDYTQCKSFHSQ  
FLGGTMILIIGGIIVASLLVFIILTI  
KYKLCNGQEKLPDV  
NNVCSQTNGGQPVLNGILPQLNPKVVRDEMELFCNCISIHSSMSSSTGSSQDCEDCYSLNSNASTLSKKW  
RHRSKSRHNIDRLMGAFASLDLRCQRKEDNCESRASTLAHYSDKEPLLGHSESRLNKLLTPMEVTKRS  
HSFDMSDFATTPCYNPERRITNIWTRRSLSVNGTLLQYDEEDLESTKGMYCSSEWMESTV

### SALM3

leucine-rich repeat and fibronectin type-III domain-containing protein 3 precursor [Mus musculus]

NCBI Reference Sequence: NP\_780687.1

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>gi|30425224|ref|NP\_780687.1| leucine-rich repeat and fibronectin type-III domain-containing protein 3 precursor [Mus musculus]

MAVLPLLLCLLPLAPASSPPQPAISSPCPRRCRCQTQSMPLSVLCPGAGLLFVPPSLDRRAELRLADNF  
IAAVRRRDLANMTGLLHLSLSRNTIRHVAAGAFADLRALRALHLDGNRLTSLGEGQLRGLVNLRHLILSN  
NQLAALAAGALDDCAETLELDLSYNNEQLPWEALGRLGNVNTLGLDHNLASVPAGAFSRLHKLARL  
D  
MTSNRLTTIPPDPFLSR  
LPPLLARPRGSPASALVLAGGNPLHCNCLEVWLRLAREDDLEACASPPALGG  
RYFWAVGEEFVCEPPVTHRSPPLAVPAGRPAALRCRAVGDPEPRVRWVSPQGRLLGNSSRARAEPNGT  
LELLVTEPEDGGTFTCIAANAAGEATAAAVELTVGPPPPPQLANSTSCDPPRDGEPEPDALTTPSAASASAKV  
ADTVAPTDGVQVTEHGATAALVQWPDQRPVPGIRMYQIYQYNSADDILVYRMIPADSRSFLLTDLASGR  
TYDLCVLAVYEDSATGLTATRPGCARFSTEPALRPCAAPHAPF  
LGGTMIIALGGVIVASVLVFIFVLL  
RYKVHGGQPPGKAKATAPVSSVCSQTNGALGPVPSAPAPEPAAPRAHTVVQLDCEPWGPSHEPAGP

## SALM4

### leucine-rich repeat and fibronectin type-III domain-containing protein 4 precursor [Mus musculus]

NCBI Reference Sequence: NP\_700437.2

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protein	1	fasta
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>gi|31559904|ref|NP\_700437.2| leucine-rich repeat and fibronectin type-III domain-containing protein 4 precursor [Mus musculus]

MAPPLLLLLLASGAAACPLPCVCQNLSESLSTLCAHRGLLFVPPNVDRTVELRLADNFIQALGPPDFRN  
MTGLVDLTLSRNAITRIGARSFGDLESRLSLHLDGNRLVELGSSSLRGPVNLQHLILSGNQLGRIAPGAF  
DDFLDSLELDLVSYNNLRQVPWAGIGSMPALHTLNLDHNLI DALPPGVFAQLSQLSRLDTSNR LATLAP  
DPLFSRGRDAEASPSPVLSFSGNPLHCNCELLWLRRRALPDDLETASPTLAGRYFWAVPEGEFSCEP  
PLIARHTQRLWVLEGQRATLRCRALGDPVPTMHWVGPDRLVGNSSRAWFPNGTLEIGVTGAGDAGAY  
T  
CIATNPAGEATARVELRVLALPHGGNTSAEGGRPGPSDIAASARTAAEGETLESEPAVQVTEVTATSGL  
VSWGLGRPADVWMFQIYQYNSEDETLYRIVPASSHHFLKHLVPGADYDLCLLALSPAAGPSDLTATR  
LLGCAHFSTLPATPLCHALQAHVLG  
GTLTVAVGGVLVAALLVFTVALLV  
RGRGAGNGRLPLKLSHVQSQT  
NGGTSPMPKSHPRSPPPRPQRSCSDLGDTGGCYGYARRLGGAWARRSHSVHGGLGAGCRGVGGS  
LEESVV

### PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 4-like [Callorhinchus milii]

NCBI Reference Sequence: XP\_007909247.1

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>gi|632984655|ref|XP\_007909247.1| PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 4-like [Callorhinchus milii]

MEKFTFAFLVGSLAAGSEACPFHCTCQNLSESLSTLCANKGLFIPINIDRRTVELRLADNFLRVIAQP

DFLNMSGLVDLTLSRNTIISLEPFAFGDLESRLSHLDNRRLIRIHEDSLRGLINLQHLIINNNQLINIA  
LSAFDDFVVTLEDLDSFNLLQRVPWEAIQSMVNHLMLNLDHNLIDYIMADTFAELFKLARLDMTSNRLQ  
TLPPDSLFSRSQTGVINPTPYTSIIILNFGGNPLHCNCCELLWLRLVREDDMETCASPAGRYFWSIP  
EEEFICEPPLITRHTKVVILEGQRATLKCRAGDPEPIHWVSPEDKIVNSSRIVSYRNGTLDILVTT  
MREDGVYTCFATNAAGESTALADLKIPLPHRGNGTLQILHHDPGSSDISTSTKPVTNSTGRSRPRDKTV  
SVTDVTGTTALIRWAQSKSPHIVWMYQIQYNCSIDETLVYRIISSSKAFILKNLISGVDYDLCILAIYD  
DSVTQLAATKVVGCIQFSTHEEYPHCHLLHAHF  
LGGTLTVIVGGIIVVTLLVFTVIMMV  
KYKVCGSARCE  
VPKLTDVYSQTNGSQTTPNGMVAQRITVLNTRGQPTGGVPVPDLSSANLPRQESRKAPPYSAKTQRKR  
YKCKQRGEGDGELATLGCQGGEGPGERTALAKQPCPQSSE

## SALM5

### **leucine-rich repeat and fibronectin type-III domain-containing protein 5 isoform 1 precursor [Mus musculus]**

NCBI Reference Sequence: NP\_848829.2

#### GenPept Identical Proteins Graphics

>gi|31559842|ref|NP\_848829.2| leucine-rich repeat and fibronectin type-III domain-containing protein 5 isoform 1 precursor [Mus musculus]

MEKFLFYFLIGIAVRAQICPKRCVCQILSPNLATLCAKKGLFVPPNIDRRRTVELRLADNFVTNIKRKD  
FANMTSLVDLTLRSNTISFITPHAFADLRNLRALHNSNRLTKITNDMSGLSNLHHLILNNNQLTLISS  
TAFDDVFALEELDSYNNELETIPWDAVEKMVSLHTLSLDHNMDNIPKGTFSHLHKMTRLVTSNKLQKL  
PPDPLFQRAQVLATSGIISPSTFALSFGGNPLHCNCCELLWLRLRSREDDLETCASALLTGRYFWSIPEE  
EFLCEPPLITRHTHEMRVLEGQRATLRCKARGDPEPAIHVISPEGKLISNATRSLVYDNGTLDILITTVK  
DTGAFTCIASNPAGEATQTVDLHIIKLPHLLNSTNHIHEPDPGSSDISTSTKSGSNASSNGDTKMSQDK  
IVVAEATSSTALLKFNFQRNIPGIRMFIQYNGTYDDTLVYRMIPPTSFLVNNLASGTMYDLCVLAIFY  
DDGITSLTATRVVGCIQFTTEQDYVRCHFMQSQFL

GGTMIIIIGGIIVASVLVFIILMI

RYKVCNNNGQ

HKVTKVSNVYSQTNGAQMQGCSVLPQSMSKQAMGHEENAQCCKVASDNAIQSSETCSSQDSSTTSALP  
PTWTSSAPVSQKQKRKTGTPSAEPQSEAVTNVESQNTNRRNSTALQLASCPPDSVTEGPTSQRATHKPS  
KFLTVPAEGSRARHRASLSGGLKDSFHYGNSQLKRSMSMNAMWT

### **PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 5 [Gallus gallus]**

NCBI Reference Sequence: XP\_421485.2

[GenPept Identical Proteins Graphics](#)

>gi|118092246|ref|XP\_421485.2| PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 5 [Gallus gallus]

MEKLLFLLFIGIAVRAQICPKRCVCQILSPNLATLCAKKGLLFVPPNIDRRTVELRLADNFVTNIKRKD  
FANMTSLVDLTLRSNTISFITPHAFADLRNLRALHLSNRALKITNDMFSGLSNLHHLILNNNQLTLISS  
TAFDDVLAEEELDSYNNLETIPWDAVEKMVLHLSLDHNMDHIPKGTFSHLHKMTRLDVTSNKLQKL  
PPDPLFQRAQVLATSGIISPSTFALSFGGNPLHCNCCELLWLRRRLSREDDLETCASPLLSGRYFWSIPEE  
EFLCEPPLITRHTHELRVLEGQRAALRCKARGDPEPAIHVISPEGKLISNATRSVVDNGTLDILITTVK  
DTGSFTCIASNPAGEATQTVDLHIIKLPHLLNSTNHIHEPDPGSSDISTSTKSGSNASSNGDTKVSQDK  
KVVAEAETSSSTALLKFNFQRNIPGIRMFQIQYNGTYDDSLVYRMIPPTSKTFLVNNLAAGTMYDLCVLA  
YDDGITSLTATRVVGCTQFTTEQDYVRCHFMQSQFL

GGTMIIIGGIIVASVLVFIILMI

RYKVCNNNG

QHKATKVSNVYSQTNGAQVQACGGALSQSASKQAVGHEEAAQCCRAASDGAGPSPEPSPGPEATAATTTS  
PSPHAWAAGTSAAQPKRKPGPKPSSEPQSEAAMSIESQNTNRNNSTALQLASRPPDSDKGVPTYKRAQS  
KPKAGADLKDTHTAPLESCPNLATRQTKRSQRTKD

#### PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 5 [Callorhinchus milii]

NCBI Reference Sequence: XP\_007891516.1

[GenPept Identical Proteins Graphics](#)

>gi|632951833|ref|XP\_007891516.1| PREDICTED: leucine-rich repeat and fibronectin type-III domain-containing protein 5 [Callorhinchus milii]

MEKLLFYLLIGMAVKAQVCPKRCVCQNLSPNLATLCAKKGLLFVPPNIDRRTVELRLGDNFITSIKRKD  
FANMTGLVDLTLRSNTINHIAPQAQFSDLCLNRLALHLSNRLTQITNEMFSRLSKLHHLIVNNNQLIEISS  
GAFSDILLSLEELDVSYNNLKTIPWEAVEKMVNHLTSLDHNMLEHIDEGTFSHLHKLIRLDMTSNKLRK  
LPPDPLFTRVQLANLGIMNPTGFVLSFGGNPLHCNCCELLWLRRRLSREDDLETCASPTHLTGRYFWSIPE  
EEFICDQPLITRHTHELRVLEGQRATLKCKAIGDPDPSIHWSSPEGKLISNMSRTVLYANGTLDILITTV  
KDTGTFTCIASNAAGETTAMVELHIIKLPHLINSTNHIHEPDPGSSDISTSTKSGSNTSNSVSDTKVKPE  
RRVAVAETSSSALIKFNLQHNIPGIRMFQIQYNGSYDDSLVYRMIPSTSFTLVNLAAGTLYDLCVLA  
IYDDGITSLTATRVVGCFETTDQDYVRCHFMPSQFL

GGTMIIIGGIIVASVLVFIILMI

RYKVCNNN

DQHKMTKVSNVYSQTNGAHLQMCGSVLSHSNSKVAMGHDDNITRCNKDPSESKTQLSESTLSQDCSTTT  
TLPHDWTASVSPSQKLKRKAGLNPSVESPMEAFTNVESLKKENTAILQKSTCAQISLKDTPTFRAHSK  
SIKFLTLPTIESRAKRRYSLDAEVSEYHCYTHSQSINSLWSKRSMMSNGMLQLANSVDGGKAVFSSSE

WIMESTV

## Appendix C: IC domain

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LINX

>HsLinx

CHLLAKHPGKPYRLILRPQAPDPMEKRIAADFPRASYLESEKSYPAGGEAGGEEPEDVQGEGLDEDAEQG  
DPSGDLQREESLAACSLVESQSKANQEEFEAGSEYSDRLPLGAEAVNIAQEINGNYRQTAG

>MmLinx

CHLLAKHPGKPYRLILRPQAPDPMEKRIAADFPRASYLESEK  
SYPARGEAGGEEPEEVPEEGLDEDVEQGDPSGDLQREESLAGCSLVESQSKANQEEFEAGSEYSDRLPLG  
AEAVNIAQEINGNYRQTAG

>GgLinx

SKYQGKTYKLIMKAQNPDQMEKHMAADFPRASYLESEKNYNPSEVGEGEAEEEDEDEDDDEGG  
RRRRRREAEETTELEREESVAASSMAESQSKANGEEFEVRSEYSDKLPLGAEAVTISQEINGNYRQRPR

>CmLinx

KYRGKTYKLIMKTQPPESLHQNAPCTFDPSASFQGSEKIYNPSEVGEESVVAESVPVSQT  
KANPEEFEACSEYSDRLPLGAEAVNISPEINGNYRQPVR

SALM1

>HsSALM1

RYKVCNHEAPSKMAAAAVNVYSQTNGAQPPPPSSAPAGAPPQGPPKVVVRNELDFTASLARASDSSSSSS  
LGSGEAAGLGRAPWRIPPSAPRPKPSLDRLMGAFASLDLKSQRKEELDSRTPAGRGAAGTSARGHHSDREP  
LLGPPAARARSLLPLEGKAKRSHSFDMGDFAAAAAGGVVPGGYSPRKVSNIWTKRSLSVNGMLPFEE  
SDLVGARGTFGSSEWVMESTV

>MmSALM1

RYKVC

NHDTPGKMAAATSVNVYSQTNGSQPPPLGGIPVQLPQAPPKVVVRNELMDFSTSLARACDSSSSSSLGS  
GEAAGLGRGPWRLPPPAPRPKPSLDRLMGAFASLDLKSQRKEELDSRTPAGRGAAGTSSRGHHSDREPLL

GPPATRARSLLPLPLEGKAKRSHSFDMGDFAAAAAVPGGYSPRRVSNIWTKRSLSVNGMLLPFEESDL  
VGARGTGSSEWVMESTV

>GgSALM1

KYKVYNNHHKNKAAKVSNVCSQTNGSHGSMARSTSKLTEGSHQECSASSSKGAVLDSGDKVTPTTH  
TTFLT  
TDPLS

>CmSALM1

RYKVYSS  
GLGDSKAVGTVNYSQTNGNGSHNGALDRSCSKPEGPGESVPEALVELPDQSQTVVLSVMCEKAGGAHTT  
A  
SATASASASVTVPTEGALPQAQRRRVQPGATGQHQHQQQLEPQTSEEQHTEASTTDSSMSVCLISSRG  
TLPGRGKPAKLSNISLLPREISRTQHRHSFDGDYSLFQSHSYPRRARTKRSLTGSGQQLHCEDRRGTFS  
TEWMLESTV

SALM2

>HsSALM2

RYKVYGDGDSRRVKGSRSLPRVSHVCQTNGAGTGAAQAPALPAQDHYEALREVESAAPAVAVEAKA  
MEAETASAEPVVLRSLGGSATSLCLPSEETSGEESRAAVGPRRSRSGALEPPTSAPPTLALVPGGAAAR  
PRPQQRYSDGDYGALFQSHSYPRRARRTKRHRSTPHLDGAGGGAAAGEDGDLGLGSARACLAFTSTEWM  
LESTV

>MmSALM2

RYK  
VYGDGDSRIKGTSRTPRVSHVCQTNGAGAQASAPPADRYEALREVAVPAAIEAKAMEAEATSTEL  
EVVLGRSLGGSATSLCLPSEETSGEESRAMTGPRRSRSGALGPPTSAPPTLALVPGGAPARPRPQQRYS  
FDGDYGAFLQSHSYPRRARRTKRHRSTPHLDGAGGGAAAGEDGDLGLGSARAARLAFTSTEWMLESTV

>GgSALM2

RYKVCNNSQGMSS  
VSNVYSQTNGAQPVQNGVLPQVNPKVVVRNELMEFNSGSVRSSISSSSSMNSRDCDNYSLQSEQTLSS  
KWRPPSRSKHNIDRLMGAFASLELKCKKEETDSRTSTAARHSDKEPLLGPESKFRSLLMLPLEGKT  
RSHSFDMGDFATSQCCTYPKKITNIWTKRSLSVNGMLLQYDDNDLTGAKGTYGSSEWVMESTV

>CmSALM2  
KYKLCNGQEKL PDV  
NNVCSQTNGGQPVLNGILPQLNPKVVGRDEMLEFNCGSIHSSMSSSTGSSQDCE DCYSLNSNASTLSKKW  
RHR SKSRHNIDRLMGAFASDLRCQRKEDNCESRASTLAHYSDKEPLLGHSESRLNKLTLPMEVTKRS  
HSFDMSDFATTPCYNP RITNIWTRRSLSVNGTLLQYDEEDELESTKG MYCSSEWVMESTV

SALM3  
>HsSALM3  
RGRGAGNGLPLKLSHVQSQTNGGPSPTPKAHPPRSPPPRPQRSCSDLGDAGCYGYARRLGGAWARRSH  
SVHGLLGAGCRGVGGSAERLEESVV

>MmSALM3  
RGRGAGNGLPLKLSHVQSQT  
NGGTSPMPKSHPPRSPPPRPQRSCSDLGDTGGCYGYARRLGGAWARRSHSVHGLLGAGCRGVGGSAER  
LEESVV

>GgSALM3

>CmSALM3

SALM4

>GgSALM4

>HsSALM4  
MRYKVHGGQPPGKAKIPAPVSSVCSQTNGALGPTPTPAPP APEPA ALRAHTVVQLDCEPW GPGHEPVGP

>MmSALM4  
RYKVHGGQPPGKAKATAPVSSVCSQTNGALGPVPSA PAPEPA APRAHTVVQLDCEPW GPSHEPAGP

>CmSALM4  
KYK VCGSARCE  
VPK LTDV YSQ TNGS QTTV PNM VSA QR ITVL NTRG QPTGGV PVPD LSSANL PRQ ESRK APPY SAKT QRKR  
YKCKQRGE GDGE LA LG CQG GE GP GERT ALAK QPCP QSSE

SALM5

>HsSALM5

RYKVCNNNGQHKVTKVSNVYSQTNGAQIQGCSVLPQSFSKQAVGHEENAQCCKATSDNVIQSSETCSSQ  
DSSTTSALPPSWTSSTSFSQKQKRKTGKSTEPQNEAVTNVESQNTNRNNSTALQLASRPPDSVTEGPTS  
KRAHIKPNALLTNVDQIVQETQRLELI

>MmSALM5

RYKVCNNNGQ  
HKVTKVSNVYSQTNGAQMQGCSVLPQSFSKQAMGHEENAQCCKVASDNAIQSSETCSSQDSSTTSALP  
PTWTSSAPVSQKQKRKTGKPSAEPQSEAVTNVESQNTNRNNSTALQLASCPPDSVTEGPTSQRAHTKPS  
KFLTVPAEGSRARHRASLSGGLKDSFHYGNSQLSKRSMSMNAMWT

>GgSALM5

RYKVCNNNG  
QHKATKVSNVYSQTNGAQVQACGGALSQSASKQAVGHEEAAQCCRAASDGAGPSPEPSPGPEATAATTTS  
PSPHAWAAGTSAAQKPKRKPGPKPSSEPQSEAAMSIESQNTNRNNSTALQLASRPPSDKGVPTYKRAQS  
KPKAGADLKDTHTAPLLESSCPNLATRQTKRSQRTKD

>CmSALM5

RYKVCNN  
DQHKMTKVSNVYSQTNGAHLQMCGSVLSHSNSKVAMGHDDNITRCNKDPSESKTQLSESTLSQDCSTTT  
TLPHDWTASVSPSQKLKRKAGLNPSVESPMEAFTNVESLKKENTAILQKSTCAQISLKDTPTFRRAHSK  
SIKFLTPTEISRAKRRYSLDAEVSEYHCYTHSQSINSLSKRSMSMNGMLLQLANSVDGGKAVFSSSE  
WIMESTV

## Appendix D: Boxshade Outputs of LIGS

Boxshade for 1.0 match

HsLinx	1 CHLLAKHPGKPYRLILRPQAPDPMEKRIAADFDFRASYLESEKSYPAGGEAGGEEPEDVQ
MmLinx	1 CHLLAKHPGKPYRLILRPQAPDPMEKRIAADFDFRASYLESEKSYPARAGEAGGEEPEEV
GgLinx	1 ----SKYQGKTYKLIMKAQNPDQMEKHMAADFDFRASYLESEKNYNPSEVGEAEEEEDE
CmLinx	1 ----YKYRGKTYKLIMKTQPPESLHQNAPCTFDFASASFQGSEKIYNPSEVCEES-----
consensus	1 K GK YrLIlr Q Pd m FDP ASy SEK Y a

HsLinx	61	-----GEGLDEDQEQGDPSGDLQREES <b>L</b> AACSLVE <b>S</b> QS <b>K</b> AN <b>Q</b> EE <b>F</b> EAG <b>S</b> EYSD <b>R</b> LPLGA
MmLinx	61	-----EEGLDEDVEQGDPSGDLQREES <b>L</b> AGCSLVE <b>S</b> QS <b>K</b> AN <b>Q</b> EE <b>F</b> EAG <b>S</b> EYSD <b>R</b> LPLGA
GgLinx	57	DEEDDDEGGRRRRRREAETTELEREES <b>V</b> AASS <b>M</b> AE <b>S</b> QS <b>K</b> ANG <b>E</b> EF <b>V</b> R <b>S</b> EYSD <b>K</b> LPLGA
CmLinx	51	----- <b>V</b> WAESV <b>P</b> V <b>S</b> QT <b>K</b> AN <b>P</b> EE <b>F</b> EAC <b>S</b> EYSD <b>R</b> LPLGA
consensus	61	<b>L</b> REES <b>I</b> a <b>S</b> <b>I</b> <b>S</b> Q <b>s</b> K <b>AN</b> <b>Q</b> EE <b>F</b> E <b>S</b> EYSD <b>R</b> <b>L</b> PLGA

HsLinx	115	<b>E</b> AVNI <b>A</b> QEINGNYRQTAG
MmLinx	115	<b>E</b> AVNI <b>A</b> QEINGNYRQTAG
GgLinx	117	<b>E</b> AVTI <b>S</b> QEINGNYRQPRP
CmLinx	83	<b>E</b> AVNI <b>S</b> PEINGNYRQPVR
consensus	121	<b>E</b> AVNI <b>QEINGNYRQ</b>

## SALM1

0.7 match

CmSALM1	1	<b>R</b> YKV <b>Y</b> SSGLGD--SK <b>A</b> VG <b>T</b> NVYSQTNG <b>N</b> GS <b>H</b> NG <b>A</b> LDRSCSK <b>P</b> E <b>G</b> PGESV-PEAL <b>V</b> LPDQ
GgSALM1	1	<b>K</b> YKV <b>Y</b> NNHHK--NK <b>A</b> AK <b>V</b> SNVC <b>S</b> QTNG <b>S</b> HGGSM <b>A</b> RSTS----- <b>K</b> -----L <b>T</b> E-
HsSALM1	1	<b>R</b> YKV <b>C</b> NHEAPSKMAA-AVS <b>N</b> VYSQTNG <b>A</b> QPPPPSSAPAGAP <b>P</b> Q <b>G</b> PP <b>K</b> V <b>V</b> VRNEL <b>L</b> D <b>F</b> T <b>A</b> -
MmSALM1	1	<b>R</b> YKV <b>C</b> NHDT <b>P</b> KGMAAAT <b>V</b> SNVYSQTNG <b>S</b> QPPPL <b>C</b> GPVGQL <b>P</b> QAPP <b>K</b> V <b>V</b> VRNEL <b>L</b> MD <b>F</b> ST-
consensus	1	<b>r</b> YKV <b>n</b> <b>aa</b> vs <b>N</b> Vy <b>S</b> QTNG <b>a</b> <b>p</b> <b>gp</b> <b>k</b> <b>v</b> <b>lve</b> <b>t</b>

CmSALM1	58	SQTVV <b>L</b> SVM <b>C</b> E <b>K</b> AGGAHTTA-S <b>A</b> T <b>A</b> S <b>A</b> S <b>A</b> SVT <b>V</b> P <b>T</b> E <b>G</b> AL <b>P</b> QA <b>Q</b> RR <b>R</b> V <b>Q</b> P <b>C</b> AT <b>G</b> Q---HQ
GgSALM1	41	----GSHQE <b>C</b> S <b>A</b> SSSK-- <b>G</b> <b>K</b> <b>A</b> V--- <b>L</b> DS <b>D</b> G <b>D</b> K <b>V</b> T <b>P</b> T <b>T</b> H <b>T</b> T <b>F</b> L <b>T</b> D <b>P</b> LS-----
HsSALM1	59	---- <b>S</b> <b>L</b> ARAS <b>D</b> <b>S</b> <b>S</b> <b>S</b> <b>S</b> <b>I</b> <b>L</b> <b>G</b> <b>S</b> <b>E</b> <b>A</b> <b>A</b> <b>G</b> <b>L</b> <b>C</b> <b>R</b> <b>A</b> <b>P</b> <b>R</b> <b>F</b> <b>K</b> <b>P</b> <b>S</b> <b>L</b> <b>D</b> <b>R</b> <b>L</b> <b>M</b> <b>G</b> <b>A</b> <b>F</b> <b>A</b> <b>S</b> <b>L</b> <b>D</b> <b>L</b> <b>K</b> <b>S</b> <b>Q</b> <b>R</b>
MmSALM1	60	---- <b>S</b> <b>L</b> ARAC <b>D</b> <b>S</b> <b>S</b> <b>S</b> <b>S</b> <b>I</b> <b>L</b> <b>G</b> <b>S</b> <b>E</b> <b>A</b> <b>A</b> <b>G</b> <b>L</b> <b>C</b> <b>R</b> <b>G</b> <b>P</b> <b>W</b> <b>R</b> <b>I</b> <b>P</b> <b>P</b> <b>A</b> <b>P</b> <b>R</b> <b>E</b> <b>K</b> <b>P</b> <b>S</b> <b>L</b> <b>D</b> <b>R</b> <b>L</b> <b>M</b> <b>G</b> <b>A</b> <b>F</b> <b>A</b> <b>S</b> <b>L</b> <b>D</b> <b>L</b> <b>K</b> <b>S</b> <b>Q</b> <b>R</b>
consensus	61	l ce sss g a a ala a kvpp g p t drl g

CmSALM1	113	<b>H</b> QQ <b>Q</b> <b>L</b> <b>E</b> <b>P</b> <b>O</b> <b>T</b> <b>S</b> <b>S</b> <b>E</b> <b>G</b> <b>H</b> <b>T</b> <b>E</b> <b>A</b> <b>S</b> <b>T</b> <b>T</b> <b>D</b> <b>S</b> <b>S</b> <b>M</b> <b>V</b> <b>C</b> <b>L</b> <b>I</b> <b>S</b> <b>S</b> <b>R</b> <b>G</b> <b>T</b> <b>L</b> <b>P</b> <b>G</b> <b>R</b> <b>G</b> <b>K</b> <b>PA</b> <b>K</b> <b>L</b> <b>S</b> <b>N</b> <b>I</b> <b>S</b> <b>L</b> <b>L</b> <b>P</b> <b>R</b> <b>E</b> <b>I</b> <b>S</b> <b>-</b> <b>R</b> <b>T</b>
GgSALM1		-----
HsSALM1	115	<b>K</b> EE <b>I</b> <b>L</b> <b>D</b> <b>S</b> <b>R</b> <b>T</b> <b>P</b> <b>A</b> <b>G</b> <b>R</b> <b>G</b> <b>A</b> <b>G</b> <b>T</b> <b>S</b> <b>A</b> <b>R</b> <b>G</b> ----- <b>H</b> <b>S</b> <b>D</b> <b>R</b> <b>E</b> -- <b>P</b> <b>L</b> <b>I</b> <b>G</b> <b>P</b> <b>P</b> <b>A</b> <b>A</b> <b>-</b> <b>R</b> <b>A</b> <b>R</b> <b>S</b> <b>L</b> <b>L</b> <b>P</b> <b>L</b> <b>I</b> <b>E</b> <b>G</b> <b>K</b> <b>A</b>
MmSALM1	116	<b>K</b> EE <b>I</b> <b>L</b> <b>D</b> <b>S</b> <b>R</b> <b>T</b> <b>P</b> <b>A</b> <b>G</b> <b>R</b> <b>G</b> <b>A</b> <b>G</b> <b>T</b> <b>S</b> <b>R</b> <b>G</b> ----- <b>H</b> <b>S</b> <b>D</b> <b>R</b> <b>E</b> -- <b>P</b> <b>L</b> <b>I</b> <b>G</b> <b>P</b> <b>P</b> <b>A</b> <b>T</b> <b>-</b> <b>R</b> <b>A</b> <b>R</b> <b>S</b> <b>L</b> <b>L</b> <b>P</b> <b>L</b> <b>I</b> <b>E</b> <b>G</b> <b>K</b> <b>A</b>

consensus 121 h le t g s r p g pa sllp i r

CmSALM1 172 QHRHSFD-GDYSLFQ-----SHSYPR---RAR[TKRSLS]GS[GQQL]----HCEDRGTF  
GgSALM1 -----  
HsSALM1 164 KRSHSFDMGDFAAAAAGGVPGGYSPPRKVSNIWTKRSLSVNGMLLPFEESDLVGARGTF  
MmSALM1 165 KRSHSFDMGDFAAAA--AAVPGGYSPPRRVSNIWTKRSLSVNGMLLPFEESDLVGARGTF  
consensus 181 h **hsfd** gdy s pr tkrslt g l rgtf

CmSALM1 216 S**STEW**M**LE**STV  
GgSALM1 -----  
HsSALM1 224 GS**SEW**V**M**E**ST**V  
MmSALM1 223 GS**SEW**V**M**E**ST**V  
consensus 241 **stewmlestv**

**SALM2**

HsSALM2	1	<b>RYK</b> VYGDGDSRRVKGS-RSLPRVSHVC <b>SQTNGA</b> GTGAAQAPAL <b>PAQDHYEALREVESQAA</b>
MmSALM2	1	<b>RYK</b> VYGDGDSRRIKGTSRTPPRVSHVC <b>SQTNGA</b> GAQQA--SAP <b>PAPDRYEALREVAVP--</b>
GgSALM2	1	<b>RYK</b> V-----NNSQGKMSS <b>VSNVY</b> SQTNGA <b>QPVQNG--VL</b> PQV-----N
CmSALM2	1	<b>KYKL</b> C-----NG-QEKLPDVNNV <b>C</b> SQTNG <b>QPVNLNG--IL</b> PQL-----N
consensus	1	<b>rYKv</b> VS V SQTNG <b>a</b> LP

HsSALM2	60	PAVAVEAK <b>A</b> ETA <b>SAEPE</b> -----VV <b>L</b>
MmSALM2	57	--AAIEAK <b>A</b> ME <b>AEAT</b> STELE-----VV <b>L</b>
GgSALM2	38	PKVVVRNEL <b>M</b> E <b>F</b> NNSG <b>V</b> RSSISSSSSMNSRDC-DNYSLQSEQGTLSSKWRPPSRSKHN <b>I</b>
CmSALM2	37	PKVVG <b>RDEM</b> <b>L</b> E <b>F</b> NCG <b>S</b> I <b>H</b> SSMSSSTGS--SQDCEDCYSLNSNASTLSKKWRHRSKSRHN <b>I</b>
consensus	61	p v mE s 1

HsSALM2	83	<b>G</b> RSI <b>GGSAT</b> SL <b>LLPSEETSGE</b> <b>E</b> SRAAVGP <b>RRS</b> - <b>R</b> SGALEPPT <b>SAPPT</b> LA <b>L</b> VPGGAARP
MmSALM2	78	<b>G</b> RSI <b>GGSAT</b> SL <b>LLPSEETSGE</b> <b>E</b> SRAMTGP <b>RRS</b> - <b>R</b> SGALGPPT <b>SAPPT</b> LA <b>L</b> VPGGAPARP
GgSALM2	97	<b>D</b> RLMG-AFAS <b>L</b> E <b>L</b> KCQKKEETT <b>D</b> SR <b>TSTAAR</b> HSD <b>K</b> EPLL <b>G</b> QPE <b>S</b> KFRS <b>I</b> L <b>M</b> I <b>P</b> LE---G
CmSALM2	95	<b>D</b> RLMG-AFAS <b>L</b> D <b>L</b> RCQRKEDNC <b>E</b> SR <b>ASTLAHY</b> S <b>D</b> K <b>EPLL</b> <b>G</b> HSE <b>S</b> RLN <b>K</b> <b>L</b> <b>L</b> T <b>L</b> PME---V
consensus	121	R 1G SL L eSR r S r LG p S L vP

HsSALM2	142	<b>R</b> PQQRY <b>SFDGDY</b> G <b>ALFQSHS</b> <b>Y</b> P <b>RR</b> ARR <b>TKR</b> <b>H</b> R <b>S</b> TPHLDGAG <b>GGAAAG</b> <b>E</b> D <b>G</b> D <b>L</b> GLGSARACL
MmSALM2	137	<b>R</b> PQQRY <b>SFDGDY</b> G <b>ALFQSHS</b> <b>Y</b> P <b>RR</b> ARR <b>TKR</b> <b>H</b> R <b>S</b> TPHLDGAG <b>GGAAAG</b> <b>E</b> D <b>G</b> D <b>L</b> GLGSARARL
GgSALM2	152	<b>K</b> TKRSH <b>SFDMGDF</b> A <b>T</b> SQCCT <b>Y</b> P <b>KK</b> ITNIWT <b>K</b> RS <b>L</b> S---VNG <b>M</b> LLQ <b>Y</b> <b>D</b> <b>D</b> <b>N</b> <b>D</b> ---TGAKG
CmSALM2	150	<b>K</b> TKRSH <b>SFDMSDF</b> ATT <b>PCYN</b> <b>Y</b> P <b>RR</b> ITNIWT <b>R</b> RS <b>L</b> S---VNG <b>T</b> LLQ <b>Y</b> <b>D</b> <b>E</b> <b>D</b> <b>L</b> ---ESTKG
consensus	181	r <b>SFD</b> A YPrr hRS G ed DL

HsSALM2	202	A <b>F</b> T <b>STEW</b> M <b>LESTV</b>
MmSALM2	197	A <b>F</b> T <b>STEW</b> M <b>LESTV</b>
GgSALM2	205	T <b>Y</b> G <b>SSEW</b> V <b>MESTV</b>
CmSALM2	203	M <b>Y</b> C <b>SSEW</b> V <b>MESTV</b>
consensus	241	f <b>StEW</b> m <b>leSTV</b>

### SALM3

HsSALM3	1	RGRGAGNGLPLKLSHVQSQTNGGPSPTPKAHPPRSPPPRPQRSCSLDLGDA-GCYGYAR
MmSALM3	1	RGRGAGNGLPLKLSHVQSQTNGGTSPMPKSHPPRSPPPRPQRSCSLDLGDTGGCYGYAR
consensus	1	RGRGAGNGLPLKLSHVQSQTNGG SP PK HPPRSPPPRPQRSCSLDLGD GCYGYAR

HsSALM3	60	RLGGAWARRSHSVHGGLLGAGCRGVGGSAERLEESVV
MmSALM3	61	RLGGAWARRSHSVHGGLLGAGCRGVGGSAERLEESVV
consensus	61	RLGGAWARRSHSVHGGLLGAGCRGVGGSAERLEESVV

### SALM4

HsSALM4	1	MRYKVHGGQPPGKA <b>K</b> I PAP <b>VSSVCSQTN</b> ALGPTPTPA-----P <b>PAP</b>
MmSALM4	1	-RYKVHGGQPPGKA <b>KATAPVSSVCSQTN</b> ALGPVP-S-----A <b>PAP</b>
CmSALM4	1	- <b>KYKVCGSARCEVPK</b> ---LTD <b>VYSQTN</b> SQTTVPNGMVSAQRITVLNTRGQPTGGV <b>PVP</b>
consensus	1	rYKV G K vs V SQTNG P P

HsSALM4	43	E-----AALRAHTVVQLD <b>CEPWGPGHEPVCP</b> -----
MmSALM4	40	E-----AAPRAHTVVQLD <b>CEPWGPSHEPAGP</b> -----
CmSALM4	56	D <b>LSSANLPQESRKAPPYSAKTQRKRYKCKQR</b> GEGDGELET <b>TLGCQGGEGPGERTALAKQP</b>
consensus	61	e C G g

HsSALM4	-----
MmSALM4	-----
CmSALM4	116 CPQSSE
consensus	121

**SALM5**

HsSALM5	1	RYKVCNNN GQHKV TKVS NVY SQT NGA Q I Q G C S V T L P Q S V S K Q A V G H E E N A Q C C K A T - S D N
MmSALM5	1	RYKVCNNN GQHKV TKVS NVY SQT NGA Q M Q G C S V T L P Q S M S K Q A M G H E E N A Q C C K V A - S D N
GgSALM5	1	RYKVCNNN GQHK A TKVS NVY SQT NGA Q V O A C G G A L S Q S A S K Q A V G H E E A A Q C C R A A - S D G
CmSALM5	1	RYKVCNNN D Q H K M T K V S N V Y S Q T N G A H I Q M C G S V L S H S N S K V A M G H D D N I T R C N K D P S E S
consensus	1	<b>RYKVCNNN</b> QHK <b>TKVS NVY SQT NGA</b> Q C L <b>S SK AvGHee</b> C Sd

HsSALM5	60	VIQSSETCSSQ--DSSTTT SALPPSWTSSTS VS QK Q KR K T G T K P S T E P Q N E A V T N V E S Q N
MmSALM5	60	AIQSSETCSSQ--DSSTTT SALPPTW TS SAPV SOK Q KR K T G T K P S A E P Q S E A V T N V E S Q N
GgSALM5	60	AGPSPEPSPGPEATAATT TSPSPHAWAAGT SAA Q K P KR K P G P K P S S E P Q S E A A M S I E S Q N
CmSALM5	61	K T Q L S E T L S Q --DCS T T T S T L P H D W T A S V S P S Q K L K R K A G L N P S V E S P M E A F T N V E S L K
consensus	61	E TTTS P W <b>QK KR K G PS E EA</b> vES

HsSALM5	118	T N R N N S T A L Q L A S R P P D S V T E G P T S K R A H I K P N A L L T N V D Q I V Q E T Q R L E L I -----
MmSALM5	118	T N R N N S T A L Q L A S C P P D S V T E G P T S Q R A H T K P S K F L T V P A E G S R A R H R A S L S G G L K D S F H
GgSALM5	120	T N R N N S T A L Q L A S R P P D S D K G V P T Y K R A Q S K P K A G A D L K D T H -----T A P L L E S S C P N L A
CmSALM5	119	K K - E N T A I L Q K S I C A Q I S L K D T P T F R R A H S K S I K F L T L P T E I S R A K R R Y S L D A E V S E Y H C
consensus	121	N s L Q s S PT RA K L

HsSALM5	-----
MmSALM5	178 Y G N S ----- Q L S L K R S M S M N A M W T -----
GgSALM5	175 T R ----- Q K T K R S Q R T K D -----
CmSALM5	178 Y T H S Q S I N S L W S K R S M S M N G M L L Q L A N S D V D G G K A V F S S E W I M E S T V
consensus	181

## Appendix E: Motifs and Fingerprints of LIGS

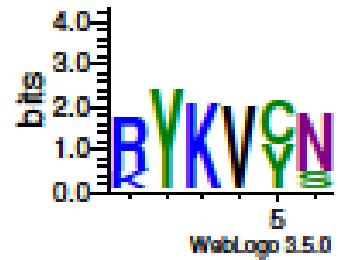
### DA1

RYKVYS

KYKVVYN

RYKVCN

RYKVCN



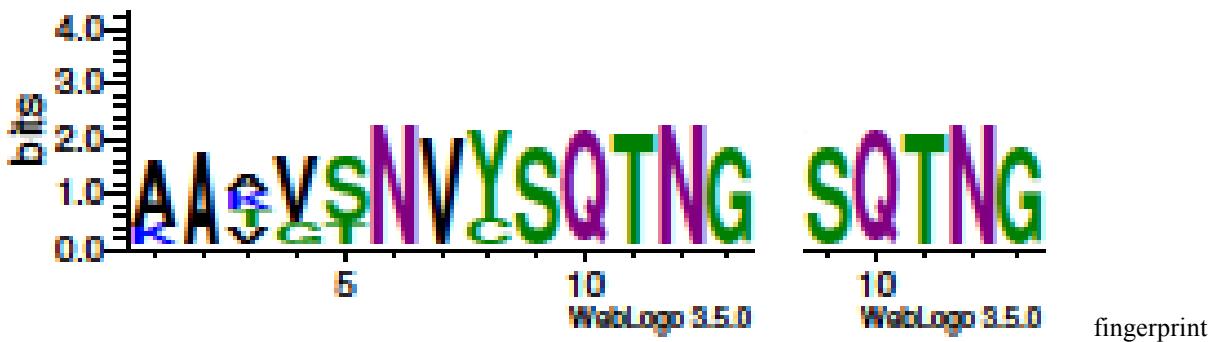
### DA2

KAVGTNVYSQTNG

AAKVSNCQTNG

A-AVSNVYSQTNG

AATVSNVYSQTNG



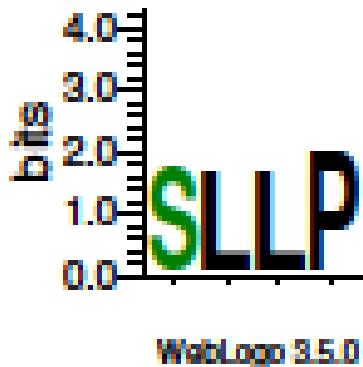
### DA3

**SLP**

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**SLLP**

**SLLP**



**WebLogo 3.5.0**

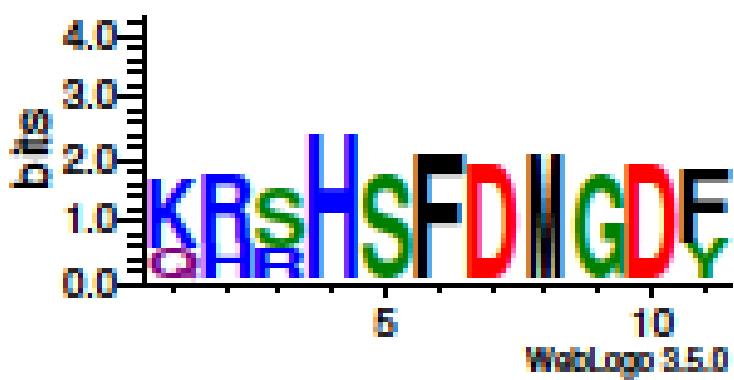
#### **DA4**

QHRHSFD-GDY

-----

KRSHSFDMGDF

KRSHSFDMGDF



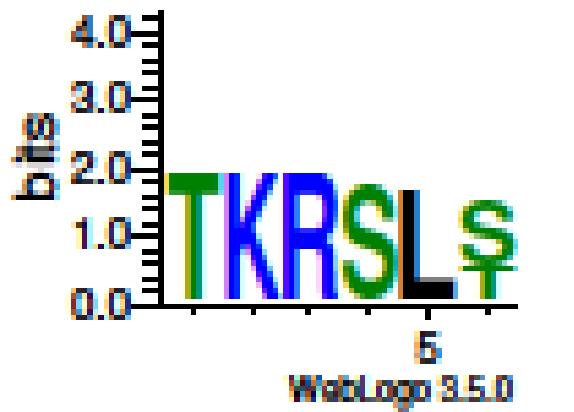
## **DA5**

TKRSLT

-----

TKRSL\$

TKRSL\$



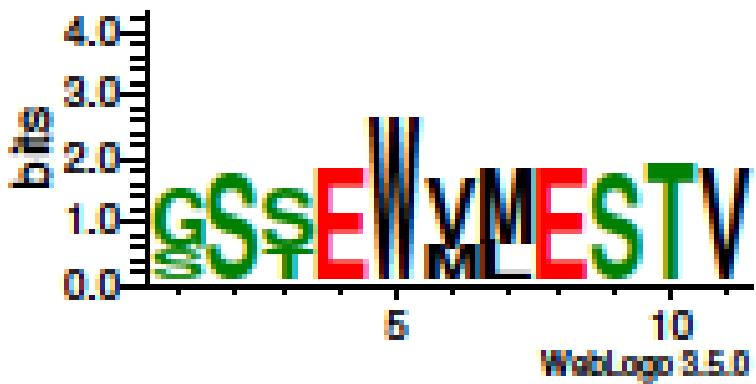
## **DA6**

SSTEWMLESTV

-----

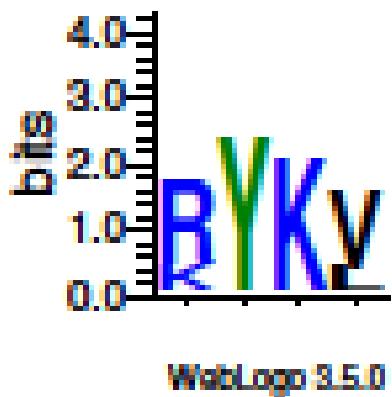
GSSEWVMESTV

GSSEWVMESTV



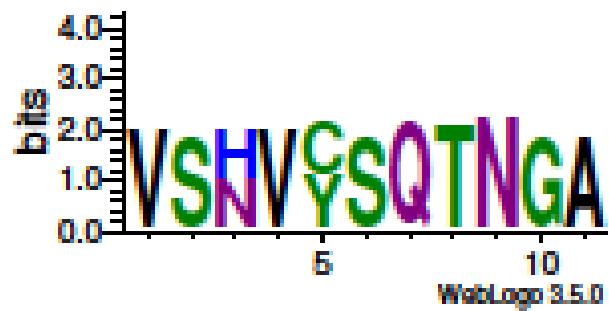
#### DA7

RYKV  
RYKV  
RYKV  
KYKL



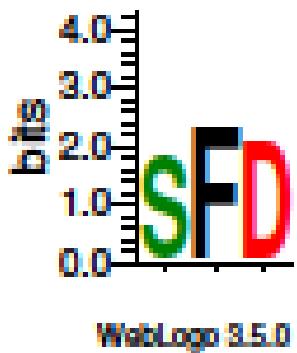
#### DA8

VSHVCSQTNGA  
VSHVCSQTNGA  
VSNVYSQTNGA  
VSNVYSQTNGA



### DA9

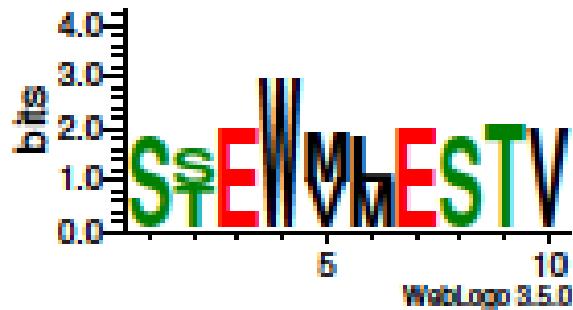
SFD  
SFD  
SFD  
SFD



### DA10

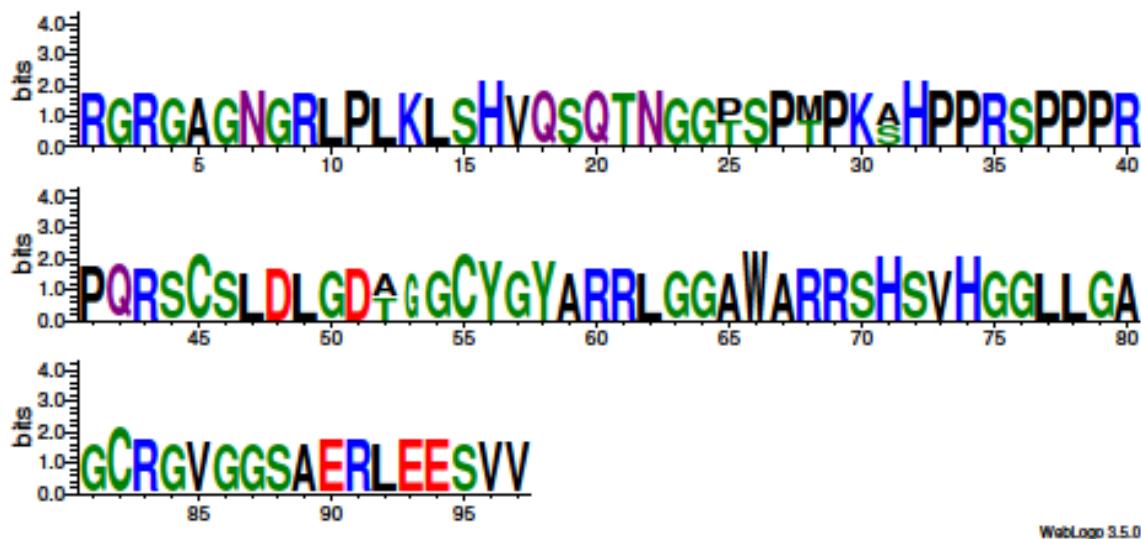
STEWMLESTV  
STEWMLESTV  
SSEWVMESTV

SSEWVMESTV



DA11

RGRGAGNGRLPLKLSHVQSQTNGGPSPPTPKAHPPRSPPPRQRSCSLDLGDA-  
GCYGYARRLGGAWARRSHSVHGLLGAGCRGVGGSAERLEESVV  
RGRGAGNGRLPLKLSHVQSQTNGGTSPMPKSHPPRSPPPRQRSCSLDGTG  
GCYGYARRLGGAWARRSHSVHGLLGAGCRGVGG  
SAERLEESVV

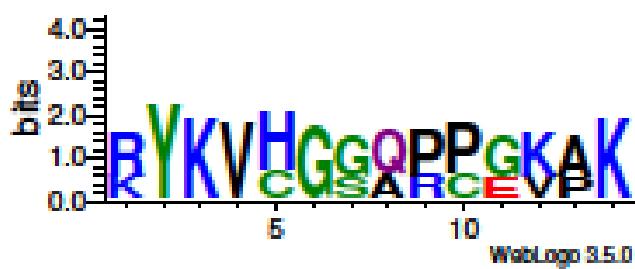
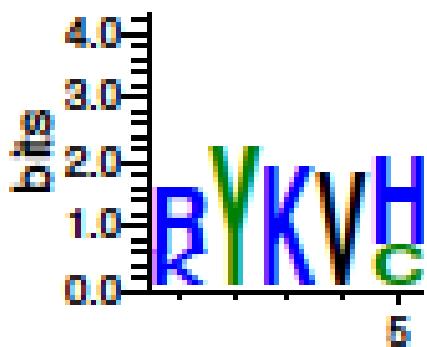


SQTNG

ESVV

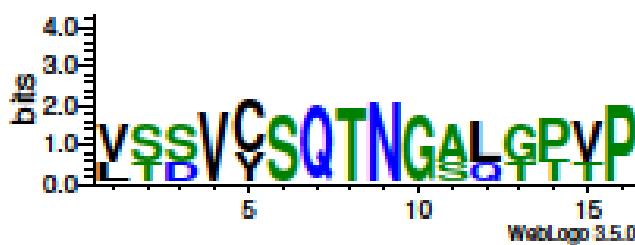
DA12

RYKVHGGQPPGKA**K**  
RYKVHGGQPPGKA**K**  
KYKVC**G**SARCEVP**K**



DA13

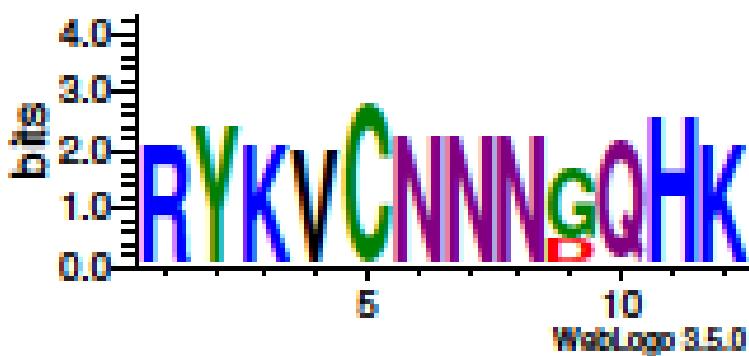
VSSVCSQTNGALGPTP  
VSSVCSQTNGALGPVP  
LTD**V**YSQTNGSQTTVP

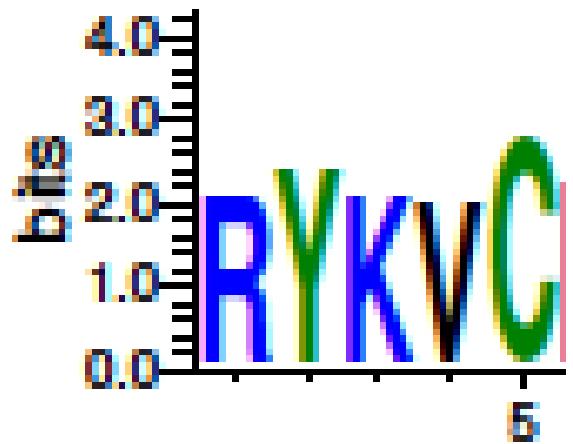


**SQTNG**  
10

DA14

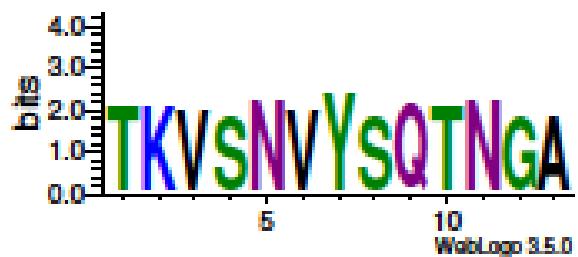
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RYKVCNNNGQHK  
RYKVCNNNGQHK  
RYKVCNNNDQHK





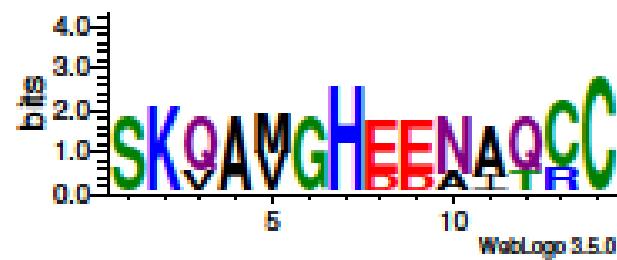
DA15

TKVSNVYSQTNGA  
TKVSNVYSQTNGA  
TKVSNVYSQTNGA  
TKVSNVYSQTNGA



DA16

SKQAVGHEE~~N~~AQCC  
SKQAMGHEE~~N~~AQCC  
SKQAVGHEEAAQCC  
SKVAMGHDDNITRC



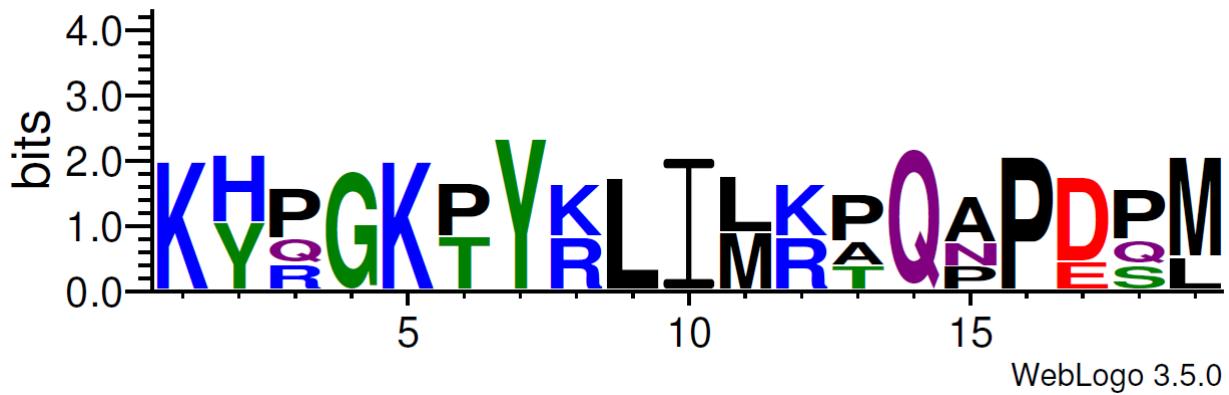
DA17

QKQKRKTGTPSTE~~P~~QNEA  
QKQKRKTGTPSAEPQSEA  
QKPKRKPGPKPSSE~~P~~QSEA  
QLKRKAGLNPSVESPMEA



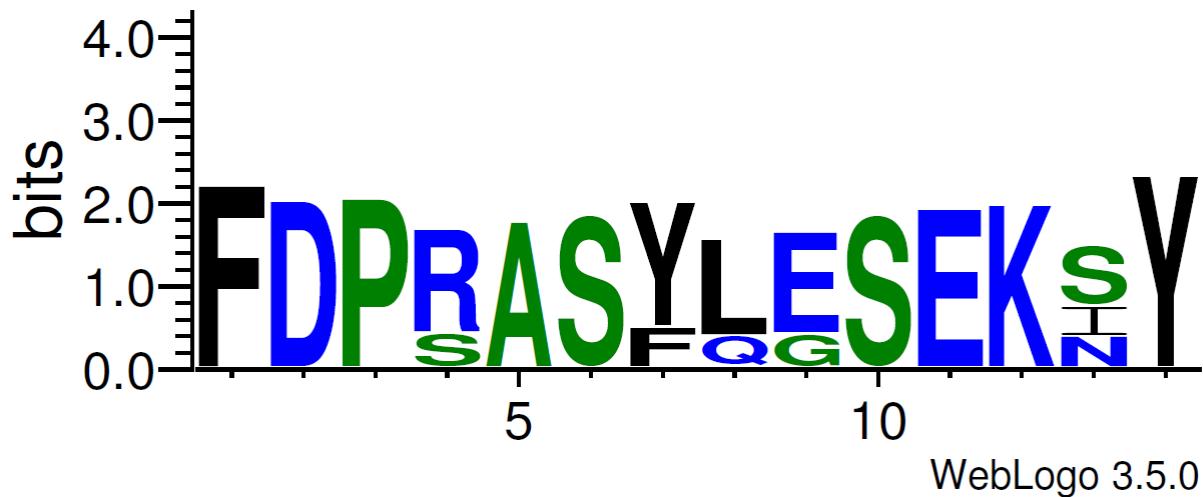
DA18

KHPGKPYRLILRPQAPDPM  
KHPGKPYRLILRPQAPDPM  
KYQGKTYKLIMKAQNPDQM  
KYRGKTYKLIMKTQPPESL



DA19

FDPRASYLESEKSY  
 FDPRASYLESEKSY  
 FDPRASYLESEKNY  
 FDPSASFQGSEKIY



DA20

LACSLVESQSKANQEEFEAGSEYSDRLPLGAEAVNIAQEINGNYRQ  
 LACCSLVESQSKANQEEFEAGSEYSDRLPLGAEAVNIAQEINGNYRQ  
 VVAASSMAESQSKANGEEFEVRSEYSDKLPLGAEAVTISQEINGNYRQ  
 VVAESVPVSQLKANPEEFEACSEYSDRLPLGAEAVNISPEINGNYRQ



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