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Computational prediction and analysis of protein structure

A dissertation submitted in partial satisfaction of the requirements for the degree

Doctor of Philosophy in Molecular Biology

by

Alejandro Daniel Meruelo

2012

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ABSTRACT OF THE DISSERTATION

Computational prediction and analysis of protein structure

by

Alejandro Daniel Meruelo

Doctor of Philosophy in Molecular Biology

University of California, Los Angeles, 2012

Professor James U. Bowie, Chair

Identifying polymer-forming SAM domains

Sterile Alpha Motif (SAM) domains are common protein modules in eukaryotic cells. It has not been possible to assign functions to uncharacterized SAM domains because they have been found to participate in diverse functions ranging from protein-protein interactions to RNA binding. Here we computationally identify likely members of the subclass of SAM domains that form polymers. Sequences were virtually threaded onto known polymer structures and then evaluated for compatibility with the polymer. We find that known SAM polymers score better than the vast majority of known non-polymers: 100% (7 of 7) of known polymers and only 8% of known non-polymers (1 of 12) score above a defined threshold value. Of 2901 SAM family members, we find 694 that score above the threshold and are likely polymers, including SAM domains from the proteins

Lethal Malignant Brain Tumor, Bicaudal-C, Liprin- β , Adenylate Cyclase and Atherin. In polymerization experiments, all of these predictions (except Adenylate Cyclase) were confirmed. As a result, the original SAM database was updated and additional predictions were obtained.

TMKink: A method to predict transmembrane helix kinks

A hallmark of membrane protein structure is the large number of distorted transmembrane helices. Because of the prevalence of bends, it is important to not only understand how they are generated but also to learn how to predict their occurrence. Here, we find that there are local sequence preferences in kinked helices, most notably a higher abundance of proline, which can be exploited to identify bends from sequence information. A neural network predictor identifies over two-thirds of all bends (sensitivity 0.70) with high reliability (specificity 0.89). It is likely that more structural data will allow for better helix distortion predictors with increased coverage in the future. The kink predictor, TMKink, is available at <http://tmkinkpredictor.mbi.ucla.edu/>.

Structural differences between mesophilic and thermophilic membrane proteins

Protein thermostability remains a focal point of interest for protein scientists. The differences in thermostability between mesophilic and thermophilic soluble proteins have been extensively studied. No differences in packing values have been found in soluble proteins. Membrane protein packing is different from soluble protein packing; thermophilic adaptation may be different as a result. Surprisingly, burial and packing values appear to be shared between mesophiles and thermophiles in both soluble and

membrane proteins. We created a non-redundant database of unpaired and paired structures for the study of thermophile-mesophile structural differences in membrane proteins. We found little or no differences in burial or packing values in both the soluble and transmembrane regions of membrane proteins.

The dissertation of Alejandro Daniel Meruelo is approved.

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Todd O. Yeates

Matteo Pellegrini

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University of California, Los Angeles

2012

To: Samantha, Dad, Mom, and Natasha

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Meruelo AD and Bowie JU. A computational approach to classifying transmembrane helical kinks. Invited talk given at *MBI Lake Arrowhead Retreat*, Lake Arrowhead, CA, October, 2009.

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Meruelo AD, Hwang BJ, and Sternberg PW. A bioinformatical technique for the identification and exploration of gene interactions using *C. elegans*, *C. briggsae*, and *C. remanei*. Presented at *SURF Summer Seminar Day*, Pasadena, CA, August, 2004.

Meruelo AD, Hwang BJ, and Sternberg PW. A bioinformatical approach to genome-wide identification of lin-3 regulatory sites across *C. elegans* and *C. briggsae*. Presented at *SURF Summer Seminar Day*, Pasadena, CA, August, 2003.

Chapter 1

Identifying polymer-forming SAM domains

SHORT COMMUNICATION

Identifying polymer-forming SAM domains

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ABSTRACT

Sterile alpha motif (SAM) domains are common protein modules in eukaryotic cells. It has not been possible to assign functions to uncharacterized SAM domains because they have been found to participate in diverse functions ranging from protein–protein interactions to RNA binding. Here we computationally identify likely members of the subclass of SAM domains that form polymers. Sequences were virtually threaded onto known polymer structures and then evaluated for compatibility with the polymer. We find that known SAM polymers score better than the vast majority of known nonpolymers: 100% (7 of 7) of known polymers and only 8% of known nonpolymers (1 of 12) score above a defined threshold value. Of 2901 SAM family members, we find 694 that score above the threshold and are likely polymers, including SAM domains from the proteins Lethal Malignant Brain Tumor, Bicaudal-C, Liprin-β, Adenylate Cyclase, and Atherin.

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Key words: protein-complex threading; protein–protein interaction; polymer prediction; sterile alpha motif; pointed domain; scaffolding proteins.

INTRODUCTION

Many common protein modules have conserved functions. For example, SH2 domains almost invariably bind to phosphotyrosines and SH3 domains almost invariably bind to proline-containing sequence motifs.¹ Thus, the presence of particular domains in a protein can often immediately suggest a functional hypothesis. On the other hand, some protein modules, such as sterile alpha motif (SAM) domains, perform a variety of different functions.² Different SAM domains can self-associate,³ bind to other SAM domains,⁴ bind to other non-SAM proteins,⁵ bind to RNA or DNA,^{6,7} or even bind lipids.⁸ Because of their diverse functions, the presence of a SAM domain does not imply a particular function, but rather a collection of possible functions. The challenge then becomes identifying which of the possible functions a particular SAM domain performs.

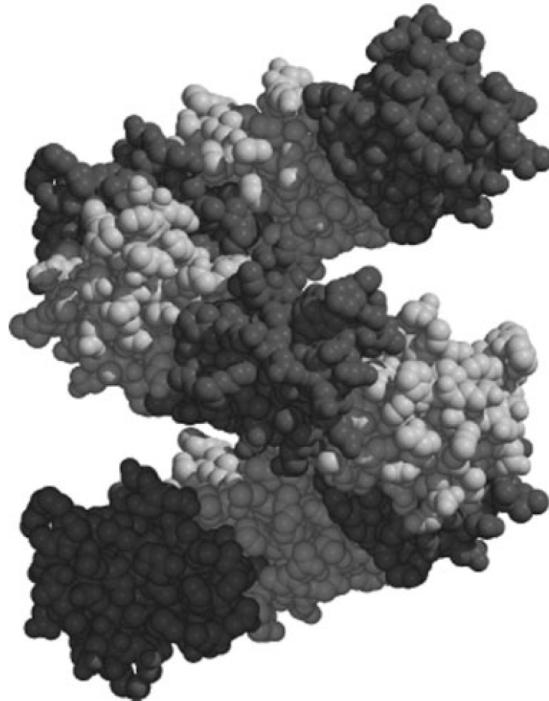
Among the SAM domains that self-associate, many of them form polymeric structures. Polymeric SAM domains have been characterized in transcriptional repressors,^{9–12} scaffolding proteins,^{13,14} and regulatory enzymes.¹⁵ So far, all the SAM polymers known to be biologically relevant have common features: they are all left-handed helices with six subunits per turn. One SAM polymer is shown in Figure 1. They differ in the precise residues used in the interface and the helical pitch, but the intersubunit contacts are made from two similar patches on the SAM domains, which have been called the Mid-Loop (ML) and End-Helix (EH) surfaces.

Although it is clear that many SAM domains form polymers, the polymerization function is difficult to characterize because polymeric SAM domains are heterogeneous assemblies that are often insoluble. It would therefore be useful to identify likely SAM polymers computationally. Here we sought to build on our current knowledge (six polymeric structures) to identify those SAM domains that are compatible with the already known polymer structures.

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**Figure 1**

SAM polymer architecture. The structure shown is the SAM domain from TEL. Every other SAM domain is shaded dark or light.

METHODS

SAM polymer template database

We constructed a database of template polymer structures initially from a collection of five known SAM polymer structures from the proteins Shank3,¹³ Diacylglycerol Kinase δ1,¹⁵ Sex-Comb on Midleg,¹¹ Translocation Ets Leukemia (TEL),¹⁰ and Polyhomeotic.⁹ We then built 11 pseudopolymers from nonpolymeric SAM domain structures by first performing a structural alignment and then substituting the sequence of the polymeric SAM with the nonpolymeric SAM. In this manner, sequences that only align to a nonpolymeric SAM can be placed into a polymeric context. The SAM domains of known structure used for the pseudopolymers were from the proteins STE11,¹⁶ EPHA4,¹⁷ EPHB2 (possibly forms an alternative type of polymer),¹⁸ ETS1,¹⁹ Mae,¹² p73,²⁰ GABPα,²¹ Erg,²¹ Fli-1 (unpublished, PDB code 1X66), Smaug,²² and VTS1.⁷ Structural alignment was performed using the combinatorial extension method (http://cl.sdsc.edu/ce/ce_align.html).²³ Each nonpolymer structure was aligned with every polymer structure and the best alignment was chosen as the template.

SAM domain sequences

About 2801 SAM domain sequences were obtained from the SMART database (<http://smart.embl-heidelberg.de>),²⁴ using the keywords SAM and SAM-PNT.

Sequence alignment

All query SAM domain sequences were aligned to the sequences of the polymer template database in two steps. First, similar sequences to the polymer template sequences were identified with PSI-BLAST²⁵ using 12 iterations. Sequences were selected if they aligned with a *P*-value less than 10^{-3} and if at least 70% of the residues could be aligned relative to the shorter of the two sequences. Similar sequences were then combined in a multiple sequence alignment using the program CLUSTALW (<http://www.ebi.ac.uk/clustalw/>).²⁶ Using this alignment, residues from similar sequences could be mapped onto the polymer structure.

Sequence-structure scoring

The interface energy of two neighboring chains, ΔG_{bind} , was computed using a dimer interface potential.²⁷ We selected interacting pairs of residues between subunits from the known polymer structures if any of the heavy atoms were within 4.5 Å and both residues had a solvent accessibility of less than 0.4. The identity of the residues used for calculation of ΔG_{bind} was from the sequence alignment (discussed earlier).

To adjust for sequence composition effects, the energy scores were compared with the energies of decoy alignments and a *Z*-score determined using the expression⁴:

$$\text{Z-score} =$$

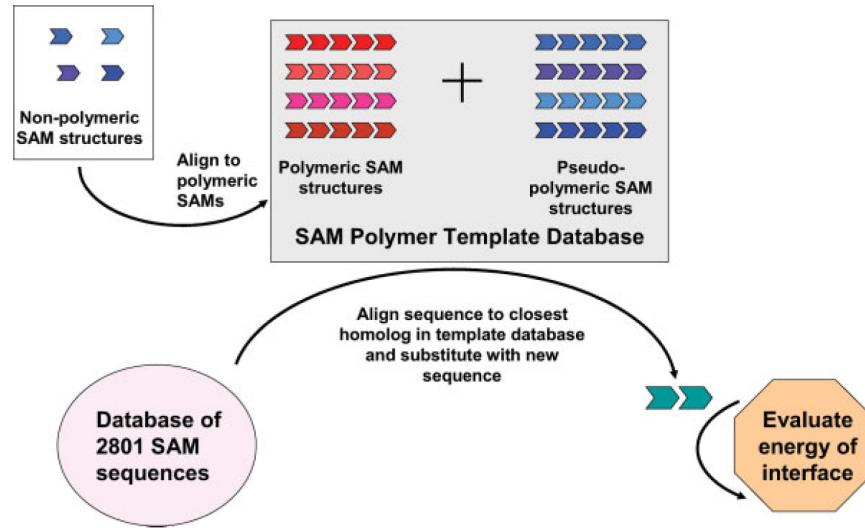
$$(\langle \Delta G_{\text{bind}}^{\text{d}} \rangle - \langle \Delta G_{\text{bind}} \rangle) / (\langle (\Delta G_{\text{bind}}^{\text{d}})^2 \rangle - \langle \Delta G_{\text{bind}}^{\text{d}} \rangle^2)^{1/2}$$

where $\Delta G_{\text{bind}}^{\text{d}}$ refers to the interface energy of the decoy alignments. Decoy alignments were created by circular permutation of the query sequence. If a sequence could be mapped to more than one template, the most positive *Z*-score was selected. To reduce false positives, we averaged the *Z*-scores of each sequence with the *Z*-scores of very close homologs, defined by a PSI-BLAST *P*-value $< 10^{-18}$, to obtain the final score, *Z*-final.

RESULTS AND DISCUSSION

The algorithm

Our algorithm is based on the protein-complex threading idea originally developed by Skolnick and co-workers.²⁷ All known SAM domain sequences were threaded onto all known polymer structures (templates) and the compatibility of the aligned sequence with the polymer interface evaluated as outlined in Figure 2. This

**Figure 2**

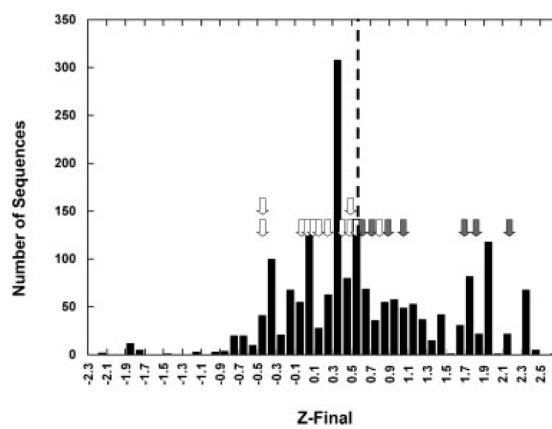
Outline of the polymer compatibility test algorithm. See text.

was accomplished by first identifying all sequences similar to each template sequence using PSI-BLAST.²⁵ The similar query sequences were then combined in a multiple sequence alignment using ClustalW,²⁶ thereby mapping positions of the query sequences onto the template structure. The amino acid type in the polymer template structure was then replaced with the amino acid type in each related query sequence. The energy of the replaced sequence in the polymer structure was then evaluated using a statistical potential function that measures the likelihood of finding different contacting amino acid pairs in a protein–protein interface.²⁷ To account for sequence composition effects, the energy was compared with the energies of 100 circularly permuted alignments to obtain a Z-Score, which is the number of standard deviations away from the mean of circularly permuted alignments. The best Z-Score for a given query sequence on all template polymers was selected. Finally, to reduce noise, a Z-final score was calculated, which is an average of the best Z-scores of the query sequence and all very closely related sequences.

Score distribution and polymer identification

Starting with 2801 SAM domain sequences, we were able to find 1885 significant alignments to the known polymers. The distribution of Z-final scores for the 1885 aligned query SAM domain sequences is shown in Figure 3. The Z-final scores for known polymers and presumed nonpolymeric SAM domains are noted in Figure 3 and are listed in Table I. For testing the scoring of the known polymers and nonpolymers, we did not allow alignments

of known polymers to their own structures, providing a more realistic scenario. As can be seen, the vast majority of known polymers score better than known nonpolymers. All of the seven known polymers score higher than a Z-final value of 0.65 and all but one of the 12 nonpolymers scored below 0.56. We therefore picked the mean of these Z-final values, 0.61, as an arbitrary threshold defining likely polymers. As we are using a previously defined

**Figure 3**

Histogram of the Z-final score distribution. The scores for the known polymers are indicated by the positions of the shaded arrows and the scores for the known nonpolymers are indicated by the positions of the white arrows. The dotted line indicates the Z-final cutoff used to separate polymers from nonpolymers.

Table I
Z-Scores of known Polymers and Nonpolymers

Polymers	Z-score	Nonpolymers	Z-score
Scm	2.253	Erg	-0.410
Ph	1.848	Fli-1	-0.406
Shank	1.687	BYR2	-0.061
Tankyrase-1	0.976	p73	0.004
DGK	0.835	Mae	0.064
TEL	0.716	Smag	0.129
Yan	0.653	EPHA4	0.290
		GABPA	0.460
		Pnt-P2	0.483
		ETS1	0.493
		STE11	0.563
		VTS1	0.792

energy function, the only variable parameter here is, in fact, the threshold.

We find that 694 SAM domain sequences of the 1885 that could be aligned to the known polymers or pseudo-polymers score higher than the cutoff of 0.61 and are potentially polymers. A list of all the SAM domains that could be aligned and their Z-final scores are given in Supporting Information. In addition to family members of known SAM domain polymers, a number of other protein families were found consistently among the high scoring proteins and may therefore contain polymeric SAM domains.

Families containing possible polymeric SAM domains

Of the 21 lethal malignant brain tumor (LMBT) homologs, all but two had Z-final scores greater than 1.197, much higher than the threshold of score of 0.61. Like the polymeric SAM-containing proteins Scm and Ph, LMBT proteins are polycomb group proteins, involved in the maintenance of transcriptional repression. The SAM domain of L(3)MBT is known to self-associate.²⁸ L(3)MBT was found to work with in conjunction with another polymeric SAM-containing protein, TEL, and the SAM domains of TEL and L(3)MBT were found to bind to each other.²⁸ It is therefore possible that the SAM domain of L(3)MBT forms a polymer that could extend the polymer formed by TEL-SAM or could even form a mixed copolymer with TEL-SAM. A similar mechanism has been proposed for the SAM polymer interactions of Ph and Scm.¹¹ It may be possible to adapt our algorithm to look specifically for copolymers, but we may need to find effective ways to reduce the combinatorial complexity.

Another protein family important for development, Bicaudal-C, was also found among the top scoring SAM domains. Out of 11 Bicaudal-C homologs, the lowest Z-final was 1.321. Bicaudal-C is an RNA binding proteins that represses expression by regulating polyadenylation.²⁹ It is possible that SAM polymerization could enable

spreading along the RNA transcripts or the SAM domain could provide a scaffolding role similar to the Shank¹³ and Tankyrase¹⁴ SAM domains.

Liprins were originally identified as proteins that bind to protein tyrosine phosphatases at focal adhesions.³⁰ There are two classes of liprins: α and β .³¹ Liprin- α is involved in organizing the active zone of neural synapses and is known to bind to a number of other proteins.³² Less is known about the liprin- β class, but they interact directly with liprin- α proteins.³¹ The liprins are somewhat unusual in that they contain three tandem SAM domains. Some of the SAM domains in the liprin- α class bind to other proteins.³³ We find here that the second SAM domain of the liprin- β class scores highly for polymer formation, suggesting that liprin- β could create a scaffold for organizing liprin- α and its binding partners.

We also found that a family of adenylate cyclases contains potential polymeric SAM domains. This family is predicted to be expressed as multiple splice variants, one that contains the SAM domain and one that does not (see <http://dylan.embl-heidelberg.de/>). The SAM-containing splice variant does not include the adenylate cyclase catalytic domain. It is therefore possible that the SAM-containing splice variant could be a scaffolding protein.

The SAM domain from the protein Atherin scores highly for polymer potential. Atherin is associated with atherosclerotic plaques and may recruit low-density lipoprotein complexes by direct binding.³⁴ Atherin polymerization could create a matrix for plaque formation. If so, inhibition of polymer formation could be a potential antiatherosclerotic therapy.

CONCLUSIONS

In this work, we have assigned a possible polymerization function to hundreds of uncharacterized SAM domains, providing a testable functional hypothesis for these proteins. The methodology could be applied to other domain types to predict polymerization.

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Appendix 1

While many previously uncharacterized SAMs were predicted to polymerize, no experimental characterization had been completed by the time of publication of the above paper. Experimental validation was later done by Mary Jane Knight *et al*¹. In her work, she developed a method to rapidly screen SAM polymers. SAM domains were fused to super-negatively charged green fluorescent protein. Using these fusions, a large library of human SAM domains were screened for polymerization using a native gel screen. All experimentally tested predictions were found to be correct as has been summarized below in Table 1-2¹; Adenylate Cyclase could not be cloned out for expression and so was not able to be experimentally studied².

Table 1-2: Experimental validation of SAM polymer predictions

Predicted polymer	Experimentally confirmed
Atherin	Yes
Bicaudal-C	Yes
Liprin- β	Yes
Lethal Malignant Brain Tumor	Yes
Adenylate Cyclase	Not tested

Because of these experimental successes, we updated our SAM database. We added 21 new structures to our mapping database to make for a total of 37 (provided below). We used the non-redundancy criteria of requiring that all minimum P-value structures with > 90% identity be thrown out. In addition, the SAM database was expanded to

include 3500 candidates from the original 3000. A number of new predicted polymers (e.g., Synaptotagmin-3) were found as a result and are tabulated in Table 1-3 below.

New structures (21): ARAP2, BIFUNCTIONAL_APOPTOSIS_REG, DCIP, EGFPS8, ELF3, EPHA2, EPHA8, EPHB1, EPHB4, FLJ21935, HCNK1, KIAA0902, LCP2, LOC64762, Neurabin, PDEF, PUTATIVE_47_KDA, SAMSN1, SASH1, Ship2, and SPHINGOMYELIN_SYNTHASE

Table 1-3: Summary of new predicted polymers

Protein	Function
UPI000155B956/44-126	PREDICTED: similar to B-cell lineage specific activator/TEL oncogene fusion protein, partial
UPI0000E4781C/1217-1284	PREDICTED: similar to mitogen-activated kinase kinase kinase 5
UPI0000F2AE6A/869-935	PREDICTED: similar to serologically defined colon cancer antigen 3
Q6BKM8_DEBHA/264-332	Similar to CA5428 CaBOI2 Candida albicans CaBOI2 budding protein
UPI000155472E/1-65	PREDICTED: similar to N-acylaminoacyl-peptide hydrolase
Q96X32_USTMA/14-80	MAP kinase pathway-interacting Ubc2
UPI00004D61B3/1152-1218	Synaptotagmin-3 (Ca sensor involved in both (i) early synaptic vesicle docking to the presynaptic membrane (ii) late steps of Ca-evoked synaptic vesicle fusion with the presynaptic membrane)
UPI0000E80BD6/971-1038	PREDICTED: similar to early development regulator 3
Q5KJP5_CRYNE/17-83	Protein kinase regulator, putative

References

1. Knight MJ, Leettola C, Gingery M, Li H, Bowie JU (2011 [cited 2012]) A human sterile alpha motif domain polymerizome. *Protein Science* 20:1697–1706.
2. Knight MJ (2010) Re: Adenylate cyclase SAM polymer.

Appendix 2

The list of new polymer and non-polymer predictions has been included below. The first line consists of the protein identifier and residue range. The second line provides the species description. The third line provides the amino acid sequence for the SAM. The last (and summary) line provides the residue range, simple protein identifier, and Z-score for polymer prediction. Z-scores with a value of 0.61 or higher were predicted as polymers and those below this value were predicted as nonpolymers.

```
>smart|SAM-uniprot|Q29KC7|Q29KC7_DROPS/849-915 GA18459-PA
[Drosophila pseudoobscura]
MQLSQHKDIQTLLTSLGLEHYIKIFVLNEIDLEMFTTLTEENLMEGITAFGARKKLLAAIHTL
LAN
849-915_Q29KC7 -2.297
>smart|SAM-uniprot|Q7QYZ7|Q7QYZ7_GIALA/5-63 GLP_164_5127_2983
[Giardia intestinalis]
VYQWLESANLQQYYPAFEQQGITPQRFITITIQDYGALGIQALPDKQKLFRLLTTLKSR
5-63_Q7QYZ7 -2.160
>smart|SAM-uniprot|Q5TRB3|Q5TRB3_ANOGA/1-73 ENSANGP00000029645
[Anopheles gambiae]
VLGWSCENVSDWARKEGLSRCIIDWIAREDIDGRCLLAITEQDVHDLQHCASPLRFGDIKRFW
FATRLLQRO
1-73_Q5TRB3 -2.032
>smart|SAM-uniprot|UPI0000F20118|UPI0000F20118/578-645
PREDICTED: SAM domain, SH3 domain and nuclear localisation
signals, 1 [Danio rerio]
CKKPRPNTLLELLERLHLEEYASSLLLNGYQTVDDLRLKERHLIELNVTDPEHRHRLLAASDC
LYVT
578-645_UPI0000F20118 -1.842
>smart|SAM-uniprot|Q4RQC8|Q4RQC8_TETNG/8-77 Chromosome 17
SCAF15006, whole genome shotgun sequence [Tetraodon
nigroviridis]
VSAWSCVQVAHWLQKEGFGEYVDLLCARHRLDGLGLLALTEADLRGPPLGLAVLGDIKRLVIAL
RRLQRR
8-77_Q4RQC8 -1.817
>smart|SAM-uniprot|UPI0000584492|UPI0000584492/3-72 PREDICTED:
hypothetical protein isoform 4 [Strongylocentrotus purpuratus]
```

VESWTCKDVAKWLSTKGFQYARQLCTNHTIDGQALLLTEADLRSPVSMKVLGDIKRQLIRQI
DLLKHQ

3-72_UPI0000584492 -1.812

>smart|SAM-uniprot|UPI00005A0829|UPI00005A0829/72-141 PREDICTED:
similar to sterile alpha motif domain containing 8 (predicted)
[Canis lupus familiaris]

IRRWTTKHVAWLKDEGFFEYVDVLCNKHRLDGITLLTLEYDLRSPPLEIKILGDIKRLMLSV
RKLQKI

72-141_UPI00005A0829 -1.800

>smart|SAM-uniprot|UPI000069F923|UPI000069F923/608-675 SAM and
SH3 domain-containing protein 1 (Proline-glutamate repeat-
containing protein). [Xenopus tropicalis]

KGLLKPNSVEDLLERINLKEHMPTFLFNGYEDLDAFKLLEEDLDELNIQNPEHKAVLLTAIEL
LQEY

608-675_UPI000069F923 -1.763

>smart|SAM-uniprot|UPI0000E80117|UPI0000E80117/819-886
PREDICTED: similar to putative adapter and scaffold protein
[Gallus gallus]

SRPPQPKSVEDLLDRINLKEHMPTFLFNGYEDLDTFKLLEEDLDELNIRDPEHRAVLLTAVEH
LQEY

819-886_UPI0000E80117 -1.751

>smart|SAM-uniprot|Q4REI4|Q4REI4_TETNG/584-651 Chromosome 10
SCAF15123, whole genome shotgun sequence [Tetraodon
nigroviridis]

ARQPKPISVEELLERINLKEHLPTFLFNGYEDLDTFKLLEEDLDELNIRDPSQHRAVLLTAVEL
LQEY

584-651_Q4REI4 -1.749

>smart|SAM-uniprot|UPI000065E67A|UPI000065E67A/973-1034 Homolog
of Homo sapiens "SAM and SH3 domain containing protein 1
[Takifugu rubripes]

FGSWLKRTVSFAQEHMPTFLFNGYEDLDTFKLLEEDLDELNIRDPSQHRAVLLTAVELLQEY

973-1034_UPI000065E67A -1.698

>smart|SAM-uniprot|Q4S703|Q4S703_TETNG/1001-1074 Chromosome 14
SCAF14723, whole genome shotgun sequence [Tetraodon
nigroviridis]

RPEVLLSGLSEHMRSLIEDVSFTQEHMPTFLFNGYEDLDTFKLLEEDLDELNIRDPSQHRAVL
LTAVELLQEY

1001-1074_Q4S703 -1.690

>smart|SAM-uniprot|UPI00006605E2|UPI00006605E2/506-573 Homolog
of Homo sapiens "SAM and SH3 domain containing protein 1
[Takifugu rubripes]

ARQPKPTSVEELLVRINLKEHLPTFLFNGYEDLDTFKLLEEEEDLDELNKDSQHRAVLLTAVEL
LQED

506-573_UPI00006605E2 -1.670

>smart|SAM-uniprot|Q299I9|Q299I9_DROPS/1-65 GA14584-PA
[Drosophila pseudoobscura]

MCPHAVREWLVLTMEMYMDRFMKNGYDTIAQCKLIVPRDLKILGIYHPGHRKLLVDGVQLMNN
A

1-65_Q299I9 -1.669

>smart|SAM-uniprot|UPI0000F20302|UPI0000F20302/609-676
PREDICTED: hypothetical protein [Danio rerio]

SRPPKPTSVEELLERISLKEHMPTFLFNGYEDLDTFKLLEEEEDLDELNISDPQHRAVLLTAVEL
LQEC

609-676_UPI0000F20302 -1.663

>smart|SAM-uniprot|UPI0000E21203|UPI0000E21203/805-872
PREDICTED: SAM and SH3 domain containing 1 [Pan troglodytes]

GRPPQPKSVEDLLDRINLKEHMPTFLFNGYEDLDTFKLLEEEEDLDELNIRDPEHRAVLLTAVEL
LQED

805-872_UPI0000E21203 -1.657

>smart|SAM-uniprot|Q5TZ33|Q5TZ33_DANRE/625-692 Novel protein
similar to vertebrate SAM and SH3 domain containing 1 [Danio
rerio]

GRPPKPSSVEELLERINLKEHMPTFLFNGYEDLDTFKLLEEEEDLDELNKDPQHRAVLMTAVEL
LQDY

625-692_Q5TZ33 -1.584

>smart|SAM-uniprot|Q7Q002|Q7Q002_ANOGA/22-92 ENSANGP00000014193
[Anopheles gambiae]

VYMWTVADTLKWLWRHCNERCVKYQELFQRHEITGRALLRITDEQTLYRMGIHDEKDRGDILRE
IIKQRLK

22-92_Q7Q002 -1.575

>smart|SAM-uniprot|P59808|SASH1_MOUSE/623-690 SAM and SH3
domain-containing protein 1 [Mus musculus]

GRPSQPKSVEDLLDRINLKEHMPTFLFNGYEDLDTFKLLEEEEDLDELNIRDPEHRAVLLTAVEL
LQEY

623-690_P59808 -1.574

>smart|SAM-uniprot|UPI0000EBCFE4|UPI0000EBCFE4/801-868
PREDICTED: similar to putative adapter and scaffold protein [Bos
taurus]

GRPPQPKSVEDLLDRINLKEHMPTFLFNGYEDLDTFKLLEEEEDLDELNIRDPEHRAVLLTAVEL
LQEY

801-868_UPI0000EBCFE4 -1.562

>smart|SAM-uniprot|UPI00006A0C3A|UPI00006A0C3A/243-310 SAM
domain-containing protein SAMSN-1 (SAM domain, SH3 domain and

nuclear localization signals protein 1) (SH3-SAM adaptor protein) (Hematopoietic adaptor containing SH3 and SAM domains 1). [Xenopus tropicalis]
SKRPKPCTLQELMERLNLDYISSLLLNGYEKLEDLKDLKESHLHELNITSSEERARLLAAIEN
LQDC

243-310_UPI00006A0C3A -1.555
>smart|SAM-uniprot|UPI0000D5574F|UPI0000D5574F/663-729
PREDICTED: similar to CG31163-PB, isoform B [Tribolium castaneum]
IKGKPISVEDLLQKINLQEYISVFVLNGYEDLELFKEIEPSLDYLGIVNSDHRSKILTAVOLL
HDL

663-729_UPI0000D5574F -1.476
>smart|SAM-uniprot|Q7QBB0|Q7QBB0_ANOGA/610-677
ENSANGP00000020309 [Anopheles gambiae]
LPTSCPTSVEELLRIGLKEYTSFVLNGYEDLELFKELEPSLDYLGIGNTEHRAKILAVAL
LHDL

610-677_Q7QBB0 -1.473
>smart|SAM-uniprot|Q4S3R6|Q4S3R6_TETNG/622-687 Chromosome 17
SCAF14747, whole genome shotgun sequence [Tetraodon nigroviridis]
EDGEDFEDLLSVLQHLGLSEYKTTFDDEKIDIESFLLCTIEDLKEMGIPLGPRKKIAKFVKERV
NK

622-687_Q4S3R6 -1.459
>smart|SAM-uniprot|UPI000051A64D|UPI000051A64D/427-494
PREDICTED: similar to CG31163-PB, isoform B, partial [Apis mellifera]
RYRQKPGSVQELLQRMNLQEHIPVVFNGYEDLELFREIEAADLDYLRIHQPEHRAKILTAVQL
LNLDL

427-494_UPI000051A64D -1.448
>smart|SAM-uniprot|Q8SZS0|Q8SZS0_DROME/737-804 LD31046p
[Drosophila melanogaster]
SHNGGPCSVEDLLIRIGLKEYTSFVLNGYEDLELFKELEPADLDYLGILNQEHRRAKLLTAVQL
LHDII

737-804_Q8SZS0 -1.446
>smart|SAM-uniprot|UPI000065F73B|UPI000065F73B/75-142 Homolog of Homo sapiens "Splice Isoform 1 of SAM-domain protein SAMSN-1 [Takifugu rubripes]
CKGAPPQSLLELLQSLNLEDYASALLPEGCQTVQDLLNPQKKQLIELNIKDPEHRYRLLAAEY
FNTE

75-142_UPI000065F73B -1.425
>smart|SAM-uniprot|Q3U754|Q3U754_MOUSE/1-60 Bone marrow macrophage cDNA, RIKEN full-length enriched library,

clone:I830086L09 product:SAM domain and HD domain, 1, full
 insert sequence [Mus musculus]
 CSFLENRGFREKKVLDIFRDNKIAGSFLPFLDEDRLEDLGVSLEERKKMIECIQQLSQS
 1-60_Q3U754 -1.418
 >smart|SAM-uniprot|A0CUC0|A0CUC0_PARTE/87-154 Chromosome
 undetermined scaffold_28, whole genome shotgun sequence
 [Paramecium tetraurelia]
 PKKWSLEDVSQWLNFVGLQQLQTTFTKNSIDGSCLELIEENDLIEDL GITNKIVRK
 KLMHWLKT GLKE
 87-154_A0CUC0 -1.410
 >smart|SAM-uniprot|Q8ML92|AVE_DROME/23-91 Protein aveugle
 [Drosophila melanogaster]
 VYLWTVSDVLKWYRRHCGEYTQYEQLFAQHDITGRALLRITDSSLQRMGVTDNRD
 REAIWREIV KQRLK
 23-91_Q8ML92 -1.408
 >smart|SAM-uniprot|Q5TZ33|Q5TZ33_DANRE/1120-1187 Novel protein
 similar to vertebrate SAM and SH3 domain containing 1 [Danio
 rerio]
 VSLVHVSSVSDWLVSIGMPMYSAPLLAAGFDTLSRVSSLNEAQARDAGLKEEH
 HIRILLSEAQL VSAA
 1120-1187_Q5TZ33 -1.397
 >smart|SAM-uniprot|UPI0000DA2491|UPI0000DA2491/41-109 PREDICTED:
 similar to SAM domain and HD domain-containing protein 1
 (Interferon-gamma-inducible protein Mg11) [Rattus norvegicus]
 LRTWGPEDVCSFLEKRGFREKKVLDIFRENKISGSFLPFLDENLLES LGVSSLEQR
 KKMIECIQ KLNQS
 41-109_UPI0000DA2491 -1.382
 >smart|SAM-uniprot|A6NCH1|A6NCH1_HUMAN/249-316 Uncharacterized
 protein CXorf9 [Homo sapiens]
 GKRPKPCTLHELLERIGLEEHSTLLLNGYQTLEDFKELRETHLNELNIMDPQHRA
 KLLTAAEL LLDY
 249-316_A6NCH1 -1.380
 >smart|SAM-uniprot|A5D8Q2|A5D8Q2_XENLA/240-307 LOC100049772
 protein [Xenopus laevis]
 SKRQKPKTVQELLERINMQEHIPTLLLNGYETIEDFKDLSENHLIELNITDPQH
 RVKLLTAAEF LLDY
 240-307_A5D8Q2 -1.376
 >smart|SAM-uniprot|UPI0000D9C6DD|UPI0000D9C6DD/144-212
 PREDICTED: SAM domain- and HD domain-containing protein 1
 [Macaca mulatta]
 YKTWGPEQVCFLLRRGGFGEPELLKNIRENKITGALLPCLDESHFENLG
 VSSLGERKKLLSYIQ RSGQI

144-212_UPI0000D9C6DD -1.374
>smart|SAM-uniprot|UPI00005A459C|UPI00005A459C/80-148 PREDICTED:
similar to SAM domain and HD domain-containing protein 1
(Dendritic cell-derived IFNG-induced protein) (DCIP) (Monocyte
protein 5) (MOP-5) [Canis lupus familiaris]
HQSWGPEQVCCFLKRSGFGDPRLLERFRENKITGLTLLHLNEADLERLGISTLGDRKKLQACIQ
QLSEI

80-148_UPI00005A459C -1.365
>smart|SAM-uniprot|Q291B9|Q291B9_DROPS/21-89 GA15868-PA
[Drosophila pseudoobscura]
VYQWTVSDVLKWYRRHCGEYSQYDQLFVQHDITGRALLRITDTSLQRMGVTDNRDREAIWREIV
KQRLK

21-89_Q291B9 -1.352
>smart|SAM-uniprot|UPI0000F2B3C2|UPI0000F2B3C2/40-108 PREDICTED:
similar to MOP-5 [Monodelphis domestica]
PQSWGPEQLCAFLNHQGFSDPALLSRFRENKIRGSQLLHLDESLLNLGVRSSEDROKLLDCVY
KLNQS

40-108_UPI0000F2B3C2 -1.349
>smart|SAM-uniprot|A1L2P0|A1L2P0_XENLA/637-702 LOC100036906
protein [Xenopus laevis]
EQQKPLPSLREALESLSAEYSSVFEKEQIDMESLLMCTVDDLKEMGIPLGPRKKIAKFVQTKR
KA

637-702_A1L2P0 -1.324
>smart|SAM-uniprot|UPI0000E47174|UPI0000E47174/3-70 PREDICTED:
similar to centaurin delta 2 isoform a variant
[Strongylocentrotus purpuratus]
QSGNGPSSVQEWLQSLHLNQFLNDFLFSGYTDLSKCCHITEADLVRLGITQVGYQKRILSRLPS
KADL

3-70_UPI0000E47174 -1.322
>smart|SAM-uniprot|Q68FK2|Q68FK2_XENLA/644-709 MGC84165 protein
[Xenopus laevis]
EQQKPLPSLQEALELLSLSAEYSSVFEKEQIDMESLLMCTVDDLKEMGIPLGPRKKIAKFVQERA
VT

644-709_Q68FK2 -1.321
>smart|SAM-uniprot|UPI0000D77D07|UPI0000D77D07/623-688
hypothetical protein LOC560750 [Danio rerio]
EEEEETGSSLSSVLEAIGLSEYLSIFENEKIDVESLLLCTVDDLKEMSIPLGPRKKLANFIKEKS
AK

623-688_UPI0000D77D07 -1.301
>smart|SAM-uniprot|UPI000155C77E|UPI000155C77E/646-711
PREDICTED: hypothetical protein [Ornithorhynchus anatinus]

SIDNGISTLQETLEMLSLSEYISTFEKEKIDMESLLMCTVDDLKEMGIPPLGPRKKIANFVKDKA
TE

646-711_UPI000155C77E -1.299
>smart|SAM-uniprot|Q9Y6Y8|S23IP_HUMAN/641-706 SEC23-interacting
protein [Homo sapiens]
VENKEVLTLQETLEALSLSEYFSTFEKEKIDMESLLMCTVDDLKEMGIPPLGPRKKIANFVEHKA
AK

641-706_Q9Y6Y8 -1.296
>smart|SAM-uniprot|UPI0000F2B03E|UPI0000F2B03E/638-702
PREDICTED: hypothetical protein [Monodelphis domestica]
VENEVLTLQETLEELSLSEYISTFEKEKIDMESLLMCTIDDLKEMGIPPLGPRKKIANFVKDKAT
K

638-702_UPI0000F2B03E -1.293
>smart|SAM-uniprot|UPI0000D8CDA0|UPI0000D8CDA0/227-294
hypothetical protein LOC450018 [Danio rerio]
SRRPRPKTLQELLERLNLEEHISLLLNGYQTVEDLRDLKEQHLVELNMTDPEHRHRLLA
AAEN
LLDS

227-294_UPI0000D8CDA0 -1.263
>smart|SAM-uniprot|Q4S4R5|Q4S4R5_TETNG/351-417 Chromosome 2
SCAF14738, whole genome shotgun sequence [Tetraodon
nigroviridis]
STPVFLQDLAEFLEQIGFSKYLPPLLQEVDVDLRIFLTLTENDLKEIGITLFGPKRKMTSAIARW
HSS

351-417_Q4S4R5 -1.255
>smart|SAM-uniprot|UPI0000F213F2|UPI0000F213F2/406-472
PREDICTED: hypothetical protein [Danio rerio]
PSYTGPKDLSFLEQIGFSKYLPLEEQDIDLRIFLTLTENDLKEIGITLFGPKRRMTSAIARW
HSN

406-472_UPI0000F213F2 -1.254
>smart|SAM-uniprot|UPI00005E9A1F|UPI00005E9A1F/423-489
PREDICTED: hypothetical protein [Monodelphis domestica]
PTYSGPQDLATLLEQIGCLKYLQVFEEQDVDLRIFLTLTESDLKEIGITLFGPKRKMTSAIARW
HSS

423-489_UPI00005E9A1F -1.229
>smart|SAM-uniprot|UPI0000ECD45F|UPI0000ECD45F/373-440 SAM
domain, SH3 domain and nuclear localisation signals, 1 [Gallus
gallus]
SKRNKPCTLQELLERVHLOEYTSTLLLNGYETLEDLKDLHESHLIELNISNPEDRARLLSAIEN
LQDN

373-440_UPI0000ECD45F -1.228
>smart|SAM-uniprot|Q9D4W6|Q9D4W6_MOUSE/406-473 Adult male testis
cDNA, RIKEN full-length enriched library, clone:4930548L11

product: SAM domain, SH3 domain and nuclear localisation signals,
 1, full insert sequence [Mus musculus]
 RRRENHQTIQEFLERIHLQEYTSTLLLNGYETLDDLKDIKESHLIELNIADPEDRARLLSAAES
 LLDE
 406-473_Q9D4W6 -1.226
 >smart|SAM-uniprot|UPI0000F2CDF8|UPI0000F2CDF8/310-377
 PREDICTED: similar to protein with homology to KIAA0790
 [Monodelphis domestica]
 SKMAKPQTVQEFLERILLQEYASTLLLNGYETLEDLKDLKESHLIELNIKNPDRKRLLSAAEN
 LLDD
 310-377_UPI0000F2CDF8 -1.221
 >smart|SAM-uniprot|UPI000155CCD0|UPI000155CCD0/237-304
 PREDICTED: similar to protein with homology to KIAA0790
 [Ornithorhynchus anatinus]
 SKRAKPKTLOEFLERIHLQEYASTLLLNGYESLEDLKYITESHLVELNIVNPEDRMRLLSAAEN
 LLDD
 237-304_UPI000155CCD0 -1.218
 >smart|SAM-uniprot|UPI000155D10F|UPI000155D10F/371-437
 PREDICTED: hypothetical protein [Ornithorhynchus anatinus]
 SHSAGPQDLASFLEQIGCLKYLQVFEQDVDLRIFLTLESIDLKEIGITLFGPKRKMTSAIARW
 HSS
 371-437_UPI000155D10F -1.216
 >smart|SAM-uniprot|UPI0000F1F822|UPI0000F1F822/113-179
 PREDICTED: hypothetical protein [Danio rerio]
 LAELNMDEVQCWFSNIGLEKCLPFIREAEFSGSHIASIDLKTLIEILQVSNLEDRERLLSAYHE
 LHP
 113-179_UPI0000F1F822 -1.215
 >smart|SAM-uniprot|UPI0000500FE5|UPI0000500FE5/238-305 SAM
 domain, SH3 domain and nuclear localization signals, 1 [Rattus
 norvegicus]
 SRRENPQTLQEFLERIHLQEYTS AFLLNGYEALDDLKDIKESHLIELNIANPDRARLLSAAES
 LLDE
 238-305_UPI0000500FE5 -1.212
 >smart|SAM-uniprot|UPI0001560616|UPI0001560616/388-455
 PREDICTED: similar to SAM domain, SH3 domain and nuclear
 localisation signals, 1 [Equus caballus]
 SEREKSKTLOEFLERIQLQEYASTLLLNGYETVEDLKDIKESHLIELNIKNPDRMRLLSAAEN
 LLDE
 388-455_UPI0001560616 -1.199
 >smart|SAM-uniprot|UPI00005A53A0|UPI00005A53A0/389-456
 PREDICTED: similar to SAM-domain protein SAMSN-1 (SAM domain,
 SH3 domain and nuclear localisation signals protein 1) (SH3-SAM

adaptor protein) (Hematopoietic adaptor containing SH3 and SAM domains 1) [Canis lupus familiaris]
SGKEKPCTLQEFLERIHLQEYTSTLLLNGYETLEDLKDIKESHLIELNIKNPDRMRLLSAAEN
LLDE

389-456_UPI00005A53A0 -1.198
>smart|SAM-uniprot|A6NJW8|A6NJW8_HUMAN/406-473 Uncharacterized
protein SAMSN1 [Homo sapiens]
SNSKKSCTLQEFLERIHLQEYTSTLLLNGYETLEDLKDIKESHLIELNIENPDDRRRLSAAEN
FLEE

406-473_A6NJW8 -1.185
>smart|SAM-uniprot|UPI0000434EA0|UPI0000434EA0/1-68 centaurin,
delta 3 [Mus musculus]
MAAPQDLDIAVWLALVHLEQYADTFRRHGLATAGAAQHLGHEELRHLGISATGHRKRILRLLRA
GSAE

1-68_UPI0000434EA0 -1.181
>smart|SAM-uniprot|UPI0000F32B5F|UPI0000F32B5F/1-68 Centaurin-
delta 3 (Cnt-d3) (Arf-GAP, Rho-GAP, ankyrin repeat and
pleckstrin homology domain-containing protein 3). [Bos taurus]
MAAPQDLDIAVWLALVHLEQYADAFRQQGLATAGAARGLGHGELRQLGVNATGHRKRILRLLQA
GTAE

1-68_UPI0000F32B5F -1.179
>smart|SAM-uniprot|Q3ZBW8|Q3ZBW8_BOVIN/237-304 Similar to SAM-
domain protein SAMSN-1 [Bos taurus]
SGVEKPETLQEFLERIHLQEYTSTLLLNGYETLEDLKDIKESHLIELNIKNPDRMRLLSAAEN
LLDE

237-304_Q3ZBW8 -1.177
>smart|SAM-uniprot|UPI0000F2B52E|UPI0000F2B52E/1-68 PREDICTED:
similar to phosphoinositide-binding proteins [Monodelphis
domestica]
MESPRDLDIEVWLAVHLCRYADTFRRHGVSTAAAQSLSQEDLRRMGIWATGHRKRILRLLQG
ASLE

1-68_UPI0000F2B52E -1.175
>smart|SAM-uniprot|A0C082|A0C082_PARTE/14-78 Chromosome
undetermined scaffold_14, whole genome shotgun sequence
[Paramecium tetraurelia]
PKWKWLDDVTIWLKFFIGLQDLDVNFRQNSVDGALLSTLDDNDLKEMGIAESSLKKLVQWIKI
G

14-78_A0C082 -1.162
>smart|SAM-uniprot|UPI0000519A8B|UPI0000519A8B/407-473
PREDICTED: similar to Phosphatidylinositol 3-kinase regulatory
subunit alpha (PI3-kinase p85-subunit alpha) (PtdIns-3-kinase
p85-alpha) (PI3K) [Apis mellifera]

VSEWTSLNVVEWMAALNLYRYADVFHKSDIKGSDLPLFLDRDKLMNMGIKDDFHQKTILACIEEL
 CQT
 407-473_UPI0000519A8B -1.141
 >smart|SAM-uniprot|UPI00005C14DE|UPI00005C14DE/4-70 PREDICTED:
 similar to phosphatidylcholine:ceramide
 cholinephosphotransferase 1 isoform 2 [Bos taurus]
 VVYWSPEKVADWLQENAMPEYCEPLEHFTGRDLINLTOEDFTKPPLCLVTSDNGQRLLHMIETL
 KME
 4-70_UPI00005C14DE -1.097
 >smart|SAM-uniprot|UPI0000D56B70|UPI0000D56B70/25-94 PREDICTED:
 similar to CG30476-PA [Tribolium castaneum]
 VYSWTVVVDVQKWLRRHCTDYYALYVDKFIQNDVTGRTLLRINDNSLLRLGITDVDHREAIWREI
 LKLRLK
 25-94_UPI0000D56B70 -1.089
 >smart|SAM-uniprot|Q7PIK6|Q7PIK6_ANOGA/822-887
 ENSANGP00000024928 [Anopheles gambiae]
 VEEWSNEQVCQWLLGIGLDHHIPAFMQHSVEGGALLQLDKPDFKILNVGGDDKKLLKRNIKELR
 RL
 822-887_Q7PIK6 -1.086
 >smart|SAM-uniprot|Q4RG97|Q4RG97_TETNG/909-974 Chromosome 12
 SCAF15104, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 MTMLDMNISQFLKSLGLDHLRDIFEREQITLDVLADMHEELKEIGINAYGHRHKLIKGVVERLL
 GG
 909-974_Q4RG97 -1.018
 >smart|SAM-uniprot|Q9V4E5|Q9V4E5_DROME/1028-1095 CG1511-PC,
 isoform C [Drosophila melanogaster]
 RGQNIFISTDLWLEHIKMSRYCHHFKEANLINAQOISRLTAQQLSDMGITLVGHQKKILHQARO
 LDTI
 1028-1095_Q9V4E5 -0.998
 >smart|SAM-uniprot|A0D4Q1|A0D4Q1_PARTE/14-80 Chromosome
 undetermined scaffold_38, whole genome shotgun sequence
 [Paramecium tetraurelia]
 PRRWTQCDVESWLEFIGLQDLSDFQRNAIDGACLEVNLDDLNELGISSNVKKKKILQWIQNG
 FIE
 14-80_A0D4Q1 -0.981
 >smart|SAM-uniprot|Q7PRV9|Q7PRV9_ANOGA/965-1032
 ENSANGP00000007421 [Anopheles gambiae]
 GNPGIFISTDLWLEGIKMSRYSQHFKEAGLVTAQQLSRLTAQQLSDMGITLVGHQKKILHQARO
 IDTI
 965-1032_Q7PRV9 -0.980

>smart|SAM-uniprot|Q4H3M5|Q4H3M5_CIOIN/900-967 Ephrin receptor
 [Ciona intestinalis]
 NSIQDVSTLDEWLDVMVKLGRYRRSFHNNGINDLESLAHISESELDRLGIA
 ASPSHRTRLQGGINT
 LRQH
 900-967_Q4H3M5 -0.970
 >smart|SAM-uniprot|Q0JDT6|Q0JDT6_ORYSJ/152-215 Os04g0378200
 protein [Oryza sativa]
 SQPKADSPLDIFLKSLGLEKYSITFQAEEVDMAALRHMTSDLKALGIPMGPRKKIMLA
 ESRA
 152-215_Q0JDT6 -0.956
 >smart|SAM-uniprot|Q5BT66|Q5BT66_SCHJA/12-81 SJCHGC02625 protein
 [Schistosoma japonicum]
 VLNWTPEDVVKWLHKYASPVIAKYSELFANSINGMCLRLMTDEWLLRLGIADQSDRSAL
 MSHI
 YRMRLK
 12-81_Q5BT66 -0.948
 >smart|SAM-uniprot|A0JMNO|A0JMNO_DANRE/4-70 Zgc:154039 [Danio
 rerio]
 VAAWSEDEVSLWLTDQGLQEYSEALRKYDGLALLNLTTEDFKRTPLSRVTSDGGRL
 LLEKIETL
 KIE
 4-70_A0JMNO -0.946
 >smart|SAM-uniprot|Q16S46|Q16S46_AEDAE/91-148 Eph receptor
 tyrosine kinase [Aedes aegypti]
 HSRIDGRTDQWLESIKMSRYSQHFKEAGLVTAQQISRLTAQQQLSDMGITLVGHQKKIL
 91-148_Q16S46 -0.942
 >smart|SAM-uniprot|UPI0000F20BB6|UPI0000F20BB6/949-1014
 PREDICTED: similar to TRF1-interacting ankyrin-related ADP-
 ribose polymerase [Danio rerio]
 ITILDMDNISQFLKSLGLEHLRDIFEREQITLDVLADMGEELKEIGINAYGHRHK
 LIKGIERLL
 GG
 949-1014_UPI0000F20BB6 -0.934
 >smart|SAM-uniprot|UPI0000ECB6F9|UPI0000ECB6F9/663-719 KS5
 protein [Gallus gallus]
 ISEWTTQVCHWLMGMNMEQYITEFTAMNIDGQQLMQLSDKLKALGVSSQNDRSTI
 663-719_UPI0000ECB6F9 -0.931
 >smart|SAM-uniprot|UPI0000D93040|UPI0000D93040/874-939
 PREDICTED: similar to tankyrase-like protein [Monodelphis
 domestica]
 VPGIDFSINQFVRNLGLEHLIDIFEREQITLDVLVEMGHKELKEIGINAYGHRHK
 IIKGVERLI
 SG
 874-939_UPI0000D93040 -0.921
 >smart|SAM-uniprot|UPI00006A0F4F|UPI00006A0F4F/113-178 Tankyrase
 2 (EC 2.4.2.30) (TANK2) (Tankyrase II) (TNKS-2) (TRF1-

interacting ankyrin-related ADP-ribose polymerase 2) (Tankyrase-like protein) (Tankyrase-related protein). [Xenopus tropicalis] VPGVELSINOFLQNLGLEHLIDIFEREQITLDVLIEMGHKEFKEIGINAYGHRHKIIKGVERLISG
 113-178_UPI00006A0F4F -0.916
 >smart|SAM-uniprot|Q4OPU2|Q4OPU2_DROME/16-83 IP04461p
 [Drosophila melanogaster]
 RTRMCGRAVRDWLVLTLMEKYIGKFLERGYDSIERCKLIIIVSDLIMLGVDNPAHRKLLLEGVRF
 LVNA
 16-83_Q4OPU2 -0.905
 >smart|SAM-uniprot|Q4SML6|Q4SML6_TETNG/180-250 Chromosome 18
 SCAF14547, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 VSRWTPEEVLSWLDQLGTWAQLYREPFQQENVNGRLLLMLGEEELLKPPFNVENQAHRRAVLAE
 LDRIKLL
 180-250_Q4SML6 -0.887
 >smart|SAM-uniprot|UPI000065DC62|UPI000065DC62/180-250 Homolog
 of Homo sapiens "Apoptosis regulator [Takifugu rubripes]"
 VSRWTPEEVLSWLDQLGPWAHLYREPFQQENVNGRLLLMLGEEELLKPPYNIGNQAHQRRAVLAELNRIKLL
 180-250_UPI000065DC62 -0.883
 >smart|SAM-uniprot|UPI0000DB6E0B|UPI0000DB6E0B/642-708
 PREDICTED: similar to Bicaudal C CG4824-PA, isoform A [Apis
 mellifera]
 ITSNRYDDLTSMLETSVGLEKYIRLFITSHEVDMATFPSLTEKDLCEIGITAWGARRKIMLLIAEM
 NKR
 642-708_UPI0000DB6E0B -0.871
 >smart|SAM-uniprot|Q5XGM0|Q5XGM0_XENLA/175-245 LOC495189 protein
 [Xenopus laevis]
 VAKWTAAEVVLWLEQLGPWASHYKEHFLSGNVNGRLLLTLAEEFSKNPYSIENSSHRKAILLE
 LDRVKT
 175-245_Q5XGM0 -0.870
 >smart|SAM-uniprot|UPI000044908D|UPI000044908D/167-237
 PREDICTED: similar to apoptosis regulator [Gallus gallus]
 VAKWTAAEVALWLEQLGPWASLYKERFLMEKVNGRLLLTLTEEDFTKEPYSIENNNHKKAITTE
 LECVKTL
 167-237_UPI000044908D -0.869
 >smart|SAM-uniprot|Q20696|SMSR1_CAEEL/11-80 Sphingomyelin
 synthase-related 1 [Caenorhabditis elegans]
 PNEWRCEDVGNWLKKIGMAKYADLIAMKHVDGKCLLALTDSDLKDPPVSINCLGDIKKILFAIEFLSQK
 11-80_Q20696 -0.868

>smart|SAM-uniprot|Q5CVX5|Q5CVX5_CRYPV/289-355 Ser/Thr protein kinase with MORN repeats at the N-terminus and a sterile alpha motif [Cryptosporidium parvum]
 CLSWNMYQVAYFVKCIGLSKYVKNFILNEIDGCTL PYIGFEELKEMGITEITHIKYIRMSL LL
 LKL
 289-355_Q5CVX5 -0.862
 >smart|SAM_PNT-uniprot|Q5TW03|Q5TW03_ANOGA/48-132
 ENSANGP00000027568 [Anopheles gambiae]
 KNGMQATCTELGISSDPKQWTTAKVHQWLELAMGKYFLPRLDNLAALFPENGAQLAALPLEEFV
 RRIPQGGDKLHGYLDL
 48-132_Q5TW03 -0.856
 >smart|SAM-uniprot|Q9VBP3|Q9VBP3_DROME/886-952 CG4719-PA
 [Drosophila melanogaster]
 PDADTITNVSGFLSSQQLHHHLIELFEREQITLDILAEMGHDDLKQVGVSAYGFRHKILKGIAQL
 RST
 886-952_Q9VBP3 -0.842
 >smart|SAM-uniprot|Q295L9|Q295L9_DROPS/894-960 GA18382-PA
 [Drosophila pseudoobscura]
 PDAESITNVSGFLSSQQLNHHLIELFEREQITLDILAEMGHDDLKQVGVSAYGFRHKILKGIAQL
 RAT
 894-960_Q295L9 -0.828
 >smart|SAM-uniprot|Q7PYH8|Q7PYH8_ANOGA/860-926
 ENSANGP00000018360 [Anopheles gambiae]
 SVPSIESSVSVFLTSIQLLEHLIDLFEREQITMDILAEMGHEDLKQVGVSAYGFRHKILKGIAQL
 RAT
 860-926_Q7PYH8 -0.819
 >smart|SAM-uniprot|UPI000044AACC|UPI000044AACC/371-436
 PREDICTED: similar to Dhd2 protein [Gallus gallus]
 SSSRGAEVVKEILKKLDLSEYCPVFEKEKIDGEALFLCAERNLEEMGIPLGPRMKLLHYISSRR
 EV
 371-436_UPI000044AACC -0.810
 >smart|SAM-uniprot|UPI00005472AC|UPI00005472AC/247-314
 PREDICTED: hypothetical protein [Danio rerio]
 HQKAKPKTLEEVLERIGLTELGSLLSMHGFQSLEDFTGLKESHLDNLNITDPEQRVKILKATEL
 IHDS
 247-314_UPI00005472AC -0.806
 >smart|SAM-uniprot|Q4SBI8|Q4SBI8_TETNG/1-59 Chromosome 15
 SCAF14667, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 VGLWLVSMDMDQYASEFTARGVDGPQLLSLDGEKLKALGVCSHSDRAVLKKKLKEIKKM
 1-59_Q4SBI8 -0.803

>smart|SAM-uniprot|UPI000023EA8E|UPI000023EA8E/160-227
 hypothetical protein FG07068.1 [Gibberella zae]
 LASWTTEMVAQAMLNAGIELSVADRFMENDISGAILITLKFEDLKELSIQSFGIRTKVWHQIQA
 LRDS
 160-227_UPI000023EA8E -0.802
 >smart|SAM-uniprot|Q4H3M4|Q4H3M4_CIOIN/961-1028 Ephrin receptor
 [Ciona intestinalis]
 LRHQVPSSLGVWLDGLGLGNYKENFVSAGYSSLDQVLQMSPSDLASLGIVETNQQNAVMEISS
 IRYK
 961-1028_Q4H3M4 -0.754
 >smart|SAM-uniprot|UPI0000D554FF|UPI0000D554FF/868-934
 PREDICTED: similar to CG4719-PA [Tribolium castaneum]
 DLIGNITNIPSFLASLNLEQLIDIFEREQITLDILAEMNHDDLQIGISAYGFRHKLIKGMERL
 MSS
 868-934_UPI0000D554FF -0.753
 >smart|SAM-uniprot|UPI0000F1EB95|UPI0000F1EB95/88-157 PREDICTED:
 hypothetical protein [Danio rerio]
 VVLWSQQDVCRWLKKHCPhNLTYVEAFSHHAITGRALLRLNGEKLERMGLVQETLRQELLQOV
 LQLQVQ
 88-157_UPI0000F1EB95 -0.751
 >smart|SAM-uniprot|UPI000155CAB3|UPI000155CAB3/86-155 PREDICTED:
 similar to sterile alpha motif domain containing 10
 [Ornithorhynchus anatinus]
 VVLWSQQDVCKWLKKHCPhNLVYVEAFAHHAITGRALLRLNAEKLQRMGISOESQRQEVLQOV
 LQLQVR
 86-155_UPI000155CAB3 -0.744
 >smart|SAM_PNT-uniprot|P29775|ETS4_DROME/260-344 DNA-binding
 protein D-ETS-4 [Drosophila melanogaster]
 KREADAICAELOQISQDPNGWSPAQVHAWLRSTLAQFRLPPADLELHFCENGAAALLSEEFV
 RRLPESGSTLHAQLEI
 260-344_P29775 -0.737
 >smart|SAM-uniprot|Q9NGC7|Q9NGC7_MYAAR/485-551 P73-like protein
 [Mya arenaria]
 QNDMQDNSVSTWLNALGLGAYIDGFHEQONLYSLLQLDDFSLDDLAKMKIGNSHRNKIWKSLLEL
 RNQ
 485-551_Q9NGC7 -0.733
 >smart|SAM-uniprot|UPI0000E48DF4|UPI0000E48DF4/406-472
 PREDICTED: similar to Viral A-type inclusion protein repeat
 containing protein [Strongylocentrotus purpuratus]
 VSKWTERDIIHWLDSIHLGHLAQNFIIDHDITGKELMEIDMSLLDEMIDSSLEREKILSTLYDL
 TNP
 406-472_UPI0000E48DF4 -0.702

>smart|SAM-uniprot|UPI0000EBD589|UPI0000EBD589/74-143 PREDICTED:
 similar to sterile alpha motif domain containing 12 isoform 2
 [Bos taurus]
 VALWTQQDVCKWLKKHCPNQYQIYSESFKQHDITGRALLRLTDKKLERMGIAQENLRQLILQQV
 LQLKVR
 74-143_UPI0000EBD589 -0.692
 >smart|SAM-uniprot|UPI00001CFD37|UPI00001CFD37/74-143 PREDICTED:
 similar to sterile alpha motif domain containing 12 [Rattus
 norvegicus]
 VALWTQQDVCKWLKKHCPNQYHIYSESFKQHDITGRALLRLTDKKLERMGVAQENQRQHILQQV
 LQLKVR
 74-143_UPI00001CFD37 -0.684
 >smart|SAM-uniprot|UPI0000F20A91|UPI0000F20A91/78-147 PREDICTED:
 similar to Sterile alpha motif domain containing 12, partial
 [Danio rerio]
 VALWSQQDVCKWLKKHCPNQHQVYSDSFQHDITGRALMRLDRKLERMGIMQESQRQFILQQV
 LQLRVR
 78-147_UPI0000F20A91 -0.676
 >smart|SAM-uniprot|UPI00005844F3|UPI00005844F3/226-298
 PREDICTED: similar to apoptosis regulator [Strongylocentrotus
 purpuratus]
 LSRWRPADVSQWLSELGAWAADGYSRQFNASEINGVLLEGLOSEGELEKEPFNIINNSLHRQAIL
 IALEKVVKQL
 226-298_UPI00005844F3 -0.651
 >smart|SAM-uniprot|Q9SFU7|Q9SFU7_ARATH/140-203 T1B9.17 protein
 [Arabidopsis thaliana]
 KSQQADSSVDSFLESLGLEKYSTAFQVEEVMDALMHMTDDDLKAMLIPMGPRKKILLALGSKP
 140-203_Q9SFU7 -0.642
 >smart|SAM-uniprot|A7A1P0|A7A1P0_YEAST/17-84 MEK kinase
 [Saccharomyces cerevisiae YJM789]
 EKTNDLPFVQLFLEEIGCTQYLDSTIQCNLVTEEEIKYLDKDILIALGVNKIGDRLKILRKSKS
 FQRD
 17-84_A7A1P0 -0.627
 >smart|SAM-uniprot|UPI0000E4A182|UPI0000E4A182/794-860
 PREDICTED: similar to neurabin [Strongylocentrotus purpuratus]
 VWDWNTTHVSQWLMANNLAQYINDFSANSINGQMLLQOLDGARLKIGVTNTNDKNTFKKKVTEL
 KAC
 794-860_UPI0000E4A182 -0.626
 >smart|SAM-uniprot|A6RE25|A6RE25_AJECA/157-224 Predicted protein
 [Ajellomyces capsulatus]
 VRSWTPTRVAEWMYESGFEDAIVERFIRNDISGSVLLDLQMDDLKELDISSFGIRHRLMNMIQG
 FRAE

157-224_A6RE25 -0.607
>smart|SAM-uniprot|Q2UBM0|Q2UBM0_ASPOR/181-248 Predicted protein
[Aspergillus oryzae]
IRGWMPTQVAHWMHIAGYDDYVIEKFMVNDITGSVLLSLQIDDLKELGIKSFGKRHQLMSSINH
LRTT

181-248_Q2UBM0 -0.601
>smart|SAM-uniprot|A1CE91|A1CE91_ASPLC/188-255 SAM and PH domain
protein [Aspergillus clavatus]
VAKWSPQDVVKWMQTLGFEDGIVEKFFINDISGILQVEDLKELELDIQSFGKRHQLMGAIRO
LRNG

188-255_A1CE91 -0.581
>smart|SAM-uniprot|A0FVA6|A0FVA6_9BURK/17-82 Adenylate/guanylate
cyclase [Burkholderia phymatum]
QRSTMTDDLTTWLAQIGLGAYAAKFISQGIDWDVLSLDSEGDLKELGLTLGDRKRLAKGLAALN
GV

17-82_A0FVA6 -0.578
>smart|SAM-uniprot|Q0CRF1|Q0CRF1_ASPTN/192-259 Predicted protein
[Aspergillus terreus]
IRKWSPSQVAHWMHIAGYDGTVIDQFLINDITGSVLLSLQIDDLKELGIQSFGKRHQIMTSIDH
LRNT

192-259_Q0CRF1 -0.568
>smart|SAM-uniprot|Q9Y1Y3|Q9Y1Y3_9METZ/817-884 Protein tyrosine
kinase [Ephydatia fluviatilis]
KTPLDYQSIEAWLEAIKMDRYVQNFHSQGLKSIQDVLLAEEDLRGMGVAIPGHVNKIMSSVRG
AHQQ

817-884_Q9Y1Y3 -0.565
>smart|SAM-uniprot|Q4WNA7|Q4WNA7_ASPLU/189-256 SAM and PH domain
protein (Boil), putative [Aspergillus fumigatus]
VASWSPQDVVMWMLQLGFEESIVEKFFINDISGAILLEANDLKELELDIQSFGKRHHLNCIRO
LKSS

189-256_Q4WNA7 -0.557
>smart|SAM-uniprot|A1DMA0|A1DMA0_NEOFI/194-261 SAM and PH domain
protein [Neosartorya fischeri]
IRAWSPSQVAHWMFIAGYDDSVIEKFIMNDITGSVLLSLQADDLKELGIKSFGKRHQLMGSIDY
LRNT

194-261_A1DMA0 -0.553
>smart|SAM-uniprot|UPI0000E4719B|UPI0000E4719B/694-761
PREDICTED: similar to cask-interacting protein 2
[Strongylocentrotus purpuratus]
VPGYIPGDVGTWLRLLDLSQYHNTLNTNRYGMEEVFHITWEDLQDIGIDMLGHQKKIMLAVRR
LKDL

694-761_UPI0000E4719B -0.519

>smart|SAM-uniprot|Q8BRB1|Q8BRB1_MOUSE/909-976 Ephrin receptor
 [Mus musculus]
 VDIATFHTTGDWLNGMRTAHCKEIFTGVEYSSCDTIAKISTDDMKVGVTVVGPQKKIISTIKA
 LETQ
 909-976_Q8BRB1 -0.512
 >smart|SAM-uniprot|P29320|EPHA3_HUMAN/908-975 Ephrin type-A
 receptor 3 precursor [Homo sapiens]
 VDITTFRTTGDWLNGVWTAHCKEIFTGVEYSSCDTIAKISTDDMKVGVTVVGPQKKIISSIKA
 LETQ
 908-975_P29320 -0.511
 >smart|SAM-uniprot|UPI000066023D|UPI000066023D/903-970 Homolog
 of Brachydanio rerio "EphA4 protein. [Takifugu rubripes]
 SEVTSVGTVGDWMDGMRTLPCKEAFSGVSYSSCDTLAKTSTEDLKKVGVТИVGPQKKIVSSLKA
 LESH
 903-970_UPI000066023D -0.504
 >smart|SAM-uniprot|UPI00001D04ED|UPI00001D04ED/409-476
 PREDICTED: similar to sterile alpha and TIR motif containing 1
 [Rattus norvegicus]
 VASWKEAEVQTWLQQIGFSQYCENFRDQQVGDLLLRLTDEELQTDLGMKSSITRKRFFRELTE
 LKTF
 409-476_UPI00001D04ED -0.503
 >smart|SAM-uniprot|Q4SB53|Q4SB53_TETNG/748-851 Chromosome
 undetermined SCAF14677, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 IHQWGTEEVGAWLDFLCLTEYKDIFTGHDVRGAELLHLERRDLKVPTPRLLPQWNLHFLSLEPG
 PSRLGGAGLTVSLLL
 748-851_Q4SB53 -0.498
 >smart|SAM-uniprot|Q1L8W1|Q1L8W1_DANRE/636-703 Novel protein
 similar to vertebrate EPH receptor A3 [Danio rerio]
 TDITSFHTMGDWLESARTLPCKDAFSGVSYSSCDTLAKTSAEDFKVGVТИVGPQKKIVSSLTT
 LETH
 636-703_Q1L8W1 -0.496
 >smart|SAM-uniprot|UPI000155F20A|UPI000155F20A/402-469
 PREDICTED: similar to sterile alpha and TIR motif containing
 protein 1 [Equus caballus]
 VASWKEAEVQTWLQQIGFSQYCOSFREQQVGDLLLRLTEEEELQTDLGMKGITRKRFFRELTE
 LKTF
 402-469_UPI000155F20A -0.493
 >smart|SAM-uniprot|UPI00005A1DDA|UPI00005A1DDA/409-476
 PREDICTED: similar to sterile alpha and TIR motif containing 1
 [Canis lupus familiaris]

VASWKEAEVQTWLQQIGFSHYCESFREQQVDGDLLLRLTEEEELQTDLGMKSGITRKRFFRELTE
 LKTF
 409-476_UPI00005A1DDA -0.492
 >smart|SAM-uniprot|Q6GP30|Q6GP30_XENLA/12-78 MGC80721 protein
 [Xenopus laevis]
 VSCWSPEALIEYFRDHNLRDCEKLVKKQGITGRRFLEMTENEIQKFPKLAVPMLIKIQQHNMKK
 DEK
 12-78_Q6GP30 -0.491
 >smart|SAM-uniprot|UPI0000D9E230|UPI0000D9E230/598-665
 PREDICTED: similar to sterile alpha and TIR motif containing 1
 [Macaca mulatta]
 VPSWKEAEVQTWLQQIGFSKYCESFREQQVDGDLLLRLTEEEELQTDLGMKSGITRKRFFRELTE
 LRTF
 598-665_UPI0000D9E230 -0.488
 >smart|SAM-uniprot|UPI0000DB748B|UPI0000DB748B/1786-1851
 PREDICTED: similar to Spinophilin CG16757-PA [Apis mellifera]
 VTDWSKEQVCQWLSGIGLERYASRFLETGINGGNLLRLESRDLKAFGICGEEKSHLKRLKELR
 AQ
 1786-1851_UPI0000DB748B -0.487
 >smart|SAM-uniprot|UPI0000D5741F|UPI0000D5741F/587-654
 PREDICTED: similar to l(3)mbt-like 3 isoform a [Tribolium
 castaneum]
 PLRWSVVNVVSEFVSTLPGLEHSSDHFRNSEIDGDAFLSLNQRDLIDILHVKGPAIKLYNVIVL
 LRRK
 587-654_UPI0000D5741F -0.478
 >smart|SAM-uniprot|Q9VR43|Q9VR43_DROME/77-142 CG15625-PA
 [Drosophila melanogaster]
 FEWDDMDIRRWINGYGYPOYMNTFRVNMITGRKLLLLDASALCAMNIKNFDHIRHISYGIRMLF
 HF
 77-142_Q9VR43 -0.470
 >smart|SAM-uniprot|UPI000065D140|UPI000065D140/453-520 Homolog
 of Homo sapiens "Sterile alpha and TIR motif containing
 protein 1 isoform a [Takifugu rubripes]
 VPNWKTCEVQTWLQQVGFSAFCNRFQELQVDGDLLLNTDQDLSADLGMTAGLTRKRFLRDLRV
 LKTY
 453-520_UPI000065D140 -0.457
 >smart|SAM-uniprot|Q4SYZ5|Q4SYZ5_TETNG/464-531 Chromosome 10
 SCAF11883, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 VPNWKTCEVQTWLQQVGFSAFCNRFQELQVDGDLLLSITDQDLSADLGMTAAALTRKRFLRDLRV
 LKTY
 464-531_Q4SYZ5 -0.445

>smart|SAM-uniprot|Q4RA58|Q4RA58_TETNG/3-55 Chromosome
 undetermined SCAF24437, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 DDSLQQYTASFQRQQVDGDKLLKMSHQELLSLGVMRVGHQELILEACDLLCAL
 3-55_Q4RA58 -0.441
 >smart|SAM-uniprot|Q01JD5|Q01JD5_ORYSA/279-344 OSIGBa0130P02.4
 protein [Oryza sativa]
 VMPEVPPTVTGLNSLGLEKYVFLFHAEEDMAALSQMGDSLKEIGVPMGPRKKILQAVAPYS
 KR
 279-344_Q01JD5 -0.438
 >smart|SAM-uniprot|Q00Z16|Q00Z16_OSTTA/199-266 Protein kinase
 family protein [Ostreococcus tauri]
 VDAWTVQDVSEWLRSIELEELVERFAKAKINGYELLRLTEKDLRESLHLERNLERVRAIRAINV
 LRAS
 199-266_Q00Z16 -0.433
 >smart|SAM_PNT-uniprot|Q4S5P4|Q4S5P4_TETNG/492-574 Chromosome 9
 SCAF14729, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LKDVTACKLLSIAADPLDWSCVQVKWLLWTEHLYRLPQVSTMFQELTGRDLCMTEADFRQR
 SSQFGDVLYAHLDIWR
 492-574_Q4S5P4 -0.419
 >smart|SAM_PNT-uniprot|UPI0000D55E79|UPI0000D55E79/305-387
 PREDICTED: similar to CG5583-PA [Tribolium castaneum]
 RKDVDTTCAALGISPDPTQWSAADVLSWLQWTSRQFGLTEPVDPDHWMNGPSLAALSEEDFTRR
 APQGGMILHAQLEIWK
 305-387_UPI0000D55E79 -0.414
 >smart|SAM-uniprot|UPI0000D55DD1|UPI0000D55DD1/895-964
 PREDICTED: similar to CG4393-PA [Tribolium castaneum]
 QNSLNADSIEKWLTQLGMQDYVGLFMSNGFDDVNFLVSKVIEDCDLSEMGITSELERQIIIDAV
 KQLPLK
 895-964_UPI0000D55DD1 -0.412
 >smart|SAM-uniprot|Q5TVD8|Q5TVD8_ANOGA/37-101 ENSANGP00000029329
 [Anopheles gambiae]
 TSLPHMELGEWLQRDLAEGDKFRSFNGVEELLFLSES迪KLGIRNNNAHRARISSLVALRE
 K
 37-101_Q5TVD8 -0.408
 >smart|SAM_PNT-uniprot|UPI0000D91D5E|UPI0000D91D5E/126-208
 PREDICTED: similar to Ets transcription factor PDEF [Monodelphis
 domestica]
 LKDIETACKLLNITADPNPGNVQKWLWTEHQYRLPPVGKAFQELGGKELCAMSEEQFRQR
 SPLCGDILHAHLDIWK
 126-208_UPI0000D91D5E -0.405

>smart|SAM-uniprot|UPI0000EBC729|UPI0000EBC729/33-99 PREDICTED:
 similar to KIAA2004 protein [Bos taurus]
 TDDWTKEDVNQWLESHKIEQKHRAILTAQDVSGAILKYLTQDHLVAMGITFGPAIQIEQLFKEL
 LET
 33-99_UPI0000EBC729 -0.375
 >smart|SAM-uniprot|P18160|KYK1_DICDI/905-972 Non-receptor
 tyrosine kinase spore lysis A [Dictyostelium discoideum]
 FKNWAPNDVAIWLESFNYGQYRKNRDNNISGRHLEGITHAMLKNDLGIEPYGHREDIINRLNR
 MIQI
 905-972_P18160 -0.374
 >smart|SAM-uniprot|Q4JFH3|Q4JFH3_CAEEL/1169-1235 Temporarily
 assigned gene name protein 137, isoform b [Caenorhabditis
 elegans]
 IPYWTSEEVCAWLSSIGMSEYGSTFRKNDIQGSELMHLERSDIMDIGITKIGHVKRLQSAIVDL
 RAO
 1169-1235_Q4JFH3 -0.333
 >smart|SAM-uniprot|UPI000065EE58|UPI000065EE58/3-55 Homolog of
 Homo sapiens "membrane associated guanylate kinase
 interacting protein-like 1 [Takifugu rubripes]
 DDSLQQYISSFQRQQVDGDKLLKMSHQELLSLGVMRVGHQELILEAVDLLCAL
 3-55_UPI000065EE58 -0.332
 >smart|SAM-uniprot|UPI0000D66E0F|UPI0000D66E0F/397-463
 PREDICTED: hypothetical protein [Mus musculus]
 HLTWSPERVAEWISDLGFPQYKECFTENFINGQKLIHVNCNSLPQMGITDFEDMKAISYHTRVL
 LGI
 397-463_UPI0000D66E0F -0.321
 >smart|SAM-uniprot|UPI000065CC80|UPI000065CC80/264-330 FRANK1
 protein (GASZ). [Takifugu rubripes]
 ECVKKLDEIQLLLHGLDLGYLTDIMTENDITWSHLLSMEKEDLEKIGVTDPGDQQKVMNALQQM
 TLD
 264-330_UPI000065CC80 -0.319
 >smart|SAM-uniprot|UPI0000D9BD12|UPI0000D9BD12/535-598
 PREDICTED: similar to CG3108-PA isoform 2 [Macaca mulatta]
 HLNWDPEKVAEWISQLGFPQYKECFITNFISGRKLIHVNCNSLPQMGITNFEDMKTEVMDLGET
 535-598_UPI0000D9BD12 -0.315
 >smart|SAM-uniprot|UPI00006112B0|UPI00006112B0/13-79 PREDICTED:
 hypothetical protein [Gallus gallus]
 VLAWGTAEVAEWVSGLGFPQYQECFRANGITGRRLILVNCSSLPAMGITDFKHMQEISRHVREL
 LGI
 13-79_UPI00006112B0 -0.311

>smart|SAM-uniprot|UPI00004A4D8E|UPI00004A4D8E/518-584
 PREDICTED: similar to Muscle M-line assembly protein unc-89
 (Uncoordinated protein 89) [Canis lupus familiaris]
 YLKWSPEKVAEWISELGFHQYKECFTTNFVCGRKLIHVNCNSLPQMGITDFEDMKVISRHTRAL
 LGI
 518-584_UPI00004A4D8E -0.303
 >smart|SAM-uniprot|UPI000069E2B4|UPI000069E2B4/55-121 SAM domain
 and HD domain-containing protein 1 (Dendritic cell-derived IFNG-
 induced protein) (DCIP) (Monocyte protein 5) (MOP-5). [Xenopus
 tropicalis]
 YREWDVEEVCLFLGSQGLGELEGVFRENKIKGRILEYLTESHLIDLQISSVALRLDLLHCLRML
 CQN
 55-121_UPI000069E2B4 -0.284
 >smart|SAM-uniprot|Q7QIZ0|Q7QIZ0_ANOGA/495-559
 ENSANGP00000007551 [Anopheles gambiae]
 GHTWVASEWLPDIGLPQYGESFMHSLVDARMLDTLSKKELEYLGVTRKFHQASIVHGIHVLRIM
 M
 495-559_Q7QIZ0 -0.280
 >smart|SAM-uniprot|Q75DM8|Q75DM8_ASHGO/1-67 ABL011Cp
 [Eremothecium gossypii]
 MATVGASAEQLLQDINCEGYLGAFRKYEISSVDQLQHLDHEILEQVGIRSIGDRIRILNKSRTL
 QAG
 1-67_Q75DM8 -0.274
 >smart|SAM-uniprot|UPI0000D55A44|UPI0000D55A44/1441-1506
 PREDICTED: similar to CG16757-PA [Tribolium castaneum]
 ISEWSKDQVSQWLIGIGMENHIPKFVELDVNGQALLLLTSADFKILGIASDDKSYLKRIKELK
 IQ
 1441-1506_UPI0000D55A44 -0.265
 >smart|SAM_PNT-uniprot|Q4H3K3|Q4H3K3_CIOIN/202-287 Transcription
 factor protein [Ciona intestinalis]
 SDHHDTKELEVFPSELSSWTNSHVVQWLTWVVEHEYKLQKVDCSRFKGITGIDLSKMTVKDFKN
 LTSTEKDADVFFSHLQ
 202-287_Q4H3K3 -0.260
 >smart|SAM-uniprot|Q6DD51|CSKI2_XENLA/535-601 Caskin-2 [Xenopus
 laevis]
 PQQIPVDLWDWLSQLGLPEYHKQLENGYESLSTVTELTwEGlQEIGIHLGHQKKLLLGVKRL
 LDL
 535-601_Q6DD51 -0.242
 >smart|SAM-uniprot|UPI0000EBC36D|UPI0000EBC36D/328-395
 PREDICTED: hypothetical protein [Bos taurus]
 TENWSEDDVSNWLCAEGLEDLVIDFKMNNIDGRELLNLTKESLADDLKIGFSQWPEIIFEKVVE
 KKQI

328-395_UPI0000EBC36D -0.240
 >smart|SAM-uniprot|Q4SUQ5|Q4SUQ5_TETNG/632-699 Chromosome
 undetermined SCAF13844, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LPEFIPSDLGEWLSAIGLSQYHKKLENGYDSISIVKDLTWEDLQEIGITKLGHQKKLMLAVKK
 LCDI
 632-699_Q4SUQ5 -0.238
 >smart|SAM-uniprot|UPI00006A0F24|UPI00006A0F24/536-603 Caskin-2.
 [Xenopus tropicalis]
 LPQQTPVLDWLSQLGLPEYHKQLSENGYESLSTVTELTwEGLQEIGIHLGHQKKLLLGVKR
 LTDL
 536-603_UPI00006A0F24 -0.233
 >smart|SAM-uniprot|A2AAI8|A2AAI8_MOUSE/554-621 Cask-interacting
 protein 2 [Mus musculus]
 LPNYIPVDLLEWLCAALGLPQYHKQLVSSGYDSMGLVADLTWEELQEIGVNKLGHQKKLMLGVKR
 LAEL
 554-621_A2AAI8 -0.231
 >smart|SAM-uniprot|Q4TAH3|Q4TAH3_TETNG/388-455 Chromosome
 undetermined SCAF7326, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LPNYIPPDNLGEWLSAIGLPQYQORLCDNGYDSITIVKDITWEDLQEIGITKLGHQKKLMLAVKR
 LCDL
 388-455_Q4TAH3 -0.230
 >smart|SAM-uniprot|UPI0000140BCB|UPI0000140BCB/555-622 Caskin-2.
 [Homo sapiens]
 LPSYIPTDLLEWLCAALGLPQYHKQLVSSGYDSMGLVADLTWEELQEIGVNKLGHQKKLMLGVKR
 LAEL
 555-622_UPI0000140BCB -0.229
 >smart|SAM-uniprot|UPI0000F2091C|UPI0000F2091C/702-769
 PREDICTED: hypothetical protein [Danio rerio]
 LPDYIPSDIGKWL SAIGLPQYQKKLAENGYDSISIVQDITWEDLQEIGITKLGHQKKLMLAVKR
 LSDV
 702-769_UPI0000F2091C -0.228
 >smart|SAM-uniprot|Q4T9Q1|Q4T9Q1_TETNG/159-219 Chromosome
 undetermined SCAF7511, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 VSSWTTQQVCHWLRLGMSMDRYVPEFSARDVDGQELVQMDGKKLKVNKRQLHLKEEEHKQGQ
 159-219_Q4T9Q1 -0.226
 >smart|SAM-uniprot|UPI0000EBDED2|UPI0000EBDED2/555-622
 PREDICTED: similar to cask-interacting protein 2 [Bos taurus]
 LPNYIPADLLEWLCAALGLPQYHKQLVSSGYDSMGLVADLTWEELQEIGVNKLGHQKKLMLGVKR
 LAEL

555-622_UPI0000EBDED2 -0.225
 >smart|SAM-uniprot|UPI00004A5EB5|UPI00004A5EB5/555-622
 PREDICTED: similar to cask-interacting protein 2 [Canis lupus familiaris]
 LPNYIPADLREWLALGLPQYHKQLVSSGYDSMGLVADLTWEELQEIGVNKLGHQKKLMLGVKR
 LAEL
 555-622_UPI00004A5EB5 -0.223
 >smart|SAM-uniprot|UPI0000E49437|UPI0000E49437/1934-2000
 PREDICTED: hypothetical protein [Strongylocentrotus purpuratus]
 PWTWGIAEVVQFLAETGEGSCAECFCRQNINGQKLMQLSKEQLVKLTGMKVAPSLKIYEQIVKL
 RSQ
 1934-2000_UPI0000E49437 -0.222
 >smart|SAM-uniprot|UPI000069DF96|UPI000069DF96/514-581 RNA-binding protein with serine-rich domain 1 (SR-related protein LDC2). [Xenopus tropicalis]
 LPEYKPANLSLWLTMIGLIQYYKVLVNGYENIDFITDITWEDLQEIGITKLGHQKKLMLAVKK
 LAEI
 514-581_UPI000069DF96 -0.221
 >smart|SAM-uniprot|UPI000065DCB1|UPI000065DCB1/483-550 Homolog of Homo sapiens "CASK interacting protein 1 [Takifugu rubripes]"
 LPEQKPANLGEWLSAIGLSQYHQVLVQNGYENIEFITDITWEDLQEIGITKLGHQKKLMLAVKR
 LAEM
 483-550_UPI000065DCB1 -0.220
 >smart|SAM-uniprot|Q4S059|Q4S059_TETNG/551-618 Chromosome undetermined SCAF14784, whole genome shotgun sequence [Tetraodon nigroviridis]
 LPEHKPANLGEWLSAIGLSQYHQVLVQNGYENIEFITDITWEDLQEIGITKLGHQKKLMLAVKR
 LAEM
 551-618_Q4S059 -0.219
 >smart|SAM-uniprot|UPI0000F1D271|UPI0000F1D271/549-616
 PREDICTED: similar to cask-interacting protein 1 [Danio rerio]
 IPQEKPSSLAEWLSAIGLNQYYQTLVQNGYDNMDFISDITLEDLKEIGFTKLGHQKKLMLAIKR
 LSES
 549-616_UPI0000F1D271 -0.218
 >smart|SAM-uniprot|Q4S703|Q4S703_TETNG/1550-1617 Chromosome 14 SCAF14723, whole genome shotgun sequence [Tetraodon nigroviridis]
 QPSGNISSVSDWLTSIGLPMYASPLAAAGVHTLSGVSSLTESGAREAGVRDERHLRRLVGEARL
 VGTE
 1550-1617_Q4S703 -0.216

>smart|SAM-uniprot|UPI0000F1DA4B|UPI0000F1DA4B/534-601
 PREDICTED: similar to cask-interacting protein 1 [Danio rerio]
 LPDQKPASLGEWLAMIGLSQYHQVLVQNGYENIDFITDITWEDLQEIGITKLGHQKKLMLAVKK
 LAEI
 534-601_UPI0000F1DA4B -0.215
 >smart|SAM-uniprot|UPI00006CD8A5|UPI00006CD8A5/320-387 protein
 kinase domain [Tetrahymena thermophila]
 LREWNTGHICKWLDSIGLGEFSPSFESNQICGQSICMLTEQELEQSIGMKALGKRKIFLKYQEM
 LKKY
 320-387_UPI00006CD8A5 -0.211
 >smart|SAM-uniprot|UPI0000DC1A24|UPI0000DC1A24/472-539 Caskin-1
 (CASK-interacting protein 1). [Rattus norvegicus]
 SASEGKANLAVWLSMIGLAQYYKVLVDNGYENIDFITDITWEDLQEIGITKLGHQKKLMLAVRK
 LAEL
 472-539_UPI0000DC1A24 -0.210
 >smart|SAM-uniprot|Q4SW70|Q4SW70_TETNG/33-99 Chromosome 9
 SCAF13686, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 FPQWGEEETCQYLKEGFGEWQEAFREREIDGARLRSILRDADLEQMGIRCPDERLRLLRSLRKL
 WQI
 33-99_Q4SW70 -0.196
 >smart|SAM-uniprot|UPI0001554AA4|UPI0001554AA4/12-78 PREDICTED:
 similar to SLP-76 [Ornithorhynchus anatinus]
 VLGWDPEDLAEYFKKLNFKDCEKAVRKFHIDGERFLNLSDNDIQKFPKLRVPILSKLSQDINKN
 EER
 12-78_UPI0001554AA4 -0.195
 >smart|SAM-uniprot|UPI0000141AD9|UPI0000141AD9/12-78 Lymphocyte
 cytosolic protein 2 (SH2 domain-containing leukocyte protein of
 76 kDa) (SLP-76 tyrosine phosphoprotein) (SLP76). [Homo sapiens]
 VLGWDPDSLADYFKKLNKDCEKAVKKYHIDGARFLNLTDIQKFPKLRVPILSKLSQEINKN
 EER
 12-78_UPI0000141AD9 -0.191
 >smart|SAM-uniprot|Q08DS6|Q08DS6_BOVIN/12-78 Similar to
 Lymphocyte cytosolic protein 2 [Bos taurus]
 VLSWDPNLTADYFKKLNKDCEKVVVKYDIDGPRFLNLAEENDIQKFPKLRVPILSKLSQEINKN
 EER
 12-78_Q08DS6 -0.187
 >smart|SAM_PNT-uniprot|A2BIB5|A2BIB5_MOUSE/31-115 Ets homologous
 factor [Mus musculus]
 CNVSSGFFGSQWHEIHPQYWTKYQVWEWLQHLLDTNQLDASCIPFQEFDISGEHLCSMSLQEFT
 RAAGSAGQLLYSNLQH
 31-115_A2BIB5 -0.182

>smart|SAM_PNT-uniprot|UPI00015613AC|UPI00015613AC/31-115
 PREDICTED: similar to Ets homologous factor [Equus caballus]
 CNVSSGFQWHEIHPQYWTQVWEWLQHLLDTNQLDASCIPFQEFDINGEHLCGMSLQEFT
 RAAGTAGQLLYSNLQH
 31-115_UPI00015613AC -0.181
 >smart|SAM-uniprot|Q4T305|Q4T305_TETNG/64-130 Chromosome 18
 SCAF10147, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LPDHKPANLADWLAHLGLSQYYQVLVQNGYENIDFISDISLEDLQEIGITKLAVEAVEHKIKEE
 DAE
 64-130_Q4T305 -0.180
 >smart|SAM-uniprot|Q5ZMC3|WSDU1_CHICK/330-396 WD repeat, SAM and
 U-box domain-containing protein 1 [Gallus gallus]
 AENWSEDDVSAWLCAQGFAELVGLFKANNIDGKELVNLTRESLIHELKMSLWLRSKILQKIEEL
 RMK
 330-396_Q5ZMC3 -0.173
 >smart|SAM-uniprot|Q8C4H2|SAMD3_MOUSE/1-66 Sterile alpha motif
 domain-containing protein 3 [Mus musculus]
 METWSVDQVCKWLVEKNLGEVPRFQEEEVSGATLLALNDRMVQQLVKKIGHQAVLMDFIKKYK
 QG
 1-66_Q8C4H2 -0.170
 >smart|SAM-uniprot|Q1AMZ6|Q1AMZ6_MYTTR/520-586 P63/p73-like
 protein [Mytilus trossulus]
 LMANGDHSISNWLTTLGLSAYIDNFHQQNLFTMEQLDDFTVEDLQKMRIGTSHRNKIWKALVEF
 HSE
 520-586_Q1AMZ6 -0.161
 >smart|SAM-uniprot|Q27937|Q27937_LOLFO/451-517 P53 tumor
 suppressor homolog [Loligo forbesi]
 KCEPTENTIAQWLTKLGLQAYIDNFQQKGLHNMFQLDEFTLEDLQSMRIGTGHRNKIWKSLDY
 RRL
 451-517_Q27937 -0.156
 >smart|SAM_PNT-uniprot|UPI0000F21820|UPI0000F21820/35-120
 PREDICTED: hypothetical protein [Danio rerio]
 SMMTSNFTSRLWTQDPHPQNWTKFQVWEWLQQTLDMHQIDATSIPFQNFQDFLDGRQLCNMSFQDF
 TRAAGSVGSILFQSLS
 35-120_UPI0000F21820 -0.154
 >smart|SAM_PNT-uniprot|UPI00006A0047|UPI00006A0047/1-74 ETS
 homologous factor (hEHF) (ETS domain-containing transcription
 factor) (Epithelium-specific Ets transcription factor 3) (ESE-
 3). [Xenopus tropicalis]
 WHDIHPQYWSKFQVWEWLQQLVDTNQLDANCIPFHDFTIAGEHLCNMTLPDFTQAAGSAGHILY
 NNLQNLKWND

1-74_UPI00006A0047 -0.151
>smart|SAM_PNT-uniprot|UPI00006A0045|UPI00006A0045/33-117 ETS homologous factor (hEHF) (ETS domain-containing transcription factor) (Epithelium-specific Ets transcription factor 3) (ESE-3). [Xenopus tropicalis]
ALPTTGYFGGQWHDIHPOYWSKFOVWEWLQQLVDTNQOLDANCIPFHDFDIAGEHLCNMTPDFT
QAAGSAGHILYNNLQN

33-117_UPI00006A0045 -0.150
>smart|SAM_PNT-uniprot|UPI00005A35DB|UPI00005A35DB/78-162 PREDICTED: similar to E74-like factor 5 ESE-2a [Canis lupus familiaris]
A FEHQ TACDSY WTSV HPEY WTKR HVWE WLQF CCDQ YKLD ANCIS FCH FN VSG LQL CSMT QEEF V
EA AGIC GEYL YFIL QN

78-162_UPI00005A35DB -0.149
>smart|SAM_PNT-uniprot|UPI00006A004B|UPI00006A004B/1-74 ETS-related transcription factor Elf-5 (E74-like factor 5) (Epithelium-specific Ets transcription factor 2) (ESE-2) (Epithelium- restricted ESE-1-related Ets factor). [Xenopus tropicalis]
W TSLL PEHWT KDH VCEWLQ YCCDSY KLD ANCIP FSH FN VSG LQL CNMTR DDF TEAAGL CGYY LY
S LLQE IRT QA

1-74_UPI00006A004B -0.146
>smart|SAM-uniprot|Q0JRM9|Q0JRM9_9VEST/481-547 P73-like protein [Haliothis tuberculata]
G QTLVDNSVASWLTSIGLSAYIDNFHEKNFLNMFQLDEF SLEDLSAMKIGTSHRNKIWKSLVEY
K QA

481-547_Q0JRM9 -0.141
>smart|SAM_PNT-uniprot|Q8VDK3|ELF5_MOUSE/35-119 ETS-related transcription factor Elf-5 [Mus musculus]
A FEHQ TACDSY WTSV HPEY WTKR HVWE WLQF CCDQ YKLD ANCIS FCH FN ISGL QLC SMT QEEFI
EA AGIC GEYL YFIL QN

35-119_Q8VDK3 -0.138
>smart|SAM_PNT-uniprot|Q5TRG1|Q5TRG1_ANOGA/64-148
ENSANGP00000027450 [Anopheles gambiae]
T SAKKVQYGADGLPVDP RDWTRANVWTWLINLAQSEG LDISPELAQKFPMNGKALCLMSLDMYL
SRVPIGGKMLYRDFRV

64-148_Q5TRG1 -0.128
>smart|SAM_PNT-uniprot|Q703F5|Q703F5_TRICA/16-102 ETS activity modulator [Tribolium castaneum]
GP TERWQRLSF DDEDNL PKD PRQWTREHVAQWINLV TQQH GLPE VPSS RFL MNGK ALCL MSL GM
FL SRVPLGGKLLYKDF

16-102_Q703F5 -0.124

>smart|SAM-uniprot|Q4P965|VTS1_ USTMA/608-672 Protein VTS1
 [Ustilago maydis]
 ADLSTLEDVPAWLQLRLHKYTPNFETSHWKEVMVMMGDKQLEDKGVAALGARRKMLKTFELVRK
 K
 608-672_Q4P965 -0.123
 >smart|SAM_PNT-uniprot|Q9VB52|Q9VB52_DROME/1278-1367 CG5954-PA,
 isoform A [Drosophila melanogaster]
 LPNYRLWQLNSAFKLDDVRTNPLHWTSWDVCEYIERALDSTDIAKVIFEQDIDGRALLMLGRKE
 LDTYLKLKVGPALKY
 1278-1367_Q9VB52 -0.121
 >smart|SAM-uniprot|UPI0000EFCDA7|UPI0000EFCDA7/60-128
 hypothetical protein An17g01280 [Aspergillus niger]
 VRSWDEKQVIAWLHSINCGQYEALFKAPANNFNGNNLIECDQKILQEMGIKKIGDRVRIFVAIK
 QLRNK
 60-128_UPI0000EFCDA7 -0.119
 >smart|SAM-uniprot|UPI00001AF4B3|UPI00001AF4B3/237-304 WD
 repeat, SAM and U-box domain-containing protein 1. [Homo
 sapiens]
 TEDWSEEDVSTWLCAQDLKDLVGIFKMNNIDGKELLNLTKESLADDLKIESLGLRSKVLRKIEE
 LRTK
 237-304_UPI00001AF4B3 -0.117
 >smart|SAM-uniprot|Q55FH8|Q55FH8_DICDI/446-512 Kelch repeat-
 containing protein [Dictyostelium discoideum]
 LTESQKKDVKYFLNSIGMSKYVNRFIEEEIDSTVFHLLTENHLKEIGIKSLGERLSIIKGVREY
 SLS
 446-512_Q55FH8 -0.116
 >smart|SAM-uniprot|UPI000056694E|UPI000056694E/327-394 WD
 repeat, SAM and U-box domain containing 1 [Mus musculus]
 TEEWSEEDVSVWLRAQGLEDLVGIFRANNIDGKELLHLTKESSLAGDLKIESLGLRSKVLRSIEE
 LRAK
 327-394_UPI000056694E -0.113
 >smart|SAM-uniprot|UPI0000EE0148|UPI0000EE0148/329-396
 PREDICTED: similar to WD repeat, sterile alpha motif and U-box
 domain containing 1 [Ornithorhynchus anatinus]
 IENWSEEVEVWLCAQGLKDLGVFKMNNIDGKELLSLTKESSLNDLKIESLGLRSKILRKIEE
 LNIK
 329-396_UPI0000EE0148 -0.112
 >smart|SAM_PNT-uniprot|Q4H3K2|Q4H3K2_CIOIN/201-285 Transcription
 factor protein [Ciona intestinalis]
 TSEENKKNKRVLPADPLIWSGPCHTAWLEVVVNEYGLHHIDLTKFSSVTGTELCRMNVEDLTR
 YTTRYNSEVLVQHLKF
 201-285_Q4H3K2 -0.111

>smart|SAM_PNT-uniprot|Q4H3K4|Q4H3K4_CIOIN/333-417 Transcription factor protein [Ciona intestinalis]
 QGNGEAETKRVIVPADPLVWTEEHVQEWEWTITEYNLSVDRSSFSNINGRALCEMTKEFHR
 ITSAKNADVFMSHLNY
 333-417_Q4H3K4 -0.110
 >smart|SAM_PNT-uniprot|UPI00005A0990|UPI00005A0990/114-198
 PREDICTED: similar to Friend leukemia integration 1
 transcription factor (Fli-1 proto-oncogene) (ERGB transcription factor) isoform 7 [Canis lupus familiaris]
 PPNNMTTNERVIVPADPTLWTQEHVRQWLEWAIKEYGLMEIDTSFFQNMDGKELCKMNKEDFLR
 ATSLYNTEVLLSHLSY
 114-198_UPI00005A0990 -0.109
 >smart|SAM_PNT-uniprot|Q4SAF7|Q4SAF7_TETNG/27-113 Chromosome 13
 SCAF14688, whole genome shotgun sequence [Tetraodon nigroviridis]
 PMVVSGYFSPLWSGCESQPQFWTKFQVWEWLQVLDMHQMDSNFPFQNFNVDGHQLCNLSHQD
 YVCAAGSVGSILFQRV
 27-113_Q4SAF7 -0.108
 >smart|SAM_PNT-uniprot|Q4T1Z3|Q4T1Z3_TETNG/112-195 Chromosome undetermined SCAF10411, whole genome shotgun sequence [Tetraodon nigroviridis]
 PSSSATEEKRVIVPADPEVWTQDHVRQWLDWAIKEYVLEEVDVVLFQALDGKALCKMTKDDMMR
 LTSAYNADILLSHLN
 112-195_Q4T1Z3 -0.107
 >smart|SAM_PNT-uniprot|UPI00005A5398|UPI00005A5398/115-199
 PREDICTED: similar to v-ets erythroblastosis virus E26 oncogene like isoform 1 isoform 1 [Canis lupus familiaris]
 PPNNMTTNERVIVPADPTLWSTDHVRQWLEWAVKEYGLPDVDILLFQNIIDGKELCKMTKDDFQR
 LTPSYNADILLSHLN
 115-199_UPI00005A5398 -0.106
 >smart|SAM-uniprot|Q5FVN8|WSDU1_RAT/329-396 WD repeat, SAM and U-box domain-containing protein 1 [Rattus norvegicus]
 IEDWSEEDVSKWLRAQGLEDLVDIFRTNNIDGKELLHLTKESSLAGDMKIESLGLRSKVLRSIDE
 LRTR
 329-396_Q5FVN8 -0.105
 >smart|SAM_PNT-uniprot|Q4Z8P1|Q4Z8P1_RAT/114-198 Friend leukemia integration 1 [Rattus norvegicus]
 PPNNMTTNERVIVPADPTLWTQEHVRQWLEWAIKEYGLMEIDTSFFQNMDGKELCKMSKEDFLR
 ATSLYNTEVLLSHLSY
 114-198_Q4Z8P1 -0.104
 >smart|SAM_PNT-uniprot|Q1L8J3|Q1L8J3_DANRE/137-200 Friend leukemia integration 1 [Danio rerio]

PPNMTTNERVIVPADPSLWSPDHVRQWLDWAIKEYGLQEIDTAMFHSTDGKELCKMSKDDFLR
 137-200_Q1L8J3 -0.103
 >smart|SAM_PNT-uniprot|A2T6E0|A2T6E0_PAPHA/1-77 ELF5 [Papio
 hamadryas]
 CDSYWTSHPEYWTKRHVWEWLQFCCDQYKLDINCISFCHFNISGLQLCSMTQEEFVEAGLCG
 EYLYFILQNIRTO
 1-77_A2T6E0 -0.102
 >smart|SAM_PNT-uniprot|Q6GNZ9|Q6GNZ9_XENLA/124-208 MGC80765
 protein [Xenopus laevis]
 PPNMTTNERVIVPADPTLWSTDHVRQWLEWAIKEYGLPDVDVLLFQNIIDGKELCKMTKEDFHR
 LTPSYNADILLSHLHY
 124-208_Q6GNZ9 -0.100
 >smart|SAM_PNT-uniprot|P41157|FLI1_XENLA/113-197 Retroviral
 integration site protein Fli-1 homolog [Xenopus laevis]
 PPNMTTNERVIVPADPALWSQDHVRQWLEWAIKEYGLVEIDCSLFQNIIDGKELCKMSKEDFLR
 STSIYNTEVLLSHLNY
 113-197_P41157 -0.099
 >smart|SAM_PNT-uniprot|UPI0000DC187D|UPI0000DC187D/111-195 v-ets
 erythroblastosis virus E26 oncogene like [Rattus norvegicus]
 PPNMTTNERVIVPADPTLWSTDHVRQWLEWAVKEYGLLDVDVLLFQNIIDGKELCKMTKDDFQR
 LTPSYNADILLSHLHY
 111-195_UPI0000DC187D -0.098
 >smart|SAM_PNT-uniprot|UPI000155479C|UPI000155479C/35-119
 PREDICTED: similar to E74-like factor 5 ESE-2a [Ornithorhynchus
 anatinus]
 TYDHQTACDSYWTSHPEYWTKRHVWEWLQFCCDQYKLDANCISFCHFNINGLQLCSMTQEEFI
 DAAGICGEYLYFILQ
 35-119_UPI000155479C -0.097
 >smart|SAM-uniprot|UPI0000D55DEA|UPI0000D55DEA/136-206
 PREDICTED: similar to Stromal interaction molecule homolog
 precursor [Tribolium castaneum]
 VHNWTVEQTTDWLVSSVDPQYVPSFISHKVTGANLPRLAANNVNYLNHLGIKDPIHKQKIVLK
 AMDVVLF
 136-206_UPI0000D55DEA -0.094
 >smart|SAM_PNT-uniprot|UPI000065F421|UPI000065F421/53-135
 Homolog of Homo sapiens "ESE-1a" [Takifugu rubripes]
 ISNLSGQSYRQIEPQYWTVDNVLEWISDHVESNKFDAHSLSLALCSMDGYSLCQMAQDQMIEAF
 GSQLGSHLYHSLQEHK
 53-135_UPI000065F421 -0.093
 >smart|SAM-uniprot|UPI0000DB6D0E|UPI0000DB6D0E/6-74 PREDICTED:
 similar to SMSr CG32380-PA, isoform A [Apis mellifera]

IIEWTQKDVAKWLSESGHEKFVDLFFDQEIDGKVLLTLKEDDLKLISMNIKKIGDIKRLYISIK
QLQRA

6-74_UPI0000DB6D0E -0.090

>smart|SAM-uniprot|UPI000065CB47|UPI000065CB47/326-393 Homolog
of *Brachydanio rerio* "Similar to U5 snRNP-specific 40 kDa
protein (HPrp8-binding). [Takifugu rubripes]

VSDWSEQDVSSWLLEEGLEGLVDKFRANNIDGTELLNLTKETLASLHIESVGLRNKLLRKVEE
LKNE

326-393_UPI000065CB47 -0.081

>smart|SAM-uniprot|Q7KUZ1|Q7KUZ1_DROME/117-188 CG9126-PC,
isoform C [*Drosophila melanogaster*]
VHNWTIEQTTDWLAQSVQLPQYVDLFKLHKVTGAALPRLAVNNLQYVGNVLGIKDPIHKQKISL
KAMDVVLF

117-188_Q7KUZ1 -0.080

>smart|SAM_PNT-uniprot|Q4TA74|Q4TA74_TETNG/63-151 Chromosome
undetermined SCAF7420, whole genome shotgun sequence [*Tetraodon nigroviridis*]
VSNTAYMSCLSGQSYRQTEPQYWTENVLEWISDHVESNKIDAHSLSLACSMNGYSLCQMAQD
QMMEVFGPQLGSHLYQ

63-151_Q4TA74 -0.074

>smart|SAM-uniprot|UPI000069E562|UPI000069E562/333-400 WD
repeat, SAM and U-box domain-containing protein 1. [*Xenopus tropicalis*]
VEAWSEEVATWLSQEGLNEVEHIFKANNIDGKELLHLTKDSLNLKIESLGLRNKLVRRIED
LKSS

333-400_UPI000069E562 -0.073

>smart|SAM-uniprot|A0AUS0|WSDU1_DANRE/344-411 WD repeat, SAM and
U-box domain-containing protein 1 [*Danio rerio*]
VSEWSEEVLAWLREEGLEAVTDAFKSNNIDGEELLSLSKETLSSDLHIESLGLRSKVMKKIEE
LKMV

344-411_A0AUS0 -0.065

>smart|SAM-uniprot|UPI0000F2BF15|UPI0000F2BF15/1120-1186
PREDICTED: similar to protein phosphatase 1, regulatory subunit
9B, spinophilin, [*Monodelphis domestica*]
ANHWTSQQVGQWLESLNLDQYTAEFAARQVDGQQLLQLDGGKLKSLGLSNSHDRALVKRKLKEL
VAA

1120-1186_UPI0000F2BF15 -0.054

>smart|SAM-uniprot|Q8J218|Q8J218_EMENI/60-126 MAPKK kinase
[*Emericella nidulans*]
VRSWDENQVISWLHSINCQQYEPLFRANNFNGNNLIECDQKILQEMGIKKIGDRVRI
FVAIKQL
RNK

60-126_Q8J218 -0.053

>smart|SAM-uniprot|Q4WTX3|Q4WTX3_ASPFU/61-127 MAP kinase kinase kinase SteC [Aspergillus fumigatus]
 VRSWDEKQVINWLHSINCGQYESLFRANNFNGNNLIECDQKILQEMGIKKVGDVRIFVAIKKL
 RNK
 61-127_Q4WTX3 -0.048
 >smart|SAM-uniprot|UPI000023D1D8|UPI000023D1D8/60-126
 hypothetical protein FG05484.1 [Gibberella zae]
 VKNWDEDQVCEYLRTVKCGEYEKIFRKNHINGENLLEMKDVLKEMGVKEVGDRVRLFLSIKKL
 RTR
 60-126_UPI000023D1D8 -0.047
 >smart|SAM-uniprot|A1C9Y4|A1C9Y4_ASPLC/60-126 MAP kinase kinase kinase Ste11/SteC [Aspergillus clavatus]
 VRSWDEKQVIawlHSINCGQYESLFRANNFNGDNLIECDQKILQEMGVKKIGDRVRIFVAIKKL
 RNK
 60-126_A1C9Y4 -0.046
 >smart|SAM-uniprot|Q0MQZ5|Q0MQZ5_PENMA/20-86 STE11-like protein [Penicillium marneffei]
 VRSWDEKKVADWLHSIRCAQYEPLFKANNFNGDNLIECDQKILQEMGVKKIGDRVRIFVAIKQL
 RNK
 20-86_Q0MQZ5 -0.019
 >smart|SAM_PNT-uniprot|UPI000065EC68|UPI000065EC68/57-145
 Homolog of Gallus gallus "Transforming protein p54/c-ets-1.
 [Takifugu rubripes]
 GETFTGAVLLYTLIYICIYADPTLWSEWEVNYWLDWCQSEFGLHSLDSELRCLOGRDLCALDKDA
 FLALISDCTAGEILWE
 57-145_UPI000065EC68 -0.017
 >smart|SAM-uniprot|UPI0000EBD8A0|UPI0000EBD8A0/68-135 PREDICTED:
 similar to centaurin delta 2, partial [Bos taurus]
 EAGDGALSVAEWLRALHLEQYTRLFEQHGLVWATDCQGLSDTRLVDMGMVLPGHRRRILAGLLR
 AHTP
 68-135_UPI0000EBD8A0 -0.013
 >smart|SAM-uniprot|UPI0000F2D237|UPI0000F2D237/3-70 PREDICTED:
 similar to centaurin, delta 2, [Monodelphis domestica]
 EEVGAMLPVAEWLRALHLEQYAGHFEQYGLVWAADCRLSDGQLLNMGVTMPGHRRRILSGLQR
 AYAT
 3-70_UPI0000F2D237 -0.012
 >smart|SAM-uniprot|UPI0000F1D36C|UPI0000F1D36C/396-462
 PREDICTED: hypothetical protein [Danio rerio]
 LSEWTNQQVCHWLMGMNMEQYIAEFTAKGVDGQHLLSMDSSKLKELGVSSQRDRATIKRRLKDM
 KKA
 396-462_UPI0000F1D36C -0.004

>smart|SAM-uniprot|Q7JQG5|Q7JQG5_DROME/321-388 LD09801p
 [Drosophila melanogaster]
 LPHFVPGSIEEWLQLLRLEEYIQPLLEQNYKTVRDVTQVTWEDLEDIGIVKLGHQKKILLAIKR
 VKDI
 321-388_Q7JQG5 -0.003
 >smart|SAM-uniprot|UPI000155CEFD|UPI000155CEFD/657-723
 PREDICTED: similar to KS5 protein [Ornithorhynchus anatinus]
 ISEWTTQQVCHWLMGMNMEQYITEFTAKNIDGQQLMLLDSDLKTLGVTSQNDRSSIKKKIKDM
 KKT
 657-723_UPI000155CEFD -0.001
 >smart|SAM-uniprot|Q4RNS2|Q4RNS2_TETNG/682-748 Chromosome 2
 SCAF15010, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 VSEWTSVQVCHWLMGMNMDQYTQEFTAKGVDGQQLLHLDSDRLKALGVSSQSDRSTLKKKMKE
 RKA
 682-748_Q4RNS2 0.000
 >smart|SAM-uniprot|Q4S122|Q4S122_TETNG/426-492 Chromosome 1
 SCAF14770, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PPYPMDSSISSLRLGCAGCLDYFTAQGLTNIYQIENYNMEDLSRLKIPSEFQHIIWKGIMEH
 RQA
 426-492_Q4S122 0.007
 >smart|SAM-uniprot|UPI0000E489DC|UPI0000E489DC/1262-1325
 PREDICTED: similar to inositol polyphosphate phosphatase-like 1
 [Strongylocentrotus purpuratus]
 RPITITEWSNLGLM0YKQOLYDNGWDNVAFLDTITDQDLISARVESIKHRRHMLESTCIMAM
 1262-1325_UPI0000E489DC 0.012
 >smart|SAM-uniprot|Q7QIQ9|Q7QIQ9_ANOGA/433-498
 ENSANGP00000014993 [Anopheles gambiae]
 NEENDSLAILRFLAAFKLEDYYPIFKNEIDMETLMMLETDVKSLGLPLGPYRRLCNAIQERR
 EA
 433-498_Q7QIQ9 0.018
 >smart|SAM-uniprot|UPI000069F923|UPI000069F923/1076-1143 SAM and
 SH3 domain-containing protein 1 (Proline-glutamate repeat-
 containing protein). [Xenopus tropicalis]
 VSPRHISSTDWLVSVGLPMYARPLAEAGITTLSTLPSLSLTHLQKAGIKEERHINKLLSATRI
 LKSM
 1076-1143_UPI000069F923 0.023
 >smart|SAM-uniprot|Q7Q0R1|Q7Q0R1_ANOGA/849-915
 ENSANGP00000006183 [Anopheles gambiae]
 PESWEVKDVATFLTINDCAVHAEQFVQKDIDGKRLLELSKDDIITLLNLKVGPA
 LKIFDLIQQL
 KCK

849-915_Q7Q0R1 0.034
>smart|SAM-uniprot|Q9HFR3|Q9HFR3_PNECA/7-73 Mekk [Pneumocystis carinii]
VRLWSEEEVGEWLESNNFGDYMDIFKENNINGDILLECNAAVLKELGVKKLGDRIRLSVCIKGL
REK
7-73_Q9HFR3 0.035
>smart|SAM-uniprot|UPI000065EAA8|UPI000065EAA8/695-761 Homolog
of Gallus gallus "KS5 protein. [Takifugu rubripes]
VSEWTNQQVCHWLMGMNMDQYTQEFTAKAVDGQQLLHLDSDLKALGVSSQSDRSTIKKKLKDL
RKA
695-761_UPI000065EAA8 0.040
>smart|SAM-uniprot|Q29L50|Q29L50_DROPS/1201-1267 GA14249-PA
[Drosophila pseudoobscura]
PDSWNVYDVSQFLRVNDCTAYCDTFSRSKIDGKRLQLTKDDIMPLLGGMKVGPA
LIISDLITQL
KCK
1201-1267_Q29L50 0.044
>smart|SAM-uniprot|UPI0000D56EC4|UPI0000D56EC4/647-713
PREDICTED: similar to CG4824-PA, isoform A [Tribolium castaneum]
IANTTWSDLPSLLSALGLDRYISL FIRHEVDLPTFTLSDKDLMTIGVTA
FGSRRKMLLAISELNKR
647-713_UPI0000D56EC4 0.047
>smart|SAM-uniprot|UPI0000ECB15E|UPI0000ECB15E/521-587 tumor
protein p73-like [Gallus gallus]
PPYPTDCSIVSFLARLGSSCVDYFTTQGLTTIYQIEHY
SMDDL VSLKIPEQFRHAIWK
GILDHRQL
521-587_UPI0000ECB15E 0.053
>smart|SAM-uniprot|P59808|SASH1_MOUSE/1157-1224 SAM and SH3
domain-containing protein 1 [Mus musculus]
LSPGCVASMSDWLISIGLPMYTSTLSDAGFSTLSQVPSLSH
SCLQEAGITEERHIRKLITA
FKLP
1157-1224_P59808 0.054
>smart|SAM-uniprot|Q7Z8J5|Q7Z8J5_YARLI/189-255 MAP kinase kinase
kinase [Yarrowia lipolytica]
VTSWDKTVAEWLRSLKCAEYIPMFEQNEICGEV
LLEIDQKLLKDMGVVKVGDRV
RINVAIKDL
RSQ
189-255_Q7Z8J5 0.057
>smart|SAM-uniprot|UPI0000DA1BC8|UPI0000DA1BC8/1162-1229
PREDICTED: similar to putative adapter and scaffold protein 1
[Rattus norvegicus]
LSPGCVTSVDWLISIGLPMYTSTLSDAGFSALSQVPSLSHTC
LQEAGITEERHIRKL
VTA
FKLP
1162-1229_UPI0000DA1BC8 0.058

>smart|SAM-uniprot|UPI000155FD3D|UPI000155FD3D/541-607
 PREDICTED: similar to TA p63 alpha isoform 1 [Equus caballus]
 PPYPTDCSLVSFLARLGCSSCLDYFTTQGLTTIYQIEHYSMDLASKIPEQFRHAIWKGILDH
 RQL
 541-607_UPI000155FD3D 0.071
 >smart|SAM-uniprot|UPI0000D55CC9|UPI0000D55CC9/803-869
 PREDICTED: similar to CG16975-PB, isoform B [Tribolium
 castaneum]
 PDEWNIYDVAQFLRVNDANCDSFSKQKVDGKTLMNLSKEDILEYTGGKVGPSLKIFDLIQQL
 KIK
 803-869_UPI0000D55CC9 0.076
 >smart|SAM-uniprot|UPI0000DB6BAA|UPI0000DB6BAA/807-873
 PREDICTED: similar to CG16975-PB, isoform B [Apis mellifera]
 PSEWNVFDVAQFLRVNDCATYCDNFSKRKIDGKTLLTKDQIIDLTGFVGPSLKIYDLIQQL
 KIK
 807-873_UPI0000DB6BAA 0.077
 >smart|SAM-uniprot|Q5XJ13|Q5XJ13_DANRE/640-706 Zgc:85964 [Danio
 rerio]
 GTITDEDELSGILRKLSLEKYQPIFEEQEVDMEAFLTLTDGDLQELGIRTDGPRQQILAAISEL
 NAG
 640-706_Q5XJ13 0.078
 >smart|SAM-uniprot|UPI0000E7FE83|UPI0000E7FE83/773-839
 PREDICTED: similar to sterile alpha motif domain containing 6
 [Gallus gallus]
 GTITEDDELTGILKKLSLEKYQPIFEEQEVDMEAFLTLTDGDLKELGIKTDSRQQILAAISEL
 NAG
 773-839_UPI0000E7FE83 0.080
 >smart|SAM-uniprot|UPI00006A1675|UPI00006A1675/613-682 Ankyrin
 repeat and SAM domain-containing protein 1 (Odin). [Xenopus
 tropicalis]
 GSYVPEQSVAQWLKSIGLLQYESKLVLNGFDDVRFLADTMMDQDLLDIGITDLQHRHQILRLA
 QSLPKP
 613-682_UPI00006A1675 0.081
 >smart|SAM-uniprot|UPI0000F1EC7A|UPI0000F1EC7A/336-407
 PREDICTED: hypothetical protein [Danio rerio]
 IHSWTEEHVHFWMEQIFEGDGELOQQYANVFRQNHTGQRLLLSESDMRDLGVTSKGHIIHLKT
 EIEKLTHD
 336-407_UPI0000F1EC7A 0.097
 >smart|SAM-uniprot|UPI0000E814A6|UPI0000E814A6/506-572
 PREDICTED: similar to P53-like transcription factor [Gallus
 gallus]

PPYNPDPSLVSFLTGLCPNCIDYFTSQGLQNIYHLQNLNSIEDLGALKIPEQYRMIIWRLQEL
 KQS
 506-572_UPI0000E814A6 0.101
 >smart|SAM-uniprot|Q4RNR9|Q4RNR9_TETNG/336-410 Chromosome 2
 SCAF15010, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 IYTWTEEHVYFWMQQIFGEGEGPSGMQVYADLFKENHITGRLLLLTEGDMRDMGVKSKGHVMH
 LQAAIHGEIEK
 336-410_Q4RNR9 0.103
 >smart|SAM_PNT-uniprot|UPI0000548281|UPI0000548281/33-122
 PREDICTED: hypothetical protein [Danio rerio]
 HLSEITFNSNSTLGSWDQVNPMQWSRQNVLIEWIGFHVVEESRFDAGLLNLNYCTMDGLTLCATSK
 EALMSMFGPELGNLLH
 33-122_UPI0000548281 0.104
 >smart|SAM-uniprot|UPI0000EBED4F|UPI0000EBED4F/453-519
 PREDICTED: similar to P73 alpha protein [Bos taurus]
 PPYHADSSLVSLTGLCPNCIEYFTSQGLQNIYHLQNLTIEDLGALKIPDQYRMTIWRLQDL
 KQS
 453-519_UPI0000EBED4F 0.115
 >smart|SAM-uniprot|UPI0000364E65|UPI0000364E65/328-394 Homolog
 of Brachydanio rerio "Similar to U5 snRNP-specific 40 kDa
 protein (HPrp8-binding). [Takifugu rubripes]
 FSDWSEEDVQAWLRQEGLQDLVGTFKSNNIDGAELSRLNKDTAAELGIESVGLRCRLMRKIEAL
 KAE
 328-394_UPI0000364E65 0.116
 >smart|SAM-uniprot|UPI0000DA2DE3|UPI0000DA2DE3/807-876
 PREDICTED: similar to cajalin 2 isoform a [Rattus norvegicus]
 RPRCPVQTVGQWLESIGLPQYENHLTANGFDNVQFMGSNVMEDQDLLEIGILNSGHRQRILQAI
 QLLPKM
 807-876_UPI0000DA2DE3 0.122
 >smart|SAM-uniprot|UPI000060934C|UPI000060934C/806-875
 PREDICTED: ankyrin repeat and sterile alpha motif domain
 containing 1B isoform 2 [Mus musculus]
 RPRCPVQTVGQWLESIGLPQYENHLMANGFDQVFMGSNVMEDQDLLEIGILNSGHRQRILQAI
 QLLPKM
 806-875_UPI000060934C 0.126
 >smart|SAM-uniprot|Q6UNX2|Q6UNX2_DANRE/491-557 Transcription
 factor TAp73 alpha [Danio rerio]
 PPYNPDPSLVSFLTSVGCQNCIDYFTSQGLQSVYHLQTLTMEDLGALKIPEQFRLAIWRGLQEM
 KQG
 491-557_Q6UNX2 0.127

>smart|SAM-uniprot|Q9NRX7|Q9NRX7_HUMAN/33-102 Putative 47 kDa protein [Homo sapiens]
 GPRCPVQTVGQWLESIGLPQYENHLMANGFDNVQFMGSNVMEDQDLLEIGILNSGHRQRILQAIQLLPKM
 33-102_Q9NRX7 0.130
 >smart|SAM-uniprot|UPI0000E46C34|UPI0000E46C34/1048-1117
 PREDICTED: similar to Ankyrin repeat and sterile alpha motif domain containing 1B [Strongylocentrotus purpuratus]
 DGPGGPQSIGAWLDSLNLGQYENMLIANGFDHLDFIGGGIIEEQLMDIGIVEEIHRKMILEATRCLPIM
 1048-1117_UPI0000E46C34 0.131
 >smart|SAM-uniprot|UPI0000660959|UPI0000660959/33-102 Homolog of Homo sapiens "Ankyrin repeat and SAM domain-containing protein 1 [Takifugu rubripes]"
 CGKALDQPVG EWLEHVGLLQYESKFLNGFDDLRFMGSNVMEDQDLRDVGITDPGHRKKILHAA RSLPKV
 33-102_UPI0000660959 0.134
 >smart|SAM-uniprot|Q6FXI1|Q6FXI1_CANCA/4-62 Candida glabrata strain CBS138 chromosome B complete sequence [Candida glabrata]
 IEFLERAGCSQYADLFEEYSLTDINDLMHIDRDILLEIGVKKLGDRIRILKESKKLQEP
 4-62_Q6FXI1 0.137
 >smart|SAM-uniprot|UPI0000E7F896|UPI0000E7F896/803-872
 PREDICTED: similar to cajalin 2 [Gallus gallus]
 RPRCPVQTVGQLENIGLPQYENHLLANGFDNVQFMGSNVMEDQDLLEIGILNSGHRQRILQAIQLLPKM
 803-872_UPI0000E7F896 0.139
 >smart|SAM-uniprot|UPI0000F2C069|UPI0000F2C069/716-785
 PREDICTED: similar to KIAA0229 [Monodelphis domestica]
 GSRMLEQSVGEWLESIGLQQYESKLLLNGFDDVHFLGSNVMEDQDLREIGISDPQHRRKLLQAA RSLPKV
 716-785_UPI0000F2C069 0.140
 >smart|SAM-uniprot|UPI0000D9AC74|UPI0000D9AC74/773-842
 PREDICTED: similar to ankyrin repeat and sterile alpha motif domain containing 1 [Macaca mulatta]
 GSRTLEQSVGEWLESIGLQQYESKLLLNGFDDVRFLGSNVMEQDLRDIGISDPQHRRKLLQAA RSLPKV
 773-842_UPI0000D9AC74 0.144
 >smart|SAM-uniprot|UPI0001555BA2|UPI0001555BA2/315-384
 PREDICTED: similar to AIDA1C transcript, partial [Ornithorhynchus anatinus]
 GPRPSERSVGEWLGSVGLQQYESKLLLNGFDDVRFLGSNVMEQDLREIGITDPQHRRKLLQAA RSLPKV

315-384_UPI0001555BA2 0.145
 >smart|SAM-uniprot|Q9VMJ2|Q9VMJ2_DROME/84-150 CG13996-PA
 [Drosophila melanogaster]
 VFKWSINDVTDWLRLNFGYYPEYEQTFRNEYIDGHKLLNLDAVALVALNVRNFEHIRHLGRGIRAL
 YRK
 84-150_Q9VMJ2 0.148
 >smart|SAM-uniprot|Q22CV7|Q22CV7_TETTH/73-140 FHA domain protein
 [Tetrahymena thermophila]
 PLIWSQEDVLEWLKFIGLDKYAPQFKENTIDGSCLNTLTSDDLQNHLKIDNAIQRKKILNWISV
 GIQT
 73-140_Q22CV7 0.150
 >smart|SAM-uniprot|Q1LX04|Q1LX04_DANRE/18-84 Novel protein
 similar to vertebrate protein phosphatase 1, regulatory
 (Inhibitor) subunit 9A [Danio rerio]
 VTEWTCQQVSCWLMGLNLEQYVSLFTAKNLDGEQLLKLDSTALKALGIESSSDRALIKKKLKDL
 KVL
 18-84_Q1LX04 0.152
 >smart|SAM-uniprot|UPI000065D045|UPI000065D045/611-680 Homolog
 of Homo sapiens "AIDA-1a [Takifugu rubripes]
 GPRCPMQSVGQWLDSIGLVQYENHLLANGFDNVQFMGSNVVEDQDLLEIGILNSAHRQRLQAI
 RLLPRV
 611-680_UPI000065D045 0.160
 >smart|SAM-uniprot|P28829|BYR2_SCHPO/1-67 Protein kinase byr2
 [Schizosaccharomyces pombe]
 MEYYTSKEVAEWLKSIGLEKYIEQFSQNНИЕGRHLNHTLPLLKDLGIENTAKGKQFLKQRDYL
 REF
 1-67_P28829 0.163
 >smart|SAM-uniprot|UPI0000F210A8|UPI0000F210A8/685-754
 PREDICTED: similar to KIAA0229 [Danio rerio]
 SGRLLEQPVGDWLEHIGLPQYESKLLLNGFDDLRYMGNDVMEEQDLREIGITDPGHRRKILSAA
 RSLPKV
 685-754_UPI0000F210A8 0.167
 >smart|SAM-uniprot|A2AM22|A2AM22_MOUSE/486-552 Transformation
 related protein 73 [Mus musculus]
 PPYHADPSLVSFLTGLGCPNCIECFTSQGLQSIYHLQNLTIEDLGALKVPDQYRMTIWRGLQDL
 KQS
 486-552_A2AM22 0.168
 >smart|SAM-uniprot|UPI00005A0D1F|UPI00005A0D1F/933-999
 PREDICTED: similar to tumor protein p73 [Canis lupus familiaris]
 PPYHADPSLVSFLTGLGCPNCIEYFTSQGLQNIYHLQNLTIEDLGALKIPDQYRMTIWRGLQDL
 KQS
 933-999_UPI00005A0D1F 0.173

>smart|SAM-uniprot|UPI000065D5D3|UPI000065D5D3/496-562 Homolog
 of Barbus barbus "P73. [Takifugu rubripes]
 PPYNPDPSLVSFLSLGCQNFIEYFASQGLQSVYHLQTLSMEDLGAMKIPQSRLAIWRGLQDM
 KQG
 496-562_UPI000065D5D3 0.181
 >smart|SAM-uniprot|Q8J0Z5|Q8J0Z5_CRYNV/94-160 STE11
 [Cryptococcus neoformans var. grubii]
 LGDWGSQELFTFLDIHKCGQYLAIFQENDINGKILLDLDVTALKSMGIDKISERVRLIGGIKDL
 RKR
 94-160_Q8J0Z5 0.186
 >smart|SAM-uniprot|Q4RQJ9|Q4RQJ9_TETNG/342-408 Chromosome 2
 SCAF15004, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 FSDWSEEDVQAWLREEGLQDLIGIFTTHNIDGAELTRLDKDAAELGIESLGLRSRLMRKIEAL
 KAE
 342-408_Q4RQJ9 0.189
 >smart|SAM-uniprot|Q4S837|Q4S837_TETNG/488-554 Chromosome 9
 SCAF14710, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PPYNPDPSLVSFLSLGCQNFIEYFASQGLQSVYHLQTLSMEDLGAMKIPQSRLAIWRALQDL
 KQD
 488-554_Q4S837 0.193
 >smart|SAM-uniprot|UPI0000DB6E48|UPI0000DB6E48/469-536
 PREDICTED: similar to Ect4 CG7915-PB, isoform B, partial [Apis
 mellifera]
 VPLWSTEDVREWVKQIGFAECAQNVESRVDGDLLLQLTEENLKEDIGLANGIRRKRTRELQN
 LKKM
 469-536_UPI0000DB6E48 0.196
 >smart|SAM-uniprot|UPI0000F51D0A|UPI0000F51D0A/4-70 Neurabin-1
 [Rattus norvegicus]
 VHEWSVQQVSHWLVLGLSLDQYVSEFSAQNISGEQLLQLDGNKLKALGMTSSQDRALVKKKLKEM
 KMS
 4-70_UPI0000F51D0A 0.198
 >smart|SAM-uniprot|Q9P6R7|VTS1_SCHPO/592-656 Protein vts1
 [Schizosaccharomyces pombe]
 VGNELPQDIPSWLRLHKTNNLKDTDWALVSLSDLQNRGIMALGARRKLLKSFQEVP
 L
 592-656_Q9P6R7 0.202
 >smart|SAM-uniprot|UPI0000F20BC0|UPI0000F20BC0/8-76 PREDICTED:
 similar to connector enhancer of kinase suppressor of Ras 2,
 [Danio rerio]

VSRWSTSQVVDWMKGLDDCLQQYIKNFEQEKGGEQLLRITHQELEDLGVSRIGHTQELILEAVD
LLCAL

8-76_UPI0000F20BC0 0.211
>smart|SAM-uniprot|UPI000156080B|UPI000156080B/85-155 PREDICTED:
similar to CNKSR family member 3 [Equus caballus]
VLDEKHLQYVEFVRGCGLDDCLQQYVHKFEREKINGEQLLQISHQDLEELGVTRIGHQELVLEA
VDLLCAL

85-155_UPI000156080B 0.212
>smart|SAM-uniprot|UPI0000EBE5C0|UPI0000EBE5C0/1-73 PREDICTED:
similar to connector enhancer of KSR2A [Bos taurus]
MPINTFCETGHKENAKKRGLDDCLQQYIKNFEREKISGDQLLRITHQELEDLGVSRIGHTQELIL
EAVDLLCAL

1-73_UPI0000EBE5C0 0.214
>smart|SAM-uniprot|UPI0000E80130|UPI0000E80130/102-170
PREDICTED: similar to CNKSR family member 3 [Gallus gallus]
VTKWSPKQVVDWTKGGLDDCLQQYVHKFEREKINGEQLLQISHQDLEDMGITRIGHQELVLEAVD
LLCAL

102-170_UPI0000E80130 0.215
>smart|SAM-uniprot|UPI000065CEF8|UPI000065CEF8/8-76 Homolog of
Homo sapiens "Splice Isoform 1 of Connector enhancer of
kinase suppressor of ras 2 [Takifugu rubripes]
VSKWSSGQVVDWMKGLDDCLQQYIKTFEREKVGKDQLLRITHQELEDLGVSRIGHTQELILEAVD
LLCAL

8-76_UPI000065CEF8 0.216
>smart|SAM-uniprot|UPI00005A5BFB|UPI00005A5BFB/8-76 PREDICTED:
similar to connector enhancer of kinase suppressor of Ras 2
isoform 2 [Canis lupus familiaris]
VSKWSPSQVVDWMKGLDDCLQQYIKNFEREKISGDQLLRITHQELEDLGVSRIGHTQELILEAVD
LLCAL

8-76_UPI00005A5BFB 0.217
>smart|SAM-uniprot|Q6GQI9|Q6GQI9_XENLA/8-76 MGC79120 protein
[Xenopus laevis]
VSKWTPSQVVDWMKGLDDCLQQYIKNFEREKIAGDQLLRITHQELEELGVTRIGHQELILEAVD
LLCAL

8-76_Q6GQI9 0.218
>smart|SAM-uniprot|UPI0000E7FAB9|UPI0000E7FAB9/8-76 PREDICTED:
similar to connector enhancer of KSR2A [Gallus gallus]
VSKWSPSQVVDWMAGLDDCLQQYIKNFEREKISGDQLLRITHQELEDLGVSRIGHTQELILEAVD
LLCAL

8-76_UPI0000E7FAB9 0.219
>smart|SAM-uniprot|UPI0000F2C1E1|UPI0000F2C1E1/8-69 PREDICTED:
similar to CNKSR family member 3 [Monodelphis domestica]

LAASSKKGLDDCLQQYVHKFEREKINGEQLLQISHQDLEELGVSRIGHTQELVLEAVDLLCAL
8-69_UPI0000F2C1E1 0.220
>smart|SAM-uniprot|UPI000155E0FD|UPI000155E0FD/984-1050
PREDICTED: similar to PPP1R9A protein isoform 1 [Equus caballus]
VHEWSVQQVSHWLMSLNLEQYVSEFSAQNITGEQLLQLDGNKLKALGMTSSQDRAVVKKKLKEM
KVS
984-1050_UPI000155E0FD 0.221
>smart|SAM-uniprot|UPI0000F208C4|UPI0000F208C4/4-72 PREDICTED:
similar to maguin-like protein, partial [Danio rerio]
ITKWTAAQVVDWIRGLDDSLQQYISNFEREKISGERLLKISHQELEELSVTRVGHQEELILEAVD
LLCAL
4-72_UPI0000F208C4 0.222
>smart|SAM-uniprot|UPI0000DB7389|UPI0000DB7389/6-74 PREDICTED:
similar to connector enhancer of ksr CG6556-PA [Apis mellifera]
VAEWKTEQVCEWLKG LDNSVLPYVHSFTNHAVNGQQLLSLRPEDLEHLGVLKLGHQEIIILEAVE
YLRNF
6-74_UPI0000DB7389 0.223
>smart|SAM-uniprot|UPI000065F984|UPI000065F984/8-76 Homolog of
Homo sapiens "Splice Isoform 1 of Connector enhancer of
kinase suppressor of ras 2 [Takifugu rubripes]
VSKWTTGQVVDWMKGLDDCLQQYVCVFERGGVCGERLLRISHAELEELGVSRIGHTQELILEAVD
LLCAL
8-76_UPI000065F984 0.224
>smart|SAM-uniprot|Q4S8W9|Q4S8W9_TETNG/8-77 Chromosome 7
SCAF14703, whole genome shotgun sequence [Tetraodon
nigroviridis]
VSKWSSGQVVDWMKGLDDCLQQYIKTFEKEKVGGDQLLRITHQELEDLGVSRIGHTQELILEAVG
PTLCRG
8-77_Q4S8W9 0.225
>smart|SAM-uniprot|UPI0000D57459|UPI0000D57459/6-74 PREDICTED:
similar to CG6556-PA [Tribolium castaneum]
VRDWSVDQVTDWLKG LDSIILQYTTFLNNNGVTGHQLNLRADDLDNLGVKSLGHQEELILESVE
HLRFH
6-74_UPI0000D57459 0.226
>smart|SAM-uniprot|UPI000060F8C6|UPI000060F8C6/1007-1073
Neurabin-1 (Neurabin-I) (Neural tissue-specific F-actin-binding
protein I) (Protein phosphatase 1 regulatory subunit 9A).
[Gallus gallus]
VQEWSVQQVSHWLMSLNLEQYVSEFSAQNINGEHLLQLDGSKLKALGMTSSQDRAIVKKKLKEM
KAS
1007-1073_UPI000060F8C6 0.233

>smart|SAM_PNT-uniprot|P51023|PNT2_DROME/166-250 ETS-like
 protein pointed, isoform P2 [Drosophila melanogaster]
 FASWEKEVQKCNITKDPREWTEEHVIYWLWAKNEFSLVSMNLDPFYKMKGRAMVDLGKEKFLA
 ITPPFTGDLWEHLDI
 166-250_P51023 0.234
 >smart|SAM-uniprot|UPI00006A1675|UPI00006A1675/683-751 Ankyrin
 repeat and SAM domain-containing protein 1 (Odin). [Xenopus
 tropicalis]
 PVLSADLSVPSWRLSIALHQYAPNFISSGYSSMETLRNLWELEIVNVLKVNLLGHRKRILASLA
 ERPYE
 683-751_UPI00006A1675 0.238
 >smart|SAM-uniprot|Q4RP02|Q4RP02_TETNG/3-55 Chromosome 10
 SCAF15009, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 DDSLQQYVSNFEREKISGEQLLKITHQDLEELGLARIGHQELVLEAVDLLCAL
 3-55_Q4RP02 0.239
 >smart|SAM-uniprot|Q7PT20|Q7PT20_ANOGA/536-599
 ENSANGP00000021615 [Anopheles gambiae]
 QPNVGMSSIAHWLKSRLHKYVWLFSNLTYDKMLGITEEYLQSLGVTKGARHKLAICIQKLKER
 536-599_Q7PT20 0.242
 >smart|SAM_PNT-uniprot|Q16VF8|Q16VF8_AEDAE/149-232 DNA-binding
 protein elg, putative [Aedes aegypti]
 DTNFKRDQQRLNIPDDPLEWTVAQVKHWIOWAVKIFQQLTSIKLQDWSISGKELCDMDHAEFKQK
 VPSDPGDLFWTHLELL
 149-232_Q16VF8 0.246
 >smart|SAM_PNT-uniprot|UPI0000DA36BA|UPI0000DA36BA/297-380
 PREDICTED: similar to C-ets-2 protein [Rattus norvegicus]
 FSGFHKEQRRLGIPKNPWLWNEQQVCQWLHWATNEFSLVNVNLQRFGMNGQMLCNLGKERFLEL
 APDFVGDLWEHLEQM
 297-380_UPI0000DA36BA 0.248
 >smart|SAM-uniprot|Q16F20|Q16F20_AEDAE/769-836 Sarm1 [Aedes
 aegypti]
 VPLWSTEDVREWVKQIGFNEYHNFFDSKVDGDLLLQLTEDNLREDIGMTNGIRRKFREELQN
 LKRM
 769-836_Q16F20 0.249
 >smart|SAM_PNT-uniprot|UPI0000D55EDE|UPI0000D55EDE/215-300
 PREDICTED: similar to DNA-binding protein D-ETS-6 [Tribolium
 castaneum]
 DSDSDIDNEMVLVPSDPLEWTNTHIKSWLSWCSRKFSLNPKPDFEKFPTTGKELCELTRTDRET
 KAGCERTGTILAKHIA
 215-300_UPI0000D55EDE 0.251

>smart|SAM_PNT-uniprot|UPI0000DB74A2|UPI0000DB74A2/209-293
 PREDICTED: similar to pointed CG17077-PD, isoform D [Apis mellifera]
 FASWEKERVRLNITKDPROWSEAAVAHWLHWAIGEFSLAGVAIQPWQNMTGKQICAMGKESFLA
 RAPAFMGDILWEHLEI
 209-293_UPI0000DB74A2 0.252
 >smart|SAM-uniprot|Q174C4|Q174C4_AEDAE/712-775 Smaug protein [Aedes aegypti]
 SPNVGMSSIAHWLKSRLHLKYVWLFSNLTYEQMMEMTEEYLANLGVTKGARHKLVLCIHKLKER
 712-775_Q174C4 0.253
 >smart|SAM_PNT-uniprot|Q5TRW4|Q5TRW4_ANOGA/27-111
 ENSANGP00000025810 [Anopheles gambiae]
 FASWEKEVQTFKITKDPROWTAEHVLIWLWNWSIKEFSLEGVNKEPFQKMSGRDIVGLGREGFLA
 IAPPFTGDILWEHLEI
 27-111_Q5TRW4 0.254
 >smart|SAM_PNT-uniprot|Q26645|Q26645_STRPU/122-205 ETS homologue [Strongylocentrotus purpuratus]
 FSSFKKEQERLGIPKDPWRSECQVVAWVHWSIKEFSLEGVSINNFRISGRDLCTLPKTDFLSR
 APPFMGDILWEHIDML
 122-205_Q26645 0.255
 >smart|SAM_PNT-uniprot|Q330K7|Q330K7_PARLI/122-205 ETS1 [Paracentrotus lividus]
 FSSFKKEQERLGIPKDPMSGWAECQVVAWVLWAIKEFSLEGVSVNNFRINGRDLCTLPKTDFLSR
 APPFMGDILWEHIDML
 122-205_Q330K7 0.256
 >smart|SAM-uniprot|Q6BRC3|Q6BRC3_DEBHA/10-79 Similar to CA4000|IPF4262 Candida albicans IPF4262 [Debaryomyces hansenii]
 FLQWDPAQVCNYINSNLGDQTSLGDCFLDNNIEGSLLPFITTEHLKEIGVTKLGTRLQIKKAT
 SELIAN
 10-79_Q6BRC3 0.259
 >smart|SAM_PNT-uniprot|UPI00005E967B|UPI00005E967B/53-136
 PREDICTED: hypothetical protein [Monodelphis domestica]
 FSGFTKEQQRLGIPKDPROWTETHVRDWVMWAVNEFSLKGVNFQKFCMSGAALCALGKDCFEL
 APDFVGDIWEHLEIL
 53-136_UPI00005E967B 0.260
 >smart|SAM-uniprot|UPI0000F2D6E7|UPI0000F2D6E7/4-70 PREDICTED:
 similar to ARAP2 [Monodelphis domestica]
 VSEVNPELRELLTNINLEQYLGCFQQSGYHTLKDCAQINDGILQKIGVLPTGHRRRILKQLEMI
 LSR
 4-70_UPI0000F2D6E7 0.265
 >smart|SAM-uniprot|UPI0000D9B349|UPI0000D9B349/4-70 PREDICTED:
 similar to centaurin delta 1 isoform a, partial [Macaca mulatta]

VSEVNVDIKDFLMSINLEQYLLHFHESGFTTVKDCAVINDSLLQKIGISPTGHRRRILKQLQII
 LSK
 4-70_UPI0000D9B349 0.267
 >smart|SAM_PNT-uniprot|P01105|MYBE_AVILE/278-361 p135Gag-Myb-
 Ets-transforming protein [Contains: Transforming protein v-Myb;
 Transforming protein v-Ets] [Avian leukemia virus]
 FSGFAKEQQRLGIPKDPQQWTETHVRDWVMWAVNEFSLKGVDQKFCMNGAALCALGKECFLEL
 APDFVGDILWEHLEIL
 278-361_P01105 0.268
 >smart|SAM_PNT-uniprot|Q8K3Q9|Q8K3Q9_MOUSE/46-129 Ets-1 [Mus
 musculus]
 FSGFTKEQQRLGIPKDPQQWTETHVRDWVMWAVNEFSLKGVDQKFCMNGAALCALGKECFLEL
 APDFVGDILWEHLEIL
 46-129_Q8K3Q9 0.270
 >smart|SAM_PNT-uniprot|P13474|ETS1A_CHICK/53-136 Transforming
 protein p54/c-ets-1 [Gallus gallus]
 FSGFAKEQQRLGIPKDPQQWTETHVRDWVMWAVNEFSLKGVDQKFCMNGAALCALGKECFLEL
 RPDFVGDILWEHLEIL
 53-136_P13474 0.271
 >smart|SAM-uniprot|UPI000155E0D0|UPI000155E0D0/4-70 PREDICTED:
 similar to ARAP2 [Equus caballus]
 VSEVNVDIKDFLMSINLEQYLLHFREFGFNTVKDCAAINDSVLHKIGISPTGHRRRILKQLQII
 LSK
 4-70_UPI000155E0D0 0.272
 >smart|SAM_PNT-uniprot|UPI0000D90565|UPI0000D90565/90-173
 PREDICTED: similar to v-ets erythroblastosis virus E26 oncogene
 homolog 2 (avian), [Monodelphis domestica]
 FSGFTKEQRRLGIPNNPWLTTEEQVCQWLFWATNEFSLMDVNFQKFIMNGQVLCNLGKERFLEL
 APDFVGDILWEHLEQM
 90-173_UPI0000D90565 0.273
 >smart|SAM_PNT-uniprot|O62803|O62803_SHEEP/43-126 Transcription
 factor Ets-2 [Ovis aries]
 FSGFKKEQRRLGIPPKSPWLWTEQQVCQWLLWATNEFSLVDVNLQRFGMTGQVLCNLGKERFLEL
 APDFVGDILWEHLEQM
 43-126_O62803 0.274
 >smart|SAM_PNT-uniprot|UPI0000EDFF36|UPI0000EDFF36/90-173
 PREDICTED: hypothetical protein [Ornithorhynchus anatinus]
 FNGFTKEQCRLGIPNNPWLTTEEQVCQWLFWATNEFSLMDVNFQRFVMNGQVLCNLGKERFLEL
 APDFVGDILWEHLEQM
 90-173_UPI0000EDFF36 0.275
 >smart|SAM-uniprot|UPI000155D1CF|UPI000155D1CF/3-70 PREDICTED:
 similar to ARAP2 [Ornithorhynchus anatinus]

SSPEAHFGIQELLTSINLGQYLPFCFESGYVTIQDCAGINNSVLLKIGVFPTGHRRRILKQLEI
 ISSK
 3-70_UPI000155D1CF 0.276
 >smart|SAM_PNT-uniprot|UPI00006A08C2|UPI00006A08C2/87-170 C-ets-
 2 protein. [Xenopus tropicalis]
 FNGFTKERCRRLGIPGNPWLWDENHVFQWLLWAAKEFSLEENVNFQKFLMNGHELCSLGKERFLAL
 APDFVGDLWEHLEEM
 87-170_UPI00006A08C2 0.277
 >smart|SAM-uniprot|UPI0000E8047B|UPI0000E8047B/15-82 PREDICTED:
 similar to ARAP2 [Gallus gallus]
 SSTGSDIDIEEFLVNINLGQYLPNFKEHGYNVVTDCVGINNSALQQMGVLPTGHRRRILKQLDT
 ALSK
 15-82_UPI0000E8047B 0.281
 >smart|SAM-uniprot|UPI0000F2C069|UPI0000F2C069/790-858
 PREDICTED: similar to KIAA0229 [Monodelphis domestica]
 CDGNAQPSVPSWLDLGLQDYIQSFLSSGYSSIDTVKNLWELELVNVLKVHLLGHRKRRIASLA
 DRPYE
 790-858_UPI0000F2C069 0.282
 >smart|SAM-uniprot|O08644|EPHB6_MOUSE/938-1005 Ephrin type-B
 receptor 6 precursor [Mus musculus]
 LDFPCLDSPQAWLSAIGLECYQDNFSKFGLSTFSDVAQLSLEDLPGLGITLAGHQKKLLHNQIOL
 LQQH
 938-1005_O08644 0.284
 >smart|SAM-uniprot|UPI0000E21804|UPI0000E21804/944-1011
 PREDICTED: ephrin receptor EphB6 isoform 2 [Pan troglodytes]
 LDFPCLDSPQAWLSAIGLECYQDNFSKFGLCTFSDVAQLSLEDLPALGITLAGHQKKLLHHIOL
 LQQH
 944-1011_UPI0000E21804 0.285
 >smart|SAM-uniprot|UPI000061005E|UPI000061005E/873-940 Ephrin
 type-A receptor 8 precursor (EC 2.7.10.1) (Tyrosine-protein
 kinase receptor EEK) (EPH- and ELK-related kinase) (HEK3).
 [Gallus gallus]
 LDFSSCLTVEDWLDSIRLGHYRDNFAMAGYSSLGMVMSMNIEDVRSLGITMMGHQKKILSSIQA
 MRSQ
 873-940_UPI000061005E 0.286
 >smart|SAM-uniprot|UPI000155E747|UPI000155E747/908-975
 PREDICTED: similar to receptor protein-tyrosine kinase [Equus
 caballus]
 PEFSAVVSVDWLQAIKMDRYKDNFTAAGYTTLEAVVHVNQDDLARIGITAVTHQNKLSSVQA
 MRTQ
 908-975_UPI000155E747 0.287

>smart|SAM-uniprot|UPI0000F200FF|UPI0000F200FF/910-977
 PREDICTED: eph-like receptor tyrosine kinase 4 [Danio rerio]
 CSAPSLASVDDWLKLIGLEQYRENFTAGYSSIESVIPMNHEDLAKMGISCSAHQRKILSSVED
 LLSG
 910-977_UPI0000F200FF 0.289
 >smart|SAM-uniprot|Q4SMF1|Q4SMF1_TETNG/641-708 Chromosome 8
 SCAF14552, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 TDLSAVSTVHEWLRALRMERYQEEFDRAHLDLDRVSLLTMEDIONLGVTLGHQRKILNAAQQ
 LRAY
 641-708_Q4SMF1 0.290
 >smart|SAM-uniprot|UPI0000E1F9B8|UPI0000E1F9B8/907-974
 PREDICTED: ephrin receptor EphA4 isoform 2 [Pan troglodytes]
 PEFSAVSVGDWLQAIKMDRYKDNFTAAGYTTLEAVVHVNVQEDLARIGITAITHQNKLSSVQA
 MRTQ
 907-974_UPI0000E1F9B8 0.291
 >smart|SAM-uniprot|UPI0000EDE459|UPI0000EDE459/908-975
 PREDICTED: similar to chromosome 10 open reading frame 53
 [Ornithorhynchus anatinus]
 PEFSAVSVTDWLQAIKMERYKDNFTAAGYTTLEAVVHMNQDDLTRIGITAITHQNKLGSVQS
 MRTQ
 908-975_UPI0000EDE459 0.292
 >smart|SAM-uniprot|Q91845|EPA4A_XENLA/908-975 Ephrin type-A
 receptor 4-A precursor [Xenopus laevis]
 PEWSQVASVLDWLQAIKMERYKDNFTAAGYTSLEAVVHVNVQDDLTRIGISSPSHQNKILSSVQG
 MRTQ
 908-975_Q91845 0.293
 >smart|SAM-uniprot|UPI0000F21D41|UPI0000F21D41/278-345
 PREDICTED: hypothetical protein [Danio rerio]
 PSFPSFSSVGEWLEAVDMGRYKDNFTAAGYCYLESVARMTVQDVLSLGITCLEHQKQILSAIQT
 LRAQ
 278-345_UPI0000F21D41 0.294
 >smart|SAM-uniprot|Q6VU50|Q6VU50_MANSE/933-1000 Eph receptor
 [Manduca sexta]
 PDLIQFASVEEWLECIKMSRYVEKF RAGGVTMDAVVDTVLQLASLGVTLVGHQKKIMNSVQS
 MRAH
 933-1000_Q6VU50 0.295
 >smart|SAM-uniprot|Q9U8V9|Q9U8V9_EPTBU/275-342 EphA [Eptatretus
 burgeri]
 TDNPRAVRTVPEWLEAIKMGRYADNF AAAGYTTLDSVAQLDADGLVRIGITLIGHQKKILSNIQT
 MKAE
 275-342_Q9U8V9 0.296

>smart|SAM-uniprot|UPI000155CF4D|UPI000155CF4D/500-567
 PREDICTED: similar to EPH receptor A10 [Ornithorhynchus
 anatinus]
 SAFPCFSSVGEWLEAIELERİKDFAAGYRSLEFVARLTIQDVANLGITTMEHQQRQLLAGIQA
 LRTQ
 500-567_UPI000155CF4D 0.297
 >smart|SAM-uniprot|Q4SFK0|Q4SFK0_TETNG/917-984 Chromosome 7
 SCAF14601, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LALSGCASPGEWLRTIKMERYEDSFLQAGLSTVDQLSQISTQDLLHMGVTLAGHQRKILSSIQS
 MTLR
 917-984_Q4SFK0 0.298
 >smart|SAM-uniprot|UPI000065CF31|UPI000065CF31/904-971 Homolog
 of Brachydanio rerio "Eph-like receptor tyrosine kinase
 rtk5. [Takifugu rubripes]
 LALSGCASPGEWLRAIKMERYEDSFLQAGLSTVDQLAQISTQDLLHMGVTLAGHQRKILSSIQT
 MTFR
 904-971_UPI000065CF31 0.299
 >smart|SAM-uniprot|UPI000155DD70|UPI000155DD70/947-1014
 PREDICTED: similar to EPH receptor A8 [Equus caballus]
 RGGSGGLTVGDWLDSIRMGRYRDHFAGGYSSLGMVLRMNAQDVRALGITLMGHQKKILGSIQT
 MRAQ
 947-1014_UPI000155DD70 0.300
 >smart|SAM-uniprot|Q4SRF4|Q4SRF4_TETNG/450-517 Chromosome
 undetermined SCAF14527, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PDFSSFCVGDWLEAIKMERYRDNFTAAGYTALESVARMSFEDVMNLGITLVGHQKKIMNSIQT
 MRAQ
 450-517_Q4SRF4 0.301
 >smart|SAM-uniprot|UPI0000F2D7AF|UPI0000F2D7AF/975-1042
 PREDICTED: similar to EPH receptor A5, [Monodelphis domestica]
 LGTGAYRSVGEWLEAIKMGRYTEIFMENGGYSSMDAVAQVTLEDLRLGVTLVGHQKKIMNSLQE
 MKVQ
 975-1042_UPI0000F2D7AF 0.302
 >smart|SAM-uniprot|UPI0000E2040F|UPI0000E2040F/914-981
 PREDICTED: ephrin receptor EphA5 isoform 1 [Pan troglodytes]
 LGSGAYRSVGEWLEAIKMGRYTEIFMENGGYSSMDAVAQVTLEDLRLGVTLVGHQKKIMNSLQE
 MKVQ
 914-981_UPI0000E2040F 0.303
 >smart|SAM-uniprot|Q1MT83|Q1MT83_DANRE/587-654 Novel protein
 similar to vertebrate Eph receptor A7 [Danio rerio]

PDFTSFRLVSEWLEAIKMERYMDNFTAAGYSSLESVARMTIEDVMSLGISLVEHQKKIMSSIQT
 MRAQ
 587-654_Q1MT83 0.304
 >smart|SAM-uniprot|Q4T1U5|Q4T1U5_TETNG/19-86 Chromosome
 undetermined SCAF10475, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PPPSACGSVSEWLRAIKMERYQQDFLQAGFTSLDVVAHLNTEDLLRVGVTLAGHQKKILSSIQT
 LRIH
 19-86_Q4T1U5 0.305
 >smart|SAM-uniprot|A3KG07|A3KG07_MOUSE/926-993 Ephrin receptor
 [Mus musculus]
 GSGNGDLTVGDWLDSIRMGRYRDHFAGGYSSLGMVLRMNAQDVRALGITLMGHQKKILGSIQT
 MRAQ
 926-993_A3KG07 0.306
 >smart|SAM-uniprot|Q9U8V7|Q9U8V7_BRABE/267-334 Eph2
 [Branchiostoma belcheri]
 ALQPVLPTVPQWLDLSDMKMGRYRDNFTNGGYSSMDLVMKMNIRDLOGIGVTMLAHQKKILNSION
 LHTQ
 267-334_Q9U8V7 0.307
 >smart|SAM-uniprot|UPI0000F21110|UPI0000F21110/898-965
 PREDICTED: similar to Cek10 protein [Danio rerio]
 PDYTTFTTVGDWLDAIKMSRYRDNFRKGCFGSSGPVDPRAEDLLRIGVTLAGHQKKILGSIQD
 MRLQ
 898-965_UPI0000F21110 0.308
 >smart|SAM-uniprot|UPI000065E0B8|UPI000065E0B8/980-1042 Homolog
 of Brachydanio rerio "EphA4 protein. [Takifugu rubripes]
 PDLSSVSSVEDWLAALKMTQYRDSFLGSGFTSLPLVTQITAEDLQRIGVSLAGHQKKILTSVQ
 980-1042_UPI000065E0B8 0.309
 >smart|SAM-uniprot|Q4SFJ9|Q4SFJ9_TETNG/699-761 Chromosome 7
 SCAF14601, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LALSGCASCVGELRTIKMERYEDSFLQAGLSTVDQLAQISTQDLLHMGVTLAGHQRKILSSIQT
 699-761_Q4SFJ9 0.310
 >smart|SAM-uniprot|UPI0000E21660|UPI0000E21660/808-875
 PREDICTED: ephrin receptor EphB4 [Pan troglodytes]
 PHYSAFGSVGEWLRAIKMGRYEEFAAGFGSFELVSQISAEDLLRIGVTLAGHQKKILASVQH
 MKSQ
 808-875_UPI0000E21660 0.311
 >smart|SAM-uniprot|UPI0000EBE37D|UPI0000EBE37D/923-990
 PREDICTED: similar to ephrin receptor EphB4 precursor isoform 2
 [Bos taurus]

PHYSAFGSGVGEWLRAIKMGRYEEFAAGFGSFELVSQISTEDLLRIGVTLAGHQQKKILASVQH
 MKSQ
 923-990_UPI0000EBE37D 0.312
 >smart|SAM-uniprot|UPI00004A6713|UPI00004A6713/208-275
 PREDICTED: similar to Ephrin type-A receptor 6 precursor
 (Tyrosine-protein kinase receptor EHK-2) (EPH homology kinase-2)
 [Canis lupus familiaris]
 PEYPLFVTVGDWLDSIKMGQYKNNFMAAGFTTFDLISRMSIDDIRRGTGVLLIGHQRRIVSSIQT
 LRLH
 208-275_UPI00004A6713 0.314
 >smart|SAM-uniprot|Q9U8W1|Q9U8W1_EPTBU/275-342 EphC1 [Eptatretus
 burgeri]
 PEYTMFCSEWLEAIKMGRYKENFTNAGYVAWTSVAQMTLEDLQRVGVTLAGHQQKRILTSIQA
 LRAQ
 275-342_Q9U8W1 0.315
 >smart|SAM-uniprot|P54758|EPHA6_RAT/957-1024 Ephrin type-A
 receptor 6 precursor [Rattus norvegicus]
 PEYPLFVTVGDWLDSIKMGQYKSNFMAAGFTTFDLISRMSIEDIRRIGVILIGHQRRIVSSIQT
 LRLH
 957-1024_P54758 0.316
 >smart|SAM-uniprot|UPI00004D424D|UPI00004D424D/898-965 Ephrin
 type-B receptor 3 precursor (EC 2.7.10.1) (Tyrosine-protein
 kinase receptor HEK-2). [Xenopus tropicalis]
 PDYTFPTVSDWLEAIKMGQYQENFLSAGFTSFHLVAQMTAEDLLRIGVTLAGHQQKKLLNSVQD
 MRLQ
 898-965_UPI00004D424D 0.317
 >smart|SAM-uniprot|Q60629|EPHA5_MOUSE/801-869 Ephrin type-A
 receptor 5 precursor [Mus musculus]
 LGSGAYRSVGEWLEAIKMGRYTEIFMENGYSSMDAVAQVTLEDLRLGVTLVGHQQKKIMSSLQ
 EMKVQ
 801-869_Q60629 0.318
 >smart|SAM-uniprot|UPI000069DB6A|UPI000069DB6A/874-941 Ephrin
 type-A receptor 6 precursor (EC 2.7.10.1) (Tyrosine-protein
 kinase receptor EHK-2) (EPH homology kinase 2). [Xenopus
 tropicalis]
 PEYHFVVTVSEWLESIKMGQYAHNFMASGFTTLDLVSRMTIDDIRGITLIGHQRRIVTSIQT
 LRLQ
 874-941_UPI000069DB6A 0.319
 >smart|SAM-uniprot|UPI0000D9CE5D|UPI0000D9CE5D/443-511
 PREDICTED: similar to cajalin 2 isoform a isoform 2 [Macaca
 mulatta]

HDGYHPTVAEWLDSIELG DYTKAFLINGYTSMDLLKKIWEVELINVLKINLIGHRK RILASLG
 DRLHD
 443-511_UPI0000D9CE5D 0.322
 >smart|SAM-uniprot|Q89T12|Q89T12_BRAJA/6-71 Blr2238 protein
 [Bradyrhizobium japonicum]
 PGTGAEGGLKRWLEGIGLAHYSDLFAQHRLDDVMADLTESDLVELGLPLGDRKRLQRAMAALF
 QA
 6-71_Q89T12 0.323
 >smart|SAM-uniprot|UPI000155CCE6|UPI000155CCE6/882-950
 PREDICTED: similar to cajalin 2 [Ornithorhynchus anatinus]
 QDGYHPTVAEWLDSIELG DYTKPFLINGYTSMDLLKKIWEVELINVLKISLIGHRK RILASLG
 DRLHE
 882-950_UPI000155CCE6 0.325
 >smart|SAM-uniprot|Q9PWR5|Q9PWR5_XENLA/902-969 Eph receptor
 tyrosine kinase precursor [Xenopus laevis]
 SEGMPFRTIAEWLDSIKMQQYTEFFMASPYNSMDKIIILMNQEDIKHLGIRQTGHQKRIAFSILG
 LKEH
 902-969_Q9PWR5 0.327
 >smart|SAM-uniprot|UPI0000E1E682|UPI0000E1E682/901-968
 PREDICTED: ephrin receptor EphA2 [Pan troglodytes]
 SEGVPFRTVSEWLESIKMQQYTEHFMAAGYTAIEKVVQMTNDDIKRIGVRLPGHQKRIAYSLLG
 LKDQ
 901-968_UPI0000E1E682 0.328
 >smart|SAM-uniprot|Q4RXA0|Q4RXA0_TETNG/969-1036 Chromosome 11
 SCAF14979, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 CDGTLFRSVPEWLESIRMSQYSESFARAGITSMEQVLALRHEDIKNIGVRLPGHMKRIAYSILG
 LKDE
 969-1036_Q4RXA0 0.332
 >smart|SAM-uniprot|Q3UNI2|Q3UNI2_MOUSE/902-969 Ephrin receptor
 [Mus musculus]
 SEGVPFRTVSEWLESIKMQQYTEHF MVAGYTAIEKVVQMSNEDIK RIGVRLPGHQKRIAYSLLG
 LKDQ
 902-969_Q3UNI2 0.333
 >smart|SAM-uniprot|UPI0000E48B74|UPI0000E48B74/6-75 PREDICTED:
 similar to connector enhancer of KSR2A [Strongylocentrotus
 purpuratus]
 IPRWSAKDVADWMKGLDFVLQPYINTILTKVTGDRLLLMTTSQDLEDLGVLKVGHQEIMLD A I
 SLLCNL
 6-75_UPI0000E48B74 0.335

>smart|SAM-uniprot|UPI0000F2B347|UPI0000F2B347/233-296
 PREDICTED: similar to sterile alpha motif domain containing 4
 isoform 2 [Monodelphis domestica]
 GTSSSSSTNVPawlksrlhkyaalfsqmtyeemmaltecqleaqnvtkgarhkivisiqklker
 233-296_UPI0000F2B347 0.338
 >smart|SAM-uniprot|Q0VA96|Q0VA96_HUMAN/231-294 SAMD4A protein
 [Homo sapiens]
 IGTSTSTNVPawlksrlhkyaalfsqmtyeemmaltecqleaqnvtkgarhkivisiqklker
 231-294_Q0VA96 0.340
 >smart|SAM-uniprot|UPI000065E100|UPI000065E100/971-1038 Homolog
 of Brachydanio rerio "Eph-like receptor tyrosine kinase 6.
 [Takifugu rubripes]
 CDGTVFRTVPEWLESIRMSQYSESFARAGITSMEQLGLRHEDIRNIGVRLPGHLKRIAYSILG
 LKDE
 971-1038_UPI000065E100 0.341
 >smart|SAM-uniprot|UPI000155DE6C|UPI000155DE6C/900-967
 PREDICTED: similar to EPH receptor A2 [Equus caballus]
 SEGVPFHTVSEWLESIKMQQYTEHFLAAGYTAIEKVVQMTNDDIKRIGVRLPGHQKRIAYSLLG
 LKDQ
 900-967_UPI000155DE6C 0.342
 >smart|SAM-uniprot|UPI000065F26A|UPI000065F26A/946-1013 Homolog
 of Brachydanio rerio "Eph-like receptor tyrosine kinase 6.
 [Takifugu rubripes]
 SDGSPFRSVSEWLESIKMSQYGENFSRAGVVTMDQVLQMKNEDIKNIGVRLPGHLKRIAYSILG
 LKDQ
 946-1013_UPI000065F26A 0.343
 >smart|SAM-uniprot|Q5Y251|Q5Y251_CRYGA/74-140 STE11p
 [Cryptococcus gattii]
 LDKWGPEQVMGFLAIHGCEHHSGIFIKHSIDGKLLLDFDMSVLTSGLITKVGERIRLMGGIKCL
 RQR
 74-140_Q5Y251 0.344
 >smart|SAM-uniprot|UPI00005E8655|UPI00005E8655/914-980
 PREDICTED: similar to Ephrin type-A receptor 1 precursor
 (Tyrosine-protein kinase receptor EPH) [Monodelphis domestica]
 SDGIPYRSVSEWLESIRMKRYILHFHSAGLDTMECVLELTTEDLKQMGITQAGHQKRILCSIQG
 FKD
 914-980_UPI00005E8655 0.345
 >smart|SAM-uniprot|UPI0000D9A9F5|UPI0000D9A9F5/917-983
 PREDICTED: ephrin receptor EphA1 isoform 2 [Macaca mulatta]
 SDGIPYRSVSEWLESIRMKRYILHFHSAGLDTMECVLELTAEDLTQMGITLPGHQKRILCSIQG
 FKD
 917-983_UPI0000D9A9F5 0.346

>smart|SAM_PNT-uniprot|UPI0000E49D06|UPI0000E49D06/256-340
 PREDICTED: hypothetical protein [Strongylocentrotus purpuratus]
 CSNYRDEQDRLNIPYDPELWTEDHTLHWLTIWASGEFANEALDHVKDIVMTGSEMCALTKEEFCE
 RFQKESEDLFWTHLEL
 256-340_UPI0000E49D06 0.362
 >smart|SAM_PNT-uniprot|Q4S0J3|Q4S0J3_TETNG/163-244 Chromosome 2
 SCAF14781, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 LEGYRKEQVRLGIPYDPVLWSADQVIHWAVWVMKEFNIDEMEIGSIHIPGRDLCAFSQEEFLQK
 VPNGEILWSHLELLRK
 163-244_Q4S0J3 0.367
 >smart|SAM_PNT-uniprot|UPI000001B0B6|UPI000001B0B6/163-244
 Transcription factor E4TF1. [Takifugu rubripes]
 LEGYRKEQVRLGIPYDPVLWSADQVIHWAVWVMKEFNIDEMEIGSIHIPGRDLCFSQEEFLQK
 VPNGEILWSHLELLRK
 163-244_UPI000001B0B6 0.371
 >smart|SAM_PNT-uniprot|UPI00005EA6C9|UPI00005EA6C9/170-251
 PREDICTED: similar to transcription factor E4TF1-60 [Monodelphis
 domestica]
 LEGYRKEQERLGIPYDPIQWSTDQVLHWVVWVMKEFSMTDIDLPTLSISGRELCSLNQEDFFQR
 VPRGEILWSHLELLRK
 170-251_UPI00005EA6C9 0.372
 >smart|SAM_PNT-uniprot|UPI000056C69D|UPI000056C69D/187-268 GA-
 binding protein transcription factor, alpha subunit [Danio
 rerio]
 LEGYRKEQVRLNIPYDPVQWTADQVIHWAVWVMKEFGIDEMEVGGIHIPGRQLCGFSQEEFLQR
 VPSGEILWSHLELLRK
 187-268_UPI000056C69D 0.373
 >smart|SAM-uniprot|Q0DLR9|Q0DLR9_ORYSJ/234-300 Os03g0850300
 protein [Oryza sativa]
 PGSMGYGGVKAWLDGLGLSRYAPVFEIHEVDDEVPLLLTLEDLKDMGIGAVGSRRKLYAAIQKL
 QRS
 234-300_Q0DLR9 0.374
 >smart|SAM-uniprot|Q94071|LIPB_CAEEL/683-749 Liprin-beta
 [Caenorhabditis elegans]
 ANKWDVHQTLRWLDDIGLPQYKDVFAENVVDGPLLLSLTANDAVERMKVVNAHHYATLARSIQFL
 KKA
 683-749_Q94071 0.382
 >smart|SAM_PNT-uniprot|UPI000069F541|UPI000069F541/170-251 GA-
 binding protein alpha chain (GABP-subunit alpha) (Transcription
 factor E4TF1-60) (Nuclear respiratory factor 2 subunit alpha).
 [Xenopus tropicalis]

LEGYRKEQERLGIPYDPLQWSVDQVLHWVLWVMKEFCLTEINVNSLGITGRELCNLNQEDFFQR
 VPRGEILWSHLELLRK
 170-251_UPI000069F541 0.384
 >smart|SAM-uniprot|UPI0000EBD618|UPI0000EBD618/1199-1264
 PREDICTED: similar to inositol polyphosphate 5-phosphatase [Bos
 taurus]
 AGGLGEAGMGAWLRAIGLERYEGLVHNGWDDLEFLSDITEEDLEEAGVQDPAHKRLLLDTLQL
 SK
 1199-1264_UPI0000EBD618 0.388
 >smart|SAM-uniprot|Q69P98|Q69P98_ORYSJ/545-611 Putative SNM1
 [Oryza sativa]
 ESNVEKRRVMEWLRLNLGSKYEEIFIKEEVDWETLQWLTEEDLLGMGITSLGPRKKIAHALCEL
 RKK
 545-611_Q69P98 0.396
 >smart|SAM-uniprot|UPI0000F2D250|UPI0000F2D250/1171-1236
 PREDICTED: similar to inositol polyphosphate phosphatase-like 1,
 [Monodelphis domestica]
 AGSLGEKGSMGAWLRALGMECYEEGLVHNGWDDLEFLSDITEEDLEEAGVQDPAHKRLLLDSLQH
 SK
 1171-1236_UPI0000F2D250 0.412
 >smart|SAM-uniprot|UPI00004DA19F|UPI00004DA19F/226-291 inositol
 polyphosphate phosphatase-like 1 [Xenopus tropicalis]
 SSSLRESSVGEWLRAIGLERYEGLIQNGWDDLEFLSDIVEEDLEEAGVLDPTHKRVILESLOA
 RK
 226-291_UPI00004DA19F 0.429
 >smart|SAM-uniprot|Q2I6J1|SHP2A_DANRE/1200-1266
 Phosphatidylinositol-3,4,5-trisphosphate 5-phosphatase 2A [Danio
 rerio]
 SSLSVDCSVGEWLQKLGLQHYEEGLLHNGWDDLEFLSDITEEDLEEAGVRDPAHKKILLASLKQ
 QQK
 1200-1266_Q2I6J1 0.435
 >smart|SAM-uniprot|UPI000065F5C0|UPI000065F5C0/1140-1202 Homolog
 of Homo sapiens "Inositol polyphosphate 5-phosphatase
 [Takifugu rubripes]
 SLVGESSVGEWLQRLGLERYEHGLLHNGWDDLEFLSDITEEDLEEAGVLDPVHKQILLQSLK
 1140-1202_UPI000065F5C0 0.438
 >smart|SAM-uniprot|Q86DA5|SARM1_CAEEL/611-678 Sterile alpha and
 TIR motif-containing protein tir-1 [Caenorhabditis elegans]
 VPGWTCADVQYWVKKIGFEYVEKFAKQMVDGDLQLTENDLKHDVGMISGLHRKRFLRELQT
 LKVA
 611-678_Q86DA5 0.443

>smart|SAM-uniprot|UPI0000D55C4D|UPI0000D55C4D/266-329
 PREDICTED: similar to Sterile alpha motif domain-containing protein 4 [Tribolium castaneum]
 SNLPVMKSILSWLKSLRLHKYSWVFHNLYQQMLDLSDESLQTIGITKGARHKLLSIAKIKER
 266-329_UPI0000D55C4D 0.445
 >smart|SAM-uniprot|Q2QQY1|Q2QQY1_ORYSJ/139-205 SAM domain family protein [Oryza sativa]
 AAEGEVVDVVEWLWGIGMGRYAAFAEAHEVDGEVLPCLTMDLRLDMGIGAVGARRKLYCAIQRL
 PPP
 139-205_Q2QQY1 0.459
 >smart|SAM-uniprot|UPI0000D9CBA4|UPI0000D9CBA4/638-705
 PREDICTED: similar to PTPRF interacting protein binding protein 1 isoform 1 [Macaca mulatta]
 FAKWTKEQVCNWLMEQGLGSYLNMGKHWIASGQTLLOASQQDLEKELGIKHSLHRKKLQLALQA
 LGSE
 638-705_UPI0000D9CBA4 0.463
 >smart|SAM-uniprot|UPI0000DA2632|UPI0000DA2632/637-704
 PREDICTED: similar to PTPRF interacting protein, binding protein 1 (liprin beta 1) [Rattus norvegicus]
 FAKWTKEQVCSWLAEQGLGSYLNSSGKHWIMSGQTLLOASQQDLEKELGIKHSLHRKKLQLALQA
 LGSE
 637-704_UPI0000DA2632 0.464
 >smart|SAM-uniprot|UPI0000F2E1B7|UPI0000F2E1B7/601-668
 PREDICTED: similar to PTPRF interacting protein binding protein 1 isoform 3 [Monodelphis domestica]
 FAKWTKEQVCNWLDQGLGSYMNNSGKHWIASGQTLLOASQQDLEKELGIKHSLHRKKLQLALQA
 LGSE
 601-668_UPI0000F2E1B7 0.465
 >smart|SAM_PNT-uniprot|UPI0000E48771|UPI0000E48771/101-182
 PREDICTED: similar to modulator of activity of ets genes [Strongylocentrotus purpuratus]
 KSPSSPLPIMSHVPADPRKWAALHVGRWLEAVSAKYALQVNKTDFVMNGRALCLMKREGFLDRV
 PENGAILFEDFRRRLR
 101-182_UPI0000E48771 0.466
 >smart|SAM-uniprot|UPI000065EBB0|UPI000065EBB0/659-726 Homolog
 of Homo sapiens "Splice Isoform 4 of Liprin-beta 1
 [Takifugu rubripes]
 FARWSKDQVCDWLQEQLGLYVNMARVWISSGQTLLOASQTDLERELGIKHPLHRKKLQLALQA
 LGSE
 659-726_UPI000065EBB0 0.469
 >smart|SAM-uniprot|Q6DCK5|Q6DCK5_XENLA/598-665 Ppfibp2-prov
 protein [Xenopus laevis]

FAKWTKDQVCSWLRDQGLGGYVNSCKQWIVSGQTLLHASQQDLEKELGIKQPLHRKKLQLALQS
 LGSE
 598-665_Q6DCK5 0.471
 >smart|SAM-uniprot|UPI00006A1F7E|UPI00006A1F7E/605-672 Liprin-
 beta-1 (Protein tyrosine phosphatase receptor type f
 polypeptide-interacting protein-binding protein 1) (PTPRF-
 interacting protein-binding protein 1) (hSGT2). [Xenopus
 tropicalis]
 FAKWTKEQVCLWLREQGLGGYVGSGKQWIVSGQTLLHASQQDLEKELGIKHPLHRKKLQLALQS
 LGSE
 605-672_UPI00006A1F7E 0.472
 >smart|SAM-uniprot|UPI0000ECD2A9|UPI0000ECD2A9/552-619 PTPRF
 interacting protein binding protein 1 [Gallus gallus]
 FAKWTKEQVCNWLQDQGLGSYISNGRHILSGQTLLQASQQDLEKELGIKHPLHRKKLQLALQA
 LGSE
 552-619_UPI0000ECD2A9 0.475
 >smart|SAM-uniprot|Q92YL0|Q92YL0_RHIME/25-91 Putative adenylate
 cyclase [Sinorhizobium meliloti]
 VDGEGAMDIGAWLRDQGLGQYEGTFRQNDIDPEVLRHLTAEDLIGVGVASVGHRRKLLAAIAAL
 REV
 25-91_Q92YL0 0.476
 >smart|SAM-uniprot|UPI0000E2294D|UPI0000E2294D/555-622
 PREDICTED: PTPRF interacting protein, binding protein 2 (liprin
 beta 2) [Pan troglodytes]
 FAQWSTERVCAWLEDGLAQYVIFARQWVSSGHTLLTATPQDMEKELGIKHPLHRKKLVAVKA
 INTK
 555-622_UPI0000E2294D 0.484
 >smart|SAM_PNT-uniprot|A0NC87|A0NC87_ANOGA/18-103
 ENSANGP00000029784 [Anopheles gambiae]
 DSDGECEQTYLPNDPHQWNAEHVSTWISWVSKNFDIFPPLEPARFPQAGSELAPFTKADFWV
 CAGSAAGGNTLAKHFA
 18-103_A0NC87 0.485
 >smart|SAM-uniprot|UPI0000DC21AA|UPI0000DC21AA/555-622 Liprin
 beta 2 [Rattus norvegicus]
 FAQWSTERVCTWLQDFGLAQYVIFARQWVASGHTLLTATPQDMEKELGIKHPLHRKKLVAVKA
 INTK
 555-622_UPI0000DC21AA 0.486
 >smart|SAM-uniprot|Q4SVD1|Q4SVD1_TETNG/583-650 Chromosome
 undetermined SCAF13765, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 FAKWSTEQVCEWLEEIGLGQYVIVARHWVTSGQTLLAATPQDLERELVLKNPLHRKKLQLAIQT
 ITSK

583-650_Q4SVD1 0.487
>smart|SAM-uniprot|UPI0000EBD637|UPI0000EBD637/553-620
PREDICTED: similar to PTPRF interacting protein, binding protein
2 (liprin beta 2) [Bos taurus]
FAQWSTERVCTWLEDGLAQYVIFARQWVTSGHTLLTATPQDMEKELGIKHPLHRKKLVLA
INTK
553-620_UPI0000EBD637 0.488
>smart|SAM-uniprot|UPI0000ECBA73|UPI0000ECBA73/556-623 Liprin-
beta-2 (Protein tyrosine phosphatase receptor type f
polypeptide-interacting protein-binding protein 2) (PTPRF-
interacting protein-binding protein 2). [Gallus gallus]
FAQWSTERVCNWLEDGLGQYVIFARQWVTSGHTLLTATPQDMEKEMGIKHPLHRKKLVLA
ISTK
556-623_UPI0000ECBA73 0.491
>smart|SAM-uniprot|UPI0000F2D7BF|UPI0000F2D7BF/552-619
PREDICTED: similar to Liprin-beta-2 (Protein tyrosine
phosphatase receptor type f polypeptide-interacting protein-
binding protein 2) (PTPRF-interacting protein-binding protein 2)
[Monodelphis domestica]
FAQWNTEQVCSWLEDGLGQYVIFARQWVSSGHTLLTATPQDMEKELGIKHPLHRKKLVLA
INTK
552-619_UPI0000F2D7BF 0.493
>smart|SAM-uniprot|UPI0000EB1088|UPI0000EB1088/581-648 Liprin-
beta-2 (Protein tyrosine phosphatase receptor type f
polypeptide-interacting protein-binding protein 2) (PTPRF-
interacting protein-binding protein 2). [Canis lupus familiaris]
FAQWSTERVCAWLEDGLGQYVIFARQWVTSGHTLLSATPQDMEKELGIKQPLHRKKLILAVKA
INTK
581-648_UPI0000EB1088 0.494
>smart|SAM-uniprot|UPI000155EA28|UPI000155EA28/554-621
PREDICTED: similar to Liprin-beta-2 (Protein tyrosine
phosphatase receptor type f polypeptide-interacting protein-
binding protein 2) (PTPRF-interacting protein-binding protein 2)
[Equus caballus]
FAQWSTERVCTWLEDGLGQYVIFARQWVTSGHTLLTATPQDMEKELGIKHPLHRKKLVLA
INTK
554-621_UPI000155EA28 0.497
>smart|SAM-uniprot|Q969H4|CNKR1_HUMAN/4-70 Connector enhancer of
kinase suppressor of ras 1 [Homo sapiens]
VETWTPGKVATWLRLDDSLQDYPFEDWQLPGKNLLQLCQPSLEALAVRSLGHQELILGGVEQL
QAL
4-70_Q969H4 0.499

>smart|SAM-uniprot|UPI0000D68174|UPI0000D68174/459-528
 PREDICTED: similar to sex comb on midleg-like 2 (Drosophila)
 [Mus musculus]
 PVNWSVNDVITFLNRLDPPLADQLYPTIKHDIDGKAMLLISSDTMIKYMGVKGVAMKFENYI
 RTLKEK
 459-528_UPI0000D68174 0.501
 >smart|SAM-uniprot|Q68M96|Q68M96_RHIME/2-61 RtsD [Sinorhizobium
 meliloti]
 DIAAWLRSLGLEEYASAFRDNDIDAQLLHLKAEDLKELGVASIGHRRKLIDAIADLRDE
 2-61_Q68M96 0.507
 >smart|SAM-uniprot|Q7JQG5|Q7JQG5_DROME/253-318 LD09801p
 [Drosophila melanogaster]
 QRQRNDELIIDWLLEMKHEEYAQLFIAAGYDLPTIARMTPEDLTAIGIKNPHHRERIKQRIDKLO
 VL
 253-318_Q7JQG5 0.520
 >smart|SAM-uniprot|UPI0000E1E745|UPI0000E1E745/4-70 PREDICTED:
 connector enhancer of kinase suppressor of Ras 1 [Pan
 troglodytes]
 VETWTPGVATWLRLGLDDSLQDYPFEDWQLPGKNLLQLCPRSLEALAVRSLGHQELILGGVEQL
 QAL
 4-70_UPI0000E1E745 0.524
 >smart|SAM-uniprot|UPI0000F1D271|UPI0000F1D271/480-546
 PREDICTED: similar to cask-interacting protein 1 [Danio rerio]
 AESKSSEAVVEWLTS AOLQFYTTNFLTAGYDLQTISRMTPEDLTAIGVTKPGHRKKMLSEISK
 NIP
 480-546_UPI0000F1D271 0.529
 >smart|SAM-uniprot|UPI0000F1DA4B|UPI0000F1DA4B/465-531
 PREDICTED: similar to cask-interacting protein 1 [Danio rerio]
 PEGKSSEAVIEWLSEFQLQVYAPNFINAGYDIPTISRMTPEDLTAIGVTKPGHRKKITSEINK
 SVN
 465-531_UPI0000F1DA4B 0.534
 >smart|SAM-uniprot|Q6DD51|CSKI2_XENLA/465-531 Caskin-2 [Xenopus
 laevis]
 LQGKDAEQIFCWLRGFQMETYVGNFISAGYDLPTIMRVTPEDLTAIGVTKPGHRKMISTEIGKL
 IVA
 465-531_Q6DD51 0.535
 >smart|SAM-uniprot|UPI00006A0F24|UPI00006A0F24/467-533 Caskin-2.
 [Xenopus tropicalis]
 LQGKDAEQIYCWLRSFQMESYVENFISAGYDLPTIMRVTPEDLTAIGVTKPGHRKKISTEIGKL
 IVA
 467-533_UPI00006A0F24 0.536

>smart|SAM-uniprot|UPI0000E2401C|UPI0000E2401C/610-676
 PREDICTED: CASK interacting protein 1 [Pan troglodytes]
 SEGKSSEAVSQWLTAFLQLQYAPNFISAGYDLPTISRMTPEDLTAIGVTKPGHRKKIAAEISGL
 SIP
 610-676_UPI0000E2401C 0.537
 >smart|SAM-uniprot|UPI000069DF96|UPI000069DF96/445-511 RNA-
 binding protein with serine-rich domain 1 (SR-related protein
 LDC2). [Xenopus tropicalis]
 LPCQSSEAVFQWLCKFQLQYAPNFINSGYDIPTISRMTPEDLTAIGVTKPGHRKKIFSEINNL
 NIG
 445-511_UPI000069DF96 0.539
 >smart|SAM-uniprot|Q4TAH3|Q4TAH3_TETNG/319-385 Chromosome
 undetermined SCAF7326, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 MKGEDSEAIYQWLSEFQLEQYSSNFIKAGYDLPTISRITPEDLTAIGVTKPGHRKKISMEIARL
 SIP
 319-385_Q4TAH3 0.540
 >smart|SAM-uniprot|UPI0000660E31|UPI0000660E31/1-63 Homolog of
 Homo sapiens "Caskin-2" [Takifugu rubripes]
 DADAIGQWLSDFQLEQYTGNFVSAGYDVPTISRMTPEDLTAIGVTKPGHRKKISMEINNLNIP
 1-63_UPI0000660E31 0.544
 >smart|SAM-uniprot|UPI0000D55AC9|UPI0000D55AC9/303-369
 PREDICTED: similar to CG12424-PA, isoform A [Tribolium
 castaneum]
 IHQGDQEIIHAWLVLDLQYDEYFPLFVSAGYDLPTIGRMTPEDLTAIGIKKPNNRKRLKAEIAQL
 NLP
 303-369_UPI0000D55AC9 0.545
 >smart|SAM-uniprot|UPI0000E8130C|UPI0000E8130C/511-577
 PREDICTED: similar to cask-interacting protein 2 [Gallus gallus]
 LEGKDAEAIYNWLSEFQLESYTVNFLSAGYDVPTISRMTPEDLTAIGVTKPGHRKKISTEIGQL
 SIA
 511-577_UPI0000E8130C 0.546
 >smart|SAM-uniprot|UPI000065E323|UPI000065E323/3-70 Homolog of
 Homo sapiens "PARX protein" [Takifugu rubripes]
 EPGESIQEVVEWLLTLRSQYTSAGFLGAGYRSLEDCGELTEERLLELKILPTGHRRRLRSLEA
 LGEK
 3-70_UPI000065E323 0.549
 >smart|SAM-uniprot|UPI0001554BD0|UPI0001554BD0/1182-1248
 PREDICTED: similar to diacylglycerol kinase eta2
 [Ornithorhynchus anatinus]
 VQKWGTDEVAAWLDLLSLGEYKETFIRHDIRGAELLHLERRDLKDLGITKVGHMKRILQGIKE
 GKG

1182-1248_UPI0001554BD0 0.552
>smart|SAM-uniprot|UPI0000F2DF5D|UPI0000F2DF5D/1151-1217
PREDICTED: similar to diacylglycerol kinase eta2 isoform 1
[Monodelphis domestica]
VQKWTGTDVAALDQLNLGEYTEIFIRHDIRGAELLHLERRDLKDLGITKVGHMKRILQGIKEL
GKS

1151-1217_UPI0000F2DF5D 0.553
>smart|SAM-uniprot|A6QPL5|A6QPL5_BOVIN/1143-1209 DGKH protein
[Bos taurus]
VHLWGTEEVAAWLEHSLCEYKDIFTRHDIRGSELLHLERRDLKDLGVTKVGHMKRILCGIKEL
SRG

1143-1209_A6QPL5 0.554
>smart|SAM-uniprot|UPI00006A04B0|UPI00006A04B0/1132-1198
Diacylglycerol kinase eta (EC 2.7.1.107) (Diglyceride kinase
eta) (DGK-eta) (DAG kinase eta). [Xenopus tropicalis]
VQKWTGTDVAWL DLLSLGEYREIFIRHDIRGSELLHLERRDLKDLGILKVGHKRILQGIKEL
SKN

1132-1198_UPI00006A04B0 0.559
>smart|SAM-uniprot|A6NFX7|A6NFX7_HUMAN/1150-1216 Uncharacterized
protein DGKH [Homo sapiens]
VQKWTGTEEVAAWLDLLNLGEYKDIFIRHDIRGAELLHLERRDLKDLGIPKVGHVVKRILQGIKEL
GRS

1150-1216_A6NFX7 0.560
>smart|SAM-uniprot|UPI00006A0B49|UPI00006A0B49/1105-1169
Diacylglycerol kinase kappa (EC 2.7.1.107) (Diglyceride kinase
kappa) (DGK-kappa) (DAG kinase kappa) (142 kDa diacylglycerol
kinase). [Xenopus tropicalis]
VDKWGTEEVAAWLDVLGLGEYKDIFIRHDIQGSELVLLERRDLKDLGIAKVGHKRILQGIKDI
T

1105-1169_UPI00006A0B49 0.562
>smart|SAM-uniprot|UPI0000F2C3EA|UPI0000F2C3EA/836-902
PREDICTED: similar to diacylglycerol kinase, delta 130kDa,
partial [Monodelphis domestica]
AHLWGTEEVAAWLEHSLCEYKEIFVRHDIRGPELLQLERRDLKDLGVTKVGHMKRILHGIRKL
SRS

836-902_UPI0000F2C3EA 0.563
>smart|SAM-uniprot|Q6A0B7|Q6A0B7_MOUSE/967-1033 MKIAA0145
protein [Mus musculus]
VHLWGTEEVAAWLEHSLCEYKDIFTRHDIRGSELLHLERRDLKDLGVTKVGHMKRILCGIKEL
SRS

967-1033_Q6A0B7 0.564

>smart|SAM-uniprot|UPI00005A4151|UPI00005A4151/1110-1176
PREDICTED: similar to diacylglycerol kinase, eta isoform 2
[Canis lupus familiaris]
VQKWGTEEVAAWLDLLNLGEYKEIFIRHDIRGAELLHLERRDLKDLGITKVGHMKRILQGIKEF
GRS
1110-1176_UPI00005A4151 0.565
>smart|SAM-uniprot|UPI0000EBD509|UPI0000EBD509/1053-1119
PREDICTED: similar to diacylglycerol kinase eta2 [Bos taurus]
VQKWGTEEVAAWLDLLNLGEYKEIFIRHDIRGAELLHLERRDLKDLGIPKVGHVVKRILQGIKEF
ERS
1053-1119_UPI0000EBD509 0.566
>smart|SAM-uniprot|A0JP53|A0JP53_MOUSE/1007-1073 Dgkh protein
[Mus musculus]
VQNWGTEEVAAWLDLLNLGEYKEIFIRHDVRGAELLHLERRDLKDLGIPKVGHMKRILQGIKEF
ERN
1007-1073_A0JP53 0.568
>smart|SAM-uniprot|UPI00006A0FDA|UPI00006A0FDA/4-72 Connector
enhancer of kinase suppressor of ras 1 (Connector enhancer of
KSR1) (hCNK1) (Connector enhancer of KSR-like) (CNK homolog
protein 1). [Xenopus tropicalis]
VSSWSPETVRNYLRLALGLDAVQVYTFQEWNVTGKDLLCLSSQQLDALGVRCIGHQEIIILEAIE
QLCAL
4-72_UPI00006A0FDA 0.569
>smart|SAM-uniprot|Q17FK9|Q17FK9_AEDAE/909-975 Tankyrase [Aedes
aegypti]
QVVAIESSVSAFLTSLQLEHLIELLEREQITMDILAEMGHEDLKQVGVSAYGFRHKILKGIA
T RAT
909-975_Q17FK9 0.573
>smart|SAM-uniprot|UPI0000EBEBAB|UPI0000EBEBAB/107-173
PREDICTED: similar to AIDA-1a [Bos taurus]
HDGYHPTSVAEWLDSIELGDYTKAFLINGYTSMDLLKKIWEVELINVADSSRQEQQITATITII
RVL
107-173_UPI0000EBEBAB 0.579
>smart|SAM-uniprot|UPI0000E46580|UPI0000E46580/1074-1140
PREDICTED: similar to diacylglycerol kinase eta
[Strongylocentrotus purpuratus]
VKLWTVEEVGAWEALLMGEYKDQFIRNDIRGSELLSLERRDLKDIGVTKVGHIKRIQQAIKDL
DKR
1074-1140_UPI0000E46580 0.590
>smart|SAM-uniprot|UPI0000D55FB2|UPI0000D55FB2/1410-1476
PREDICTED: similar to CG31187-PA [Tribolium castaneum]

VASWGTQEVAWLQLSEYIDSFVKNDIRGRELTLARRDLKDLGVTKVGHVKRILQAIKDL
TQG

1410-1476_UPI0000D55FB2 0.592
>smart|SAM_PNT-uniprot|UPI000155B956|UPI000155B956/44-126
PREDICTED: similar to B-cell lineage specific activator/TEL
oncogene fusion protein, partial [Ornithorhynchus anatinus]
APSAHRPVAPLAGGLQPVSWSRDDVGQWLKWAEKEFSLRPIDNNTFEMNGKALLLTKE
DFRYR
CPHSGDVLYELLQHIL

44-126_UPI000155B956 0.593
>smart|SAM_PNT-uniprot|UPI000065E96A|UPI000065E96A/1-87 Homolog
of Homo sapiens "Splice Isoform G of Transcription factor
ETV7 [Takifugu rubripes]
VGSPPTLEDLWRLPGRLRINPSLWDKEDVALWLHWAQKEYSLRRPEKGRFEMNGRALCLLTKE
FRHRCPSSGDVLFIEL

1-87_UPI000065E96A 0.594
>smart|SAM_PNT-uniprot|Q5R3L3|Q5R3L3_HUMAN/31-117 Ets variant
gene 7 (TEL2 oncogene) (Ets variant gene 7 (TEL2 oncogene),
isoform CRA_f) [Homo sapiens]
QINLLGEGGICKLPGRLRIQPALWSREDVLHWLRWAEQEYSLPCTAEHGFE
MNGRALCILTKDD
FRHRAPSSGDVLYELL

31-117_Q5R3L3 0.595
>smart|SAM_PNT-uniprot|UPI0000E816B8|UPI0000E816B8/20-103
PREDICTED: similar to Ets transcription factor TEL-2b [Gallus
gallus]
PRGRHTPADEGHSLALPGRLRIQPSSLWSKDDVIHWLRWAEREYS
LQQTDESKFEMNGKALCILT
KDDFRFRAPGSGMDGG

20-103_UPI0000E816B8 0.596
>smart|SAM-uniprot|UPI0000E4781C|UPI0000E4781C/1217-1284
PREDICTED: similar to mitogen-activated kinase kinase kinase 5
[Strongylocentrotus purpuratus]
GVEAKQEELSSWLQAVGLDEDSIRRFILEEDTLPLVLNSMTRDDLRS
LKIRVGAQVRIWDAVQQ
HRMN

1217-1284_UPI0000E4781C 0.598
>smart|SAM_PNT-uniprot|P97360|ETV6_MOUSE/39-125 Transcription
factor ETV6 [Mus musculus]
PRALRMEEDSIHLPTHRLQPIYWSRDDVAQWLKWAE
NEFSLRPIESNKFEMNGKALLLTKE
DFRYRSPHSGDVLYELL

39-125_P97360 0.599
>smart|SAM_PNT-uniprot|Q0VC65|ETV6_BOVIN/38-124 Transcription
factor ETV6 [Bos taurus]
PRALRMEEDSIRLPAHLRLQPMFW
SRDDVAQWLKWAE
NEFSLRPIDSNTFEMNGKALLLTKE
DFRYRSPHSGDVLYELL

38-124_Q0VC65 0.600
>smart|SAM_PNT-uniprot|UPI000155BF36|UPI000155BF36/75-161
PREDICTED: similar to Ets transcription factor TEL-2b
[Ornithorhynchus anatinus]
SFPLITEEEICRLPGQLRIQPSLWSKEDVIHWRWAKEYSLRRIVENKFEMNGKALCILTKDD
FRYRAPSSGDVLYELL

75-161_UPI000155BF36 0.601
>smart|SAM_PNT-uniprot|Q01842|POK_DROME/31-117 Ets DNA-binding
protein pokkuri [Drosophila melanogaster]
APSSQLAELKTQLPPSLPSDPRLWSREDVLVFLRCVREFDLPKLDLFDLQMNNGKALCLLTRAD
FGHRCPGAGDVLHNVL

31-117_Q01842 0.602
>smart|SAM_PNT-uniprot|O96416|O96416_DROVI/31-117 ETS DNA
binding protein Yan [Drosophila virilis]
PPSSQLAELKTQLPPSLPSDPRLWSRDDVLDLFLRCVREFDLPKLDLFDLQMNNGKALCLLTRAD
FGHRCPGAGDVLHNVL

31-117_O96416 0.608
>smart|SAM-uniprot|Q294L8|Q294L8_DROPS/1733-1799 GA16074-PA
[Drosophila pseudoobscura]
ANNWSVNEVVTWLETMQLSEYVDSFLKNDIRGKELLTLGRRDLKDLGVVKVGHVKRILQAIKDL
SEN

1733-1799_Q294L8 0.609
>smart|SAM-uniprot|Q7PTK0|Q7PTK0_ANOGA/1658-1724
ENSANGP00000013418 [Anopheles gambiae]
VSNWGVGEVVTWLEAMQLAEYVDSFIKNDIRGKELLTLARRDLKDLGVTKVGHVKRILQAIKDL
GAG

1658-1724_Q7PTK0 0.612
>smart|SAM_PNT-uniprot|Q16SU2|Q16SU2_AEDAE/95-181 Ets [Aedes
aegypti]
APPSPLADLKTLQLPPQLNTDPRIWGREGEEAVFLRCEREFDLPKFDLDFQMNNGKALCVLTKND
LAERSPGAGDVLHNVL

95-181_Q16SU2 0.613
>smart|SAM-uniprot|UPI0000DB7AC8|UPI0000DB7AC8/1293-1358
PREDICTED: similar to CG31187-PA [Apis mellifera]
VTTWGVQEVCWLENLQLGEYTEKFISHDIRGRELLSLARRDLKELGIIVKGVHKRILQAINDL
NN

1293-1358_UPI0000DB7AC8 0.614
>smart|SAM-uniprot|Q17HC2|Q17HC2_AEDAE/1646-1712 Diacylglycerol
kinase, delta, kappa, eta [Aedes aegypti]
VNNWGVGEVVTWLEAMQMTEYVDSFIKNDIRGKELLTLARRDLKDLGVTKVGHVKRILQAIKDL
GST

1646-1712_Q17HC2 0.615

>smart|SAM_PNT-uniprot|UPI0000D55CB4|UPI0000D55CB4/47-133
 PREDICTED: similar to Ets DNA-binding protein pokkuri (Protein yan) (Protein anterior open) [Tribolium castaneum]
 PPSSPLLDYKSQQLPSNLASDPRVWSREDVATFLRWAEREFDLQPIDMDMFQMNGKAICLLTRTD
 LAERAPGSGDVLYNVL
 47-133_UPI0000D55CB4 0.617
 >smart|SAM_PNT-uniprot|A4PIG5|A4PIG5_CIOIN/15-97 Ets
 transcription factor Elf [Ciona intestinalis]
 AKEDVAASDLAQLFATSPWEWSEDGFAWMESHVNQFDLDASNKLKNLHINGKELQMFQSQDEFEK
 KVPYGNVILWAHLQFLS
 15-97_A4PIG5 0.636
 >smart|SAM_PNT-uniprot|Q66PI1|Q66PI1_9BIVA/69-154 ETS-family
 transcription factor [Chlamys farreri]
 PCPFDEKNMKSWAEKHPEHWNNEVLDWIYFVAAENHLDVARLRGENFQTITGQKLCQMTQQDF
 IDKDQEYGSYYFELFH
 69-154_Q66PI1 0.647
 >smart|SAM_PNT-uniprot|Q4H3K1|Q4H3K1_CIOIN/408-495 Transcription
 factor protein [Ciona intestinalis]
 PPNVTLNPDKHKRSTLPQDPSTWTRNHVKEWLNVSSKEYNLQELDLKKFASTDGYKLCQMTMRD
 LCRITSKANAEVILNK
 408-495_Q4H3K1 0.658
 >smart|SAM-uniprot|Q6R5A4|Q6R5A4_DANRE/743-809 Bicaudal-C [Danio
 rerio]
 PSPAHADDLIELLAQLGLEKYIDIFRQOEIDYQTFLTLSDEDLKEVGVSTFGARRKMLLAIADL
 SKK
 743-809_Q6R5A4 0.677
 >smart|SAM-uniprot|UPI000065F979|UPI000065F979/1-59 Homolog of
 Brachydanio rerio "Bicaudal-C. [Takifugu rubripes]"
 LPELLSQLGLIKYIDIFEQQEIDYPTFLTLSDEDLKEVGVFTFGARRKMLLAIADLSKS
 1-59_UPI000065F979 0.681
 >smart|SAM-uniprot|Q758Y4|VTS1_ASHGO/415-479 Protein VTS1
 [Eremothecium gossypii]
 CDPKLLKNIPAWLKSRLHKEYSASLNGKSWQELIDLDDAILEDMGVSALGARRKLLKAFAIVKE
 C
 415-479_Q758Y4 0.682
 >smart|SAM-uniprot|Q6C9X3|Q6C9X3_YARLI/154-221 Similar to
 sp|P39969 Saccharomyces cerevisiae YER114c BOI2 budding protein
 [Yarrowia lipolytica]
 VMSWTPIQVARWFENRGFEPAIPKLFVEHKISGAILTELELQHLKELDITSFGTRFEVYKEIEK
 LRGD
 154-221_Q6C9X3 0.683

>smart|SAM-uniprot|UPI000155D868|UPI000155D868/810-875
 PREDICTED: similar to Protein bicaudal C homolog 1 (Bic-C)
 [Equus caballus]
 SSFKGSDLPELFSQLGLGKYTDVFQQQEIDLQTLTDQDLKELGITTGARRKMLLAISELN
 KN
 810-875_UPI000155D868 0.684
 >smart|SAM-uniprot|Q99MQ1|BICC1_MOUSE/872-938 Protein bicaudal C
 homolog 1 [Mus musculus]
 NSCFKGSDLPELFSQLGLGKYTDVFQQQEIDLQTLTDQDLKELGITTGARRKMLLAISEL
 SKN
 872-938_Q99MQ1 0.685
 >smart|SAM-uniprot|UPI0000E80747|UPI0000E80747/1087-1153
 PREDICTED: similar to bicaudal-C [Gallus gallus]
 NSLFKGSDLPELFSQLGLGKYTDVFQQQEIDLQTLTDQDLKELGITTGARRKMLLAISEL
 NKN
 1087-1153_UPI0000E80747 0.687
 >smart|SAM-uniprot|Q08831|VTS1_YEAST/448-512 Protein VTS1
 [Saccharomyces cerevisiae]
 TDPKLLKNIPMWLKSRLHKEYSDALSGTPWIELIYLDDETLEKKGVLAGARRKLLKAFGIVID
 Y
 448-512_Q08831 0.688
 >smart|SAM-uniprot|Q6CY29|VTS1_KLULA/384-448 Protein VTS1
 [Kluyveromyces lactis]
 CDPKLLKNVPAWLKSRLHKEYSEALGSKPWFEIYLDDEALENMGSALGARRKLLKAFSIVRE
 Y
 384-448_Q6CY29 0.689
 >smart|SAM-uniprot|UPI0000F2AE6A|UPI0000F2AE6A/869-935
 PREDICTED: similar to serologically defined colon cancer antigen
 3 [Monodelphis domestica]
 NSSLGSDLPELFSQLGLGKYTDVFQQQEIDLQTLTDQDLKELGITTGARRKMLLAISEL
 NKN
 869-935_UPI0000F2AE6A 0.693
 >smart|SAM-uniprot|Q6BSL1|VTS1_DEBHA/512-576 Protein VTS1
 [Debaryomyces hansenii]
 SNPDLNNIPAWLKLLRLHKEYTDCLKDIYWKDLIEYDDLLEKRGVKALGARRKLLKAFDAVKE
 S
 512-576_Q6BSL1 0.695
 >smart|SAM-uniprot|A3LX24|A3LX24_PICST/566-630 Predicted protein
 [Pichia stipitis]
 ANIELLNNIPAWLKLLRLHKEYTDCLKDTPWKELIELDNEQLELKGVAAALGARRKLLKAFDAVKLT
 T
 566-630_A3LX24 0.696

>smart|SAM-uniprot|Q6P0K1|Q6P0K1_DANRE/4-70 Connector enhancer
 of kinase suppressor of Ras 1 [Danio rerio]
 VASWSTERVTEWLKG LDAPLQQYPFSEWQLSGSDLLQLSSRLENIGVHKIGHQELILEAVEKL
 CAL
 4-70_Q6P0K1 0.699
 >smart|SAM-uniprot|UPI000065EE82|UPI000065EE82/1126-1192 Homolog
 of Brachydanio rerio "Bicaudal-C. [Takifugu rubripes]
 NSSLGSDLPELFNSNLGLGKYTDVFQQQEIDLQTLTDQDLKELGITTFGARRKMLLAISVV
 PAA
 1126-1192_UPI000065EE82 0.700
 >smart|SAM-uniprot|Q54XX5|Q54XX5_DICDI/376-441 Ankyrin repeat-
 containing protein [Dictyostelium discoideum]
 FLKNVQDIFNWLSKSIDLEQYWLNFKKEEIMFDLIDERTLDLSLGITYSGHRLKIIRNCRILR
 DQ
 376-441_Q54XX5 0.702
 >smart|SAM-uniprot|A0D9Q7|A0D9Q7_PARTE/264-331 Chromosome
 undetermined scaffold_42, whole genome shotgun sequence
 [Paramecium tetraurelia]
 MQEWQLEDVCNWLDLHLSEYKDEFIKNQMTGKTLYALTDNDLKQDLGISVLGHRKQILOQSIEE
 YKKY
 264-331_A0D9Q7 0.704
 >smart|SAM-uniprot|UPI0000E45E6F|UPI0000E45E6F/1894-1960
 PREDICTED: similar to KIAA1617 protein [Strongylocentrotus
 purpuratus]
 PLHWSVDRVVKFIFKTDASCALAKVFKDQEIDGQALLLTLPTVQECMELKLGPAIKLCHIERV
 KIA
 1894-1960_UPI0000E45E6F 0.714
 >smart|SAM-uniprot|Q0CUP3|Q0CUP3_ASPTN/540-604 Protein VTS1
 [Aspergillus terreus]
 TDPNLLKDIPSWLRLHLKYTENLKDLDKWTTELVELNDKALEERGVNALGARNKMLKVFEQVRE
 A
 540-604_Q0CUP3 0.724
 >smart|SAM-uniprot|Q2UKF0|Q2UKF0_ASPOR/542-606 Predicted protein
 [Aspergillus oryzae]
 TDPALLKDIPSWLRLHLKYTENLKDLDKWTTELVELDDKALEERGVNALGARNKMLKVFEQVRE
 A
 542-606_Q2UKF0 0.728
 >smart|SAM-uniprot|A1D4C6|A1D4C6_NEOFI/538-602 SAM domain
 protein [Neosartorya fischeri]
 TDPNLLKDIPSWLRLHLKYTDNLKDLDKWTTELIELDDKALEDRGVNALGARNKMLKVFEQVRE
 A
 538-602_A1D4C6 0.729

>smart|SAM-uniprot|Q4IBN1|VTS1_GIBZE/546-610 Protein VTS1
 [Gibberella zae]
 TDPTLLQDIPSWLRLHKYTDNLKDMKWTDLIELDDKALEERGVNALGARRKMLKVFEQVKE
 A
 546-610_Q4IBN1 0.731
 >smart|SAM_PNT-uniprot|UPI0000D99E09|UPI0000D99E09/48-132
 PREDICTED: similar to E74-like factor 3 (ets domain
 transcription factor, epithelial-specific) [Macaca mulatta]
 QMSLEGTEKTSWSGEQPQFWSKTQVLDWISYQVEKNKYDASAIDFSRCMDGATLCNCALEELR
 LVFGPLGDQLHAQLRD
 48-132_UPI0000D99E09 0.739
 >smart|SAM-uniprot|Q5BGC4|VTS1_EMENI/536-600 Protein vts1
 [Emericella nidulans]
 TDPNLLKDIPSWLRLHKYTDNLKDLKWTTELIELNDKOLEERGVNALGARNKMLKVFEQVKE
 A
 536-600_Q5BGC4 0.742
 >smart|SAM-uniprot|Q6CME5|Q6CME5_KLULA/245-312 Similar to
 sp|P39969 Saccharomyces cerevisiae YER114c BOI2 budding protein
 [Kluyveromyces lactis]
 VKNWVPEQVTAYLISLGFDVESASRFQKHKISGTILLELELAHLKELEINSFGTRFEIFKEIEA
 LKEI
 245-312_Q6CME5 0.832
 >smart|SAM-uniprot|Q751J9|Q751J9_ASHGO/223-290 AGL293Cp
 [Eremothecium gossypii]
 AELWTPEQVTAYLLSTGFDEESSNKFQEHKISGAILLELELAHLKELEINSFGTRFEIFKEIEA
 IKET
 223-290_Q751J9 0.837
 >smart|SAM-uniprot|Q6BKM8|Q6BKM8_DEBHA/264-332 Similar to
 CA5428|CaBOI2 Candida albicans CaBOI2 budding protein
 [Debaryomyces hansenii]
 ADTWTPKQVSSYFAIILGFDMDVAGKFARHKITGAILFELDLTYLKELDIDSFGTRFEVYKEIE
 KLKQL
 264-332_Q6BKM8 0.839
 >smart|SAM-uniprot|A3M0F7|A3M0F7_PICST/186-254 RHO protein
 signal transduction [Pichia stipitis]
 ASSWTPQQVSSYFAVLGFDMDVAGKFSRHKITGPILFELDLGHLKELDIDSFGTRFEVYKEIE
 KLKEL
 186-254_A3M0F7 0.840
 >smart|SAM-uniprot|Q6FTC8|Q6FTC8_CANGA/232-299 Candida glabrata
 strain CBS138 chromosome G complete sequence [Candida glabrata]
 VENWTPDEVTAYFIGHGFDIKSASSFQQHKISGKILLELQLEHLKELDISSFGTRFEMYKQIEM
 IREV

232-299_Q6FTC8 0.841
>smart|SAM-uniprot|Q4SVU7|Q4SVU7_TETNG/3-70 Chromosome
undetermined SCAF13721, whole genome shotgun sequence [Tetraodon
nigroviridis]
EPPECVREVVGWLHTLHLSQYTPSFLGAGYRTLDDCRELTEGRLLELDVLPRGHRRRMLRSLEA
LGVT

3-70_Q4SVU7 0.842
>smart|SAM-uniprot|Q7R022|Q7R022_GIALA/14-80 GLP_456_60559_61104
[Giardia intestinalis]
VEFWSKHDVSVWLSSIDMKKYNQQLRKHDVCGRSIMLLTEDDLKQMGIDSLGDRKALLYEIKRL
DRW

14-80_Q7R022 0.869
>smart|SAM-uniprot|A0BV34|A0BV34_PARTE/226-293 Chromosome
undetermined scaffold_13, whole genome shotgun sequence
[Paramecium tetraurelia]
MQDWNTNDVCVWLECLGLSQYKENFQKNQMVGELHNLTDELKEELGIQILGHRKQILOQINM
HKKY

226-293_A0BV34 0.903
>smart|SAM-uniprot|Q299K5|Q299K5_DROPS/2-66 GA12577-PA
[Drosophila pseudoobscura]
PNHNIVCEWLRTIGLAQYGESFLENGYDELEICKQIGEIDLDAIGVDNLSHRGKLLKSVRMLRE
K

2-66_Q299K5 0.915
>smart|SAM-uniprot|UPI0000DB7366|UPI0000DB7366/1-67 PREDICTED:
similar to CG31163-PB, isoform B [Apis mellifera]
MTMASNIVVEWLRSIHLGQYSESFIGDNGYDDLEICKQIGDPDLDAIGVFNQTHRARLLQSVKTL
REE

1-67_UPI0000DB7366 0.926
>smart|SAM-uniprot|Q9VD37|Q9VD37_DROME/2-66 CG13859-PA
[Drosophila melanogaster]
PHHNIVCEWLRTIGLAHYGESFLENGYDELEICKQIGEIDLDAIGVDNPSHRGKLLKSVRMLRE
K

2-66_Q9VD37 0.928
>smart|SAM-uniprot|UPI0000F2C2C7|UPI0000F2C2C7/1-65 PREDICTED:
similar to SAM domain containing 1 [Monodelphis domestica]
MCTNIVYEWLKALQLSQYAESFVDNGYDDLEVCKQIGDPDLDAIGVLAQHRRRIHDAVRLRE
Q

1-65_UPI0000F2C2C7 0.936
>smart|SAM-uniprot|A5D8S3|A5D8S3_DANRE/1-68 LOC100002699 protein
[Danio rerio]
MMTNGPNIVYEWLKTLQLCQYVEAFVDNGYDDLEVCKQIGDPDLDAIGVFIPHHRQRIHDAVQR
LKDD

1-68_A5D8S3 0.944
>smart|SAM-uniprot|UPI000155472E|UPI000155472E/1-65 PREDICTED:
similar to N-acylaminoacyl-peptide hydrolase [Ornithorhynchus
anatinus]
MCAHVVYEWLKTLRLPQYAESFVDNGYDDLEVCKEIGDPDLDAIGVSAPRERRIQQAVRRLRL
D
1-65_UPI000155472E 0.945
>smart|SAM-uniprot|Q6FWY7|Q6FWY7_CANGA/201-268 Candida glabrata
strain CBS138 chromosome C complete sequence [Candida glabrata]
IIAWAPKDVAEYMLSRGFDFTTAKFQKHQITGIIILLEMOTNMLKEIEISSFGIRFELEKEINH
LRSV
201-268_Q6FWY7 0.956
>smart|SAM-uniprot|UPI0000E21813|UPI0000E21813/911-977
PREDICTED: ephrin receptor EphA1 isoform 1 [Pan troglodytes]
PNFSAGCCVSEWLESIRMKRYILHFHSAGLDTMECVLELTAEQLTOMGITLPGHQKRILCSIQG
FKD
911-977_UPI0000E21813 1.030
>smart|SAM-uniprot|Q5LL22|Q5LL22_SILPO/1-66 Adenylate/guanylate
cyclase [Silicibacter pomeroyi]
MSRTIGSVTSWLADIGLEVYAAERFEQADIDLEVLEHLSQDLTDLGVTSLGRRKILARIADM
DA
1-66_Q5LL22 1.125
>smart|SAM-uniprot|UPI0000E4894C|UPI0000E4894C/1397-1459
PREDICTED: similar to conserved hypothetical protein
[Strongylocentrotus purpuratus]
FGEEPPLLSIFLKDLGYEKYDTNFLKAGISMIEMPYLSEEKLENIGIPIGPRLRILQEAQVMI
1397-1459_UPI0000E4894C 1.141
>smart|SAM-uniprot|A0CL90|A0CL90_PARTE/236-303 Chromosome
undetermined scaffold_20, whole genome shotgun sequence
[Paramecium tetraurelia]
MQDWNIEEVCIWLDCLGLSQYKENFIKNHMIGDTLHDLDVELKEELGIEILGHRKLILQQINM
HKKY
236-303_A0CL90 1.171
>smart|SAM-uniprot|UPI0000ECAB95|UPI0000ECAB95/3-70 Centaurin-
delta 3 (Cnt-d3) (Arf-GAP, Rho-GAP, ankyrin repeat and
pleckstrin homology domain-containing protein 3). [Gallus
gallus]
SPCTPDSDIADWLATIHLERYRDLFKQHGFHLARDVAALDNKQLOQLGITATGHRKRILNLAEK
TRLL
3-70_UPI0000ECAB95 1.209
>smart|SAM_PNT-uniprot|Q4H3F7|Q4H3F7_CIOIN/129-211 GA repeat
binding protein alpha homolog [Ciona intestinalis]

PKVSQPEKVAELEIPGDPLLWNKTQVYQWMVVWAKEFNLDTSIIMDPDLDGIELNRMSQTEFVS
TFAYGNVLWSHHALLK

129-211_Q4H3F7 1.210
>smart|SAM-uniprot|Q96X32|Q96X32_USTMA/14-80 MAP kinase pathway-interacting Ubc2 [Ustilago maydis]
VNWKWSEQQVVDWLSSVGLSKYARDFKSNGITGDVLVLLDDEALRDIGVVTIGQLALLAAIYRL
KQQ

14-80_Q96X32 1.242
>smart|SAM-uniprot|Q675P2|Q675P2_OIKDI/520-586 SH3 and multiple ankyrin repeat domains 3-like protein [Oikopleura dioica]
AACWTTKEVVTWLEANFEQYSSEFQENDISGENLLDLTKEELTEMGVTKIGHRKTLTLIQLR
KEP

520-586_Q675P2 1.285
>smart|SAM-uniprot|UPI0000F21B52|UPI0000F21B52/1039-1105
PREDICTED: similar to SH3 and multiple ankyrin repeat domains 2 [Danio rerio]
IPLWSKHDVADWLDTLNLAEHKKAFLENDIEGSHLPNLQKEDLVDLGVTRVGHRMNIEKALKLL
MDR

1039-1105_UPI0000F21B52 1.292
>smart|SAM-uniprot|Q9JLU4|SHAN3_RAT/1749-1815 SH3 and multiple ankyrin repeat domains protein 3 [Rattus norvegicus]
LQLWSKFDVGDWLESIHLGEHRDRFEDHEIEGAHPALTKEFVELGVTRVGHRMNIERALRQL
DGS

1749-1815_Q9JLU4 1.306
>smart|SAM-uniprot|UPI0000250F45|UPI0000250F45/2101-2167 SH3 and multiple ankyrin repeat domains protein 1 (Shank1) (GKAP/SAPAP-interacting protein) (SPANK-1) (Synamon) (Somatostatin receptor-interacting protein) (SSTR-interacting protein) (SSTRIP). [Rattus norvegicus]
LGFWTKF DVADWLEWLGLSEHRAQFLDHEIDGSHLPALTKEDYVDLGVTRVGHRMNIDRALKFF
LER

2101-2167_UPI0000250F45 1.307
>smart|SAM-uniprot|UPI00004D61B3|UPI00004D61B3/1152-1218
Synaptotagmin-3 (Synaptotagmin III) (SytIII). [Xenopus tropicalis]
VHFWTKYDVADWLDYLNLngREHFLDNEIDGSHLPSLTKEDYIDLGVTRVGHRMNIERALKFF
IER

1152-1218_UPI00004D61B3 1.309
>smart|SAM-uniprot|Q5RGE7|Q5RGE7_DANRE/1535-1601 Novel protein similar to vertebrate SH3 and multiple ankyrin repeat domains family protein [Danio rerio]

LALWSKYDVGEWLESVGLGEHRARFLEHEIEGAHPALTKEEIDGSHLPNLQKEDLIDLGVTRVGRMNIERALKQL
LES

1535-1601_Q5RGE7 1.313

>smart|SAM-uniprot|UPI0000EBE749|UPI0000EBE749/1908-1974
PREDICTED: similar to Shank2E [Bos taurus]
VHLWTKPDVADWLESNLGEHKEAFMDNEIDGSHLPNLQKEDLIDLGVTRVGRMNIERALKQL
LDR

1908-1974_UPI0000EBE749 1.314

>smart|SAM-uniprot|A0E8G4|A0E8G4_PARTE/210-277 Chromosome
undetermined scaffold_82, whole genome shotgun sequence
[Paramecium tetraurelia]
MQEWNIDEVCTWLDYLGLSQYQEKFIFKNHMIGEILHDLTVELKDELGIDISAHRNLILSKQIY
IRNI

210-277_A0E8G4 1.315

>smart|SAM-uniprot|Q52KW0|SHAN2_XENLA/1226-1292 SH3 and multiple
ankyrin repeat domains protein 2 [Xenopus laevis]
IHMWTQDVAEWLESIHLGEHREMFDNEIDGTHLPNLQKEDLIDLGVTRVGRMNIERALKQL
LDR

1226-1292_Q52KW0 1.316

>smart|SAM-uniprot|Q4TAT2|Q4TAT2_TETNG/1380-1446 Chromosome
undetermined SCAF7261, whole genome shotgun sequence [Tetraodon
nigroviridis]
LTLWNKYDVGDWLESVGLAEHRQRFQEHEIEGSHLPALTKEEIDGSHLPNLQKEDLIDLGVTRVGRMNIERALKQL
LDG

1380-1446_Q4TAT2 1.317

>smart|SAM-uniprot|UPI00005A206C|UPI00005A206C/1475-1541
PREDICTED: similar to SH3 and multiple ankyrin repeat domains 3
(Shank3) (Proline-rich synapse associated protein 2) (ProSAP2)
(SPANK-2) [Canis lupus familiaris]
LQFWSKFDVGDWLESIHLGEHRRDFEDHEIEGAHPALTKEEIDGSHLPNLQKEDLIDLGVTRVGRMNIERALKQL
DGS

1475-1541_UPI00005A206C 1.320

>smart|SAM-uniprot|UPI0000584890|UPI0000584890/972-1038
PREDICTED: similar to SH3 and multiple ankyrin repeat domains
protein 2 (Shank2) (Proline-rich synapse-associated protein 1)
(ProSAP1) (Cortactin-binding protein 1) (CortBP1) (GKAP/SAPAP-
interacting protein) (SPANK-3) [Strongylocentrotus purpuratus]
VSEWTVEDVSDWLEELNLGEYKESFTDNAISGEHLSLGKEDLSELGVKRLGHRLTIIKALQKL
QNQ

972-1038_UPI0000584890 1.343

>smart|SAM-uniprot|Q876M8|Q876M8_ASPFU/64-130 Ste50p
[Aspergillus fumigatus]

ITEWTAEECANWLAALGLRQYCATFLENEIVGEALIALKHEELKEMGITSVGHRLTILKSVYET
KVK

64-130_Q876M8 1.414
>smart|SAM-uniprot|Q82Y00|Q82Y00_NITEU/3-63 Guanylate
cyclase:TPR repeat:SAM domain [Nitrosomonas europaea]
NDRVSAWLDSLGLTVYHESFEHNAITWDVLPELNEDDLEALGVLLGHRKILLRAIAQLSQN
3-63_Q82Y00 1.419
>smart|SAM_PNT-uniprot|O97145|O97145_STRPU/108-190 Ets4
transcription factor [Strongylocentrotus purpuratus]
IPDILEDCHKLNLIPNVYCWTPEDEVQKWLVCANRFELGELEMGHFYINGPTLATLQDVDFRHR
APKCGDILYSVVCLLK
108-190_O97145 1.423
>smart|SAM-uniprot|UPI000023E42A|UPI000023E42A/66-132
hypothetical protein FG04101.1 [Gibberella zae]
ITEWTVEECADFIGTIGLPQYADRFMENEIVGEALVALQHDDLKSMGIASVGHRLTILKSVYDV
KKA

66-132_UPI000023E42A 1.426
>smart|SAM-uniprot|Q0MQZ4|Q0MQZ4_PENMA/64-126 STE50-like protein
[Penicillium marneffei]
ISDWTAAECADFVVGLGLRQYRTAFIVGEALIALKHEELKEMGITSAGHRLTILKSVYETKVK
64-126_Q0MQZ4 1.427
>smart|SAM-uniprot|UPI000051A5A7|UPI000051A5A7/349-415
PREDICTED: similar to CG10743-PA, isoform A [Apis mellifera]
SDKMDNTAVRWLDDIGLPQHKEAFQNGKVDGRMLHRLTTEDLLNLGVTAQLHAASLRGGIQIL
REL

349-415_UPI000051A5A7 1.456
>smart|SAM-uniprot|P36622|STE4_SCHPO/8-73 Sexual differentiation
protein ste4 [Schizosaccharomyces pombe]
YWNWNNEAVCNWIEQLGFPHKEAFEDYHILGKDIDLLSSNDLRDMGIESVGHRIDILSAIQSMK
KQ

8-73_P36622 1.510
>smart|SAM-uniprot|UPI000155C33B|UPI000155C33B/257-323
PREDICTED: similar to PTPRF interacting protein, binding protein
1 (liprin beta 1) [Ornithorhynchus anatinus]
HGKDFNWVTRWLDDIGLPQYKTQFDEARVDGRVLHYLTLDLSSLKVVSVLHHLSIKRAIQVL
RIN

257-323_UPI000155C33B 1.533
>smart|SAM-uniprot|Q29HR7|Q29HR7_DROPS/358-423 GA15414-PA
[Drosophila pseudoobscura]
VSTWTVEQVVKYLARFYPDEAEAFKQQDVDGASLLLLTREDVINGFGFKLGPALRVFQIIILGLQ
SH

358-423_Q29HR7 1.543

>smart|SAM-uniprot|UPI0000F2D7BF|UPI0000F2D7BF/624-690
 PREDICTED: similar to Liprin-beta-2 (Protein tyrosine phosphatase receptor type f polypeptide-interacting protein-binding protein 2) (PTPRF-interacting protein-binding protein 2) [Monodelphis domestica]
 SAQLDHIWVTRWLDDIGLPQYKDQFHESRVDGRMLQYLTVDLLFLKVTSQLHHLSIKCAIHVL
 HVN
 624-690_UPI0000F2D7BF 1.544
 >smart|SAM-uniprot|UPI00006090D0|UPI00006090D0/710-776
 PREDICTED: similar to mKIAA1230 protein isoform 1 [Mus musculus]
 YGKLDFNWVTRWLDDIGLPQYKTQFDEGRVDGRMLHYMTVDDLLSLKVSVLHHLSIKRAIQVL
 RIN
 710-776_UPI00006090D0 1.547
 >smart|SAM-uniprot|UPI0000ECD2A9|UPI0000ECD2A9/624-690 PTPRF interacting protein binding protein 1 [Gallus gallus]
 HGKLDYHWVTRWLDDIGLPQYKTQFDEGVGDGRMLHYMTVDDLLSLKVSVLHHLSIKRAIQVL
 RIN
 624-690_UPI0000ECD2A9 1.549
 >smart|SAM-uniprot|UPI00006A1F7E|UPI00006A1F7E/677-743 Liprin-beta-1 (Protein tyrosine phosphatase receptor type f polypeptide-interacting protein-binding protein 1) (PTPRF-interacting protein-binding protein 1) (hSGT2). [Xenopus tropicalis]
 YGKLDYRWVTRWLDDIGLPQYKTQFDDAKIDGRMLHYLAVIDLLSLKVSVLHHLSIKRAIQVL
 RIN
 677-743_UPI00006A1F7E 1.550
 >smart|SAM-uniprot|UPI000155E6CB|UPI000155E6CB/676-742
 PREDICTED: similar to PTPRF interacting protein binding protein 1 isoform 3 [Equus caballus]
 HGKDFNWWVTRWLDDIGLPQYKTQFDEGRVDGRMLHYMTVDDLLSLKVSVLHHLSIKRAIQVL
 RIN
 676-742_UPI000155E6CB 1.552
 >smart|SAM-uniprot|Q6DD79|Q6DD79_XENLA/677-743 LOC398525 protein [Xenopus laevis]
 YGRLDYRWVTRWLDDIGLPQYKTQFDDSKIDGRMLHYLAVEDLLSLKVSVLHHLSIKRAIQVL
 RIN
 677-743_Q6DD79 1.553
 >smart|SAM-uniprot|UPI0000DC21AA|UPI0000DC21AA/627-693 Liprin beta 2 [Rattus norvegicus]
 SALLDHVVWVTRWLDDIGLPQYKDQFYESRVDGRMLQYLTVDLLFLKVTSQLHHLSIKCAIHVL
 HVN
 627-693_UPI0000DC21AA 1.554

>smart|SAM-uniprot|UPI00001C1EF8|UPI00001C1EF8/627-693 Liprin-beta-2 (Protein tyrosine phosphatase receptor type f polypeptide-interacting protein-binding protein 2) (PTPRF-interacting protein-binding protein 2). [Homo sapiens]
 SALLDHIWVTRWLDDIGLPQYKDQFHESRVDRRMLQYLTVDLLFLKVTSQHHLSIKCAIHVL
 HVN
 627-693_UPI00001C1EF8 1.555
 >smart|SAM-uniprot|Q4S5D4|Q4S5D4_TETNG/631-697 Chromosome 19
 SCAF14731, whole genome shotgun sequence [Tetraodon nigroviridis]
 RGKLDHNWVTRWLDDIGLPQYKSHFEEARVDGRVLHHMTVEDLLSLKVGSVLHHLSIKRAIQVL
 RIN
 631-697_Q4S5D4 1.559
 >smart|SAM-uniprot|UPI000155460B|UPI000155460B/653-719
 PREDICTED: similar to coiled-coil like protein 1
 [Ornithorhynchus anatinus]
 SAQLDHIWVTRWLDDIGLPQYKDQFHESRVDRVHQYLTVDLLFLKVTSQHHLSIKCAIHVL
 HTH
 653-719_UPI000155460B 1.566
 >smart|SAM-uniprot|Q6DCK5|Q6DCK5_XENLA/670-736 Ppfibp2-prov
 protein [Xenopus laevis]
 YGRLDYRWVTRWLDDIGLPQYKTQFDDAKLDGRMLHYLAVEDLLSLKVVSVLHHLSIKRAIQQL
 RIN
 670-736_Q6DCK5 1.569
 >smart|SAM-uniprot|UPI0000E4628B|UPI0000E4628B/78-144 PREDICTED:
 similar to PTPRF interacting protein binding protein 1, partial
 [Strongylocentrotus purpuratus]
 QGDLDYHWVTRWLDDVGLPQYKDTFLEAKVDGRMLHYMTVDDLLLKVTSAFHHVSIMRGIQCL
 RLH
 78-144_UPI0000E4628B 1.577
 >smart|SAM-uniprot|Q86AT8|Q86AT8_DICDI/237-303 Similar to
 Dictyostelium discoideum (Slime mold). Ankyrin repeat containing
 protein [Dictyostelium discoideum]
 ARIKKYKDLFDWLQKHGFEQYKDAFLKEEMFLDELGEMSEDILNKMGITSTGTRLRILKETS
 ANE
 237-303_Q86AT8 1.616
 >smart|SAM-uniprot|UPI0000E80BD7|UPI0000E80BD7/242-309
 PREDICTED: hypothetical protein [Gallus gallus]
 IRKWTVDDVYNFIMSLPGCSDYAQTFKDHAIDGETLPLLTEEHLLDTMGLKLGPALKIRS
 QVSR
 RLGN
 242-309_UPI0000E80BD7 1.663

>smart|SAM-uniprot|Q9VKL0|Q9VKL0_DROME/795-862 CG31868-PA
 [Drosophila melanogaster]
 VNSWSVDDVCGFVGgidicaeyvqsfrdqsidgtglplltedhlvnslgmklgpalkrsilak
 klgg
 795-862_Q9VKL0 1.672
 >smart|SAM-uniprot|UPI0000DA1F03|UPI0000DA1F03/316-383
 PREDICTED: similar to sterile alpha motif domain containing 7
 [Rattus norvegicus]
 IHKWTVDVYNFIRSLPGCSDYAQVFKDHAIDGETLPLLTEQHLRGTMGLKLGPA
 LKIQSQVSOHVGN
 316-383_UPI0000DA1F03 1.673
 >smart|SAM-uniprot|Q4SVD1|Q4SVD1_TETNG/678-744 Chromosome
 undetermined SCAF13765, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 ASDGRLFSFAGWLDDIGLPQYKDOFNEGRVDGQMLQYLTVNDLLYLVTSQ
 LHLSIKCAIHVLHVN
 678-744_Q4SVD1 1.680
 >smart|SAM-uniprot|P78365|P78365_HUMAN/366-433 Polyhomeotic 2
 homolog [Homo sapiens]
 SHQVNVEDVYEFIRSLPGCQEIAEEFRAQEIDGQALLLKEDHLM
 SVMNICKLGPA
 KIYARISM
 LKDS
 366-433_P78365 1.763
 >smart|SAM-uniprot|UPI00005A45C5|UPI00005A45C5/760-827
 PREDICTED: similar to 1(3)mbt-like isoform II [Canis lupus
 familiaris]
 VAKWTIDEVFSFVQTLTGCEDQARLFKDEMIDGEAFLLL
 TQADIVKIMSVKLGPALKIYNAILM
 FKNA
 760-827_UPI00005A45C5 1.769
 >smart|SAM-uniprot|Q5RGR0|Q5RGR0_DANRE/664-731 Novel protein
 similar to human and mouse 1(3)mbt-like 3 [Danio rerio]
 VASWSIDEVIQFIQGLPGCKEQVRTFRDEQIDGEAFLLL
 TQVDLVKILSIKLGPA
 KIYNSILM
 FKTT
 664-731_Q5RGR0 1.772
 >smart|SAM-uniprot|UPI0000D9E7C6|UPI0000D9E7C6/540-607
 PREDICTED: similar to 1(3)mbt-like 3 isoform 2 [Macaca mulatta]
 VARWTVDVAEFVQSLLGCEEHAKCFKKEQIDGKA
 FLLL
 TQIDIVKVMKIKLGPA
 KIYNSILM
 FRNS
 540-607_UPI0000D9E7C6 1.773
 >smart|SAM-uniprot|UPI0000E813CE|UPI0000E813CE/756-823
 PREDICTED: similar to mKIAA0681 protein [Gallus gallus]
 VAKWTIDEVFSFVQTLTGCEDQAKLFKDEMIDGEAFLLL
 TQADIVKIMSVKLGPALKIYNAILM
 FKNA

756-823_UPI0000E813CE 1.774
 >smart|SAM-uniprot|Q32NI4|Q32NI4_XENLA/598-665 MGC131153 protein
 [Xenopus laevis]
 VGKWSADEVAQFVHQLPGCKDQSSVFKDEQIDGEAFLLLTQSDLVKILGLKLGPALKIYNSILM
 FHTL
 598-665_Q32NI4 1.775
 >smart|SAM-uniprot|UPI0000E24C72|UPI0000E24C72/540-607
 PREDICTED: hypothetical protein isoform 5 [Pan troglodytes]
 VARWTVDEVAEFVQSLLGCEEHAKCFKKEQIDGKAFLLLTQTDIVKVMKIKLGPALKIYNSILM
 FRHS
 540-607_UPI0000E24C72 1.776
 >smart|SAM-uniprot|UPI00006D58F6|UPI00006D58F6/324-391
 PREDICTED: similar to sterile alpha motif domain containing 7
 isoform 2 [Macaca mulatta]
 IQKWTWDDVHSFISSLPGCSDYAQVFKDHAIDGETLPPLLTEEHLRGTMGLKLGPALKIQSQVSQ
 HVGS
 324-391_UPI00006D58F6 1.777
 >smart|SAM-uniprot|UPI00001CF8F1|UPI00001CF8F1/404-471
 PREDICTED: similar to sterile alpha motif domain containing 11
 [Rattus norvegicus]
 INKWTWDDVCNFVGGLSGCGEYARVFREQGIDGETLPPLLTEEHLLNTMGLKLGPALKIRAQVAK
 RLGR
 404-471_UPI00001CF8F1 1.778
 >smart|SAM-uniprot|UPI0000E2569F|UPI0000E2569F/768-835
 PREDICTED: similar to mKIAA0681 protein isoform 3 [Pan
 troglodytes]
 VAKWTIDEVFGFVQTLTGCEDQARLFKDEMIDGEAFLLLTQADIVKIMSVKLGPALKIYNAILM
 FKNA
 768-835_UPI0000E2569F 1.779
 >smart|SAM-uniprot|UPI0000E1F217|UPI0000E1F217/531-598
 PREDICTED: sterile alpha motif domain containing 11, partial
 [Pan troglodytes]
 VTKWTWDDVCSFVGGLFGCGEYTRVFREQGINGETLPPLLTEEHLLSTMGLKLGPALKIRAQVAR
 RLGR
 531-598_UPI0000E1F217 1.780
 >smart|SAM-uniprot|UPI000065D5A4|UPI000065D5A4/347-414 Homolog
 of Homo sapiens "SAMD11 protein [Takifugu rubripes]
 ISKWSVEDVCGFISSLAGCAEYTQVFREQQAIDGETLPPLLTEEHLLSTMGLKLGPALKIRSQVAR
 RVGR
 347-414_UPI000065D5A4 1.781
 >smart|SAM-uniprot|UPI0000F1D97F|UPI0000F1D97F/389-456
 PREDICTED: hypothetical protein [Danio rerio]

IRKWTVDDVYSFISEIPSACAEYAQTFKEHMIDGETPLLLTEDHLLDTLGLKLGPAKIRSQLSR
RLGN

389-456_UPI0000F1D97F 1.782
>smart|SAM-uniprot|UPI000065E8F8|UPI000065E8F8/241-308 Homolog
of Brachydanio rerio "Polyhomeotic Ph2beta homolog (Phc2
protein). [Takifugu rubripes]

IRKWTVNDVYNFINSIPACSEYAQTFKDHMIDGETPLLLSEEHLLDTLGLKLGPAKIRSQVES
RFIH

241-308_UPI000065E8F8 1.783
>smart|SAM-uniprot|Q4S7Z0|Q4S7Z0_TETNG/1222-1289 Chromosome 9
SCAF14710, whole genome shotgun sequence [Tetraodon
nigroviridis]

VVAWSVDEVFRFVQNLIGCEEQARVFKEEMIDGEAFLLLTQTDIVKIMSIKLGPAKISNAILM
FKST

1222-1289_Q4S7Z0 1.787
>smart|SAM-uniprot|Q4SK78|Q4SK78_TETNG/238-305 Chromosome 15
SCAF14568, whole genome shotgun sequence [Tetraodon
nigroviridis]

VRKWTVNDVYSFISSIPTCSEYAQTFKDHMIDGETPLLLSEEHLLDTLGLKLGPAKIRSQVSQ
QIQT

238-305_Q4SK78 1.792
>smart|SAM-uniprot|UPI00004D9966|UPI00004D9966/801-865
Lethal(3)malignant brain tumor-like 3 protein (L(3)mbt-like 3
protein) (H-l(3)mbt-like protein). [Xenopus tropicalis]
VGKWSADEVAQFVHQLPGCTEQSSVFRDEQIDGEAFLLLTQNDLVKILGLKLGPAKIYNSILM
F

801-865_UPI00004D9966 1.793
>smart|SAM-uniprot|A3EXT2|A3EXT2_9HEMI/161-228 Polyhomeotic-like
protein 2-like protein [Maconellicoccus hirsutus]
PWKWTVDVCEFIKKLPEASDYVEEFSTHEIDGQALMLKENHLISVMNNMKLGLALKIVNKINA
LREP

161-228_A3EXT2 1.794
>smart|SAM-uniprot|UPI0000F31BF7|UPI0000F31BF7/525-592 Sterile
alpha motif domain-containing protein 11. [Bos taurus]
VSKWTVDDVCSFVGGLSGCGEYAPVFREQIDGETPLLLTEEHLLTTMGLKLGPAKIRAQVAK
RLGR

525-592_UPI0000F31BF7 1.796
>smart|SAM-uniprot|UPI0000F2E1F2|UPI0000F2E1F2/325-392
PREDICTED: hypothetical protein [Monodelphis domestica]
IRKWAVDDVFDFITGLPGCSDYAQIFRDHAIDGETPLLLTEEHLLDTMGLKLGPAKIRAQPT
MMII

325-392_UPI0000F2E1F2 1.797

>smart|SAM-uniprot|UPI0000E45C59|UPI0000E45C59/790-857
 PREDICTED: similar to polyhomeotic Ph2alpha homolog
 [Strongylocentrotus purpuratus]
 PSSWSVENVAMFIRSLPGCAGYADEFQSQEIDGQALMLLKEDHLMTALSMKLGPAALKIINKINT
 LDK
 790-857_UPI0000E45C59 1.798
 >smart|SAM-uniprot|Q4RVQ7|Q4RVQ7_TETNG/418-485 Chromosome 9
 SCAF14991, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 ISKWSVEDVCGFISSLAGCAEYTQVFREQAIDGETLPLLSEKHLNTMGLKLGPALKIRSQVAR
 RVGR
 418-485_Q4RVQ7 1.800
 >smart|SAM-uniprot|Q4TBZ6|Q4TBZ6_TETNG/213-279 Chromosome
 undetermined SCAF7068, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PSQWSVEEVTAFITTLPGCSDVAEAFLQEQIDGQALLLLTEDHLMTSMNICKLGPAALKICAHINA
 LKH
 213-279_Q4TBZ6 1.809
 >smart|SAM-uniprot|Q16MU8|Q16MU8_AEDAE/1474-1541
 Lethal(3)malignant brain tumor [Aedes aegypti]
 PLCWSVEDVASYVERFPGCSMVGNQIREEQVNGMAFLSMTQEDLIKYL DVKLGPALKIYNRIIH
 LRLQ
 1474-1541_Q16MU8 1.812
 >smart|SAM-uniprot|UPI000065FB40|UPI000065FB40/212-279 Homolog
 of Brachydanio rerio "Polyhomeotic Ph2beta homolog (Phc2
 protein). [Takifugu rubripes]
 PGQWNIEDVYEFISALPGCLEIAEEFRSQEIDGQALLLLKEDHLMATMNICKLGPAIKIFQAQINS
 LKDS
 212-279_UPI000065FB40 1.814
 >smart|SAM-uniprot|UPI0000437DA3|UPI0000437DA3/760-827
 Polyhomeotic-like protein 2. [Danio rerio]
 PTKWNVEDVYEFICSLPGCHEIAEEFRSQEIDGQALMLLKEDHLMSTMNICKLGPAALKIFARISM
 LKDS
 760-827_UPI0000437DA3 1.818
 >smart|SAM-uniprot|UPI00006607F9|UPI00006607F9/636-702 Homolog
 of Brachydanio rerio "Polyhomeotic Ph1 homolog. [Takifugu
 rubripes]
 PAQWSVEEVCRFISSLQGCEELAAQFLSQEIDGQALLLLREEHLISTMNNICKLGPAALKICASINT
 LRE
 636-702_UPI00006607F9 1.819

>smart|SAM-uniprot|Q4SY21|Q4SY21_TETNG/235-301 Chromosome
 undetermined SCAF12233, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PTEWNTEEVSQFIASLQGCKELASQFLSQEIDGQALLLLKEEHLMSAMNIKLGPAALKICAHINS
 LRT
 235-301_Q4SY21 1.820
 >smart|SAM-uniprot|UPI000051A220|UPI000051A220/1150-1217
 PREDICTED: similar to 1(3)mbt-like 3 isoform b [Apis mellifera]
 PRRWSNEEVVIKFIQSVPNCKEIGNIFRQHNDGEAFLMLTQEDLVSLGLRLGPAIKLYNSIVL
 LRRR
 1150-1217_UPI000051A220 1.821
 >smart|SAM-uniprot|A4QNV8|A4QNV8_DANRE/786-853 Zgc:56685 protein
 [Danio rerio]
 PTKWNVQEVFIRSLPGCQEIADEFRAQEIDGQALLLLKEDHLMSAMNIKLGPALKIFARINM
 LKDS
 786-853_A4QNV8 1.822
 >smart|SAM-uniprot|UPI00005A5775|UPI00005A5775/325-392
 PREDICTED: similar to sterile alpha motif domain containing 7
 [Canis lupus familiaris]
 IQKWTVNDVHNFISGLPGCSDYAQVFKDHAIDGETLPLLTEEHLRSTLGLKLGPALKIQSQVSQ
 HVES
 325-392_UPI00005A5775 1.823
 >smart|SAM-uniprot|UPI0000660A3A|UPI0000660A3A/212-278 Homolog
 of Homo sapiens "Homolog of polyhomeotic 3 [Takifugu
 rubripes]"
 PSQWSVEEVTAFITLPGCSDVAEAFLQEIDGQALLLTEDHLMTSMNIKLGPAALKICAHINS
 LKN
 212-278_UPI0000660A3A 1.824
 >smart|SAM-uniprot|UPI0000F204D5|UPI0000F204D5/492-559
 PREDICTED: hypothetical protein [Danio rerio]
 PVDWTVADVASYFTAAGFPEQAIAFRTQEIDGKSLLLQRNDVLTGLSIRLGPAALKIYERHVKV
 LQKT
 492-559_UPI0000F204D5 1.825
 >smart|SAM-uniprot|UPI00005EB781|UPI00005EB781/914-981
 PREDICTED: similar to polyhomeotic like 3 (Drosophila), isoform
 1 [Monodelphis domestica]
 PSIWTWDEVWAFIHSLPGCQDIADEFRAQEIDGQALLLLKEDHLMSAMNIKLGPAALKICARINS
 LKES
 914-981_UPI00005EB781 1.826
 >smart|SAM-uniprot|A0JMR9|A0JMR9_XENLA/776-843 MGC154340 protein
 [Xenopus laevis]

PSHWSVEEVYEFISLQGCQDLAEDFRSQEIDGQALLLKEEHLMSALNIKLGPA
LKICAKINLKET

776-843_A0JMR9 1.827

>smart|SAM-uniprot|UPI000065DD1C|UPI000065DD1C/23-90 Homolog of
Homo sapiens "Dnaj-like protein [Takifugu rubripes]
PANWGVADVNYFKATGFEEQATAFQDQEIDGKSLLLMTTRNDVLTGLSIKLGP
ALKIYEYHVKP
LQTO

23-90_UPI000065DD1C 1.829

>smart|SAM-uniprot|Q4V7W5|PHC2_XENLA/277-344 Polyhomeotic-like
protein 2 [Xenopus laevis]
PTKWNVEDVYDFVRSIPLGCQEISEEFRAQEIDGQALLLKEDHLM
SAMNIKLGPA
LKLYARISM
LKDS

277-344_Q4V7W5 1.830

>smart|SAM-uniprot|UPI0000E80BD6|UPI0000E80BD6/971-1038
PREDICTED: similar to early development regulator 3 [Gallus
gallus]

PSVWTVDEVWAFIHSLPGCQDI
ADEFRAQEIDGQALLLKEDHLM
SAMNIKLGPA
LKICARINS
LKES

971-1038_UPI0000E80BD6 1.831

>smart|SAM-uniprot|UPI00006A0796|UPI00006A0796/412-479
Polyhomeotic-like protein 1 (hPH1) (Early development regulatory
protein 1). [Xenopus tropicalis]
PSRWSVEEVYEFISLQGCQD
LAEDFRSQEIDGQALLLKEDHLM
SAMNIKLGPA
LKICAKINLKET

412-479_UPI00006A0796 1.834

>smart|SAM-uniprot|UPI0000F1D982|UPI0000F1D982/822-889
PREDICTED: similar to polyhomeotic like 3 (Drosophila), [Danio
rerio]

PSQWSVEQVCSFISTLPGCHDVA
AEFGCQEIDGQALLLTEDHLM
SAMNLKLGP
ALKICAHINT
LKQT

822-889_UPI0000F1D982 1.835

>smart|SAM-uniprot|Q4RIF1|Q4RIF1_TETNG/605-672 Chromosome 11
SCAF15043, whole genome shotgun sequence [Tetraodon
nigroviridis]
PTKWNVEEVYDFIRSLPGCQEIA
DEFRSQEIDGQALLLKEDHLM
STMNIKLGPA
LKIFARINM
LKDS

605-672_Q4RIF1 1.838

>smart|SAM-uniprot|P39769|PHP_DROME/1510-1577 Polyhomeotic-
proximal chromatin protein [Drosophila melanogaster]
ISSWSVDDVSNFIRELPGCQDYVDDFIQQEIDGQALLLK
EKHLVNAMGMKLGPA
LKIVAKVES
IKEV

1510-1577_P39769 1.853

>smart|SAM-uniprot|UPI0000DBF086|UPI0000DBF086/240-309 sex comb
 on midleg-like 2 (Drosophila) [Rattus norvegicus]
 PSTWSVDEVIQFMKHTDPHISGPLADLFRQHEIDGKALLLKSELIMKYMGLKLGPALKLCYYI
 EKLKER
 240-309_UPI0000DBF086 1.861
 >smart|SAM-uniprot|UPI0000E25C7E|UPI0000E25C7E/255-324
 PREDICTED: sex comb on midleg-like 1 [Pan troglodytes]
 PSTWSVEAVVLFLKQTDPVALCPLVDLFRSHEVDGKALLLTSDVLLKHFGVKLGTAVKLCYYI
 DRLKQG
 255-324_UPI0000E25C7E 1.863
 >smart|SAM-uniprot|Q1JQ83|Q1JQ83_XENLA/630-699 MGC130971 protein
 [Xenopus laevis]
 PAMWSVDDVMRFVKEADPQSLAPHAELFRRHEIDGTALLLKSDMIMKYMGLKLGPALKLCYHI
 ERLKQG
 630-699_Q1JQ83 1.865
 >smart|SAM-uniprot|Q4R8H0|Q4R8H0_MACFA/590-658 Testis cDNA
 clone: QtsA-12509, similar to human sex comb on midleg homolog 1
 (Drosophila) (SCMH1), [Macaca fascicularis]
 PSSWTVEDVMQFVQEADPQLGPHADLFRKHEIDGKALLLRSGMMMKYMGLKLGPALKLSYHID
 RLKQG
 590-658_Q4R8H0 1.867
 >smart|SAM-uniprot|UPI0000F32863|UPI0000F32863/223-292 Sex comb
 on midleg-like protein 1. [Bos taurus]
 PSIWSVDEVTMFLQDVDPQTDFDPLVDLFRDHGIDGKALLLRSDMMIKYMGLKVGTALKICHYI
 DRLKRE
 223-292_UPI0000F32863 1.868
 >smart|SAM-uniprot|UPI0000DA230B|UPI0000DA230B/803-870
 PREDICTED: similar to Lethal(3)malignant brain tumor-like
 protein (L(3)mbt-like) (L(3)mbt protein homolog) (H-1(3)mbt
 protein) (H-L(3)MBT) (L3MBTL1) [Rattus norvegicus]
 VSKWTIEEVFGFVQTLTGSEDQARLFKEEMIDGEAFLLLTQADIVKIMSVKLGPALKIYNAILM
 FKNN
 803-870_UPI0000DA230B 1.869
 >smart|SAM-uniprot|Q4SV88|Q4SV88_TETNG/618-687 Chromosome 1
 SCAF13775, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PSSWSIEEVMQFVRDADPTALAPHAELFRKHEIDGKALMLRSDMIMKYMGLKLGPALKLCHHI
 ERLKQG
 618-687_Q4SV88 1.871
 >smart|SAM-uniprot|UPI0000D61EC3|UPI0000D61EC3/363-430
 Polyhomeotic-like protein 2 (hPH2) (Early development regulatory
 protein 2). [Homo sapiens]

VSKWNVEDVYEFIRSLPGCQEIAEEFRAQEIDGQALLLKEDHLMSAMNIKLGPA
 LKSIYARISM
 LKDS
 363-430_UPI0000D61EC3 1.872
 >smart|SAM-uniprot|UPI00005A15E8|UPI00005A15E8/702-769
 PREDICTED: similar to l(3)mbt-like 3 [Canis lupus familiaris]
 VARWTVEEVAFVQSLLGCEEHAKCFKKEQIDGKAFLLTQTDIVKVMKIKLGPA
 LKSIYNSILM
 FRNS
 702-769_UPI00005A15E8 1.873
 >smart|SAM-uniprot|UPI00005A2CEB|UPI00005A2CEB/553-621
 PREDICTED: similar to Polycomb protein SCMH1 (Sex comb on midleg
 homolog 1) isoform 1 [Canis lupus familiaris]
 PSSWTVEDVMQFVREADPQLGPHADLFRKHEIDGKALLLRSDMMMKYMGLKLGP
 ALKLSFHID
 RLKQG
 553-621_UPI00005A2CEB 1.874
 >smart|SAM-uniprot|UPI0000F2C803|UPI0000F2C803/474-541
 PREDICTED: similar to l(3)mbt-like 4 (Drosophila), [Monodelphis
 domestica]
 VAKWTVDEVANFVQSLPGCEDHSRYFKKEQIDGKAFLLTQTDIVKVMRIKLGPA
 LKSIYNSILM
 FRNA
 474-541_UPI0000F2C803 1.876
 >smart|SAM-uniprot|Q9UQR0|SCML2_HUMAN/628-697 Sex comb on
 midleg-like protein 2 [Homo sapiens]
 PSTWSVDEVIQFMKHTDPQISGPLADLFRQHEIDGKALFLLKSDVMMKYMGLKLGP
 ALKLCYYI
 EKLKEG
 628-697_Q9UQR0 1.877
 >smart|SAM-uniprot|UPI0000F2147C|UPI0000F2147C/553-620
 PREDICTED: hypothetical protein [Danio rerio]
 VTHWSVEEVADFIHSLPGCEEQAKQFREEQIDGKAFLLTQRDIVKIMSVKLGP
 ALKSIYNSILM
 FKHA
 553-620_UPI0000F2147C 1.878
 >smart|SAM-uniprot|UPI0000ECA191|UPI0000ECA191/622-690 Polycomb
 protein SCMH1 (Sex comb on midleg homolog 1). [Gallus gallus]
 PSCWTVEEVMFIREADPQLGPHADLFRKHEIDGKALLLRSDMMMKYMGLKLGP
 ALKTYHID
 KLKQG
 622-690_UPI0000ECA191 1.889
 >smart|SAM-uniprot|Q96GD3|SCMH1_HUMAN/590-658 Polycomb protein
 SCMH1 [Homo sapiens]
 PSSWTVEDVMQFVREADPQLGPHADLFRKHEIDGKALLLRSDMMMKYMGLKLGP
 ALKLSYHID
 RLKQG
 590-658_Q96GD3 1.890

>smart|SAM-uniprot|UPI0000660F04|UPI0000660F04/161-228 Homolog
 of Homo sapiens "Splice Isoform 4 of Lethal [Takifugu
 rubripes]
 KPDASVLQVFRFVQNLIGCEEQARVFKDEMIDGEAFLLTQTDIVKIMSIKLGPALKISNAILM
 FKST
 161-228_UPI0000660F04 1.908
 >smart|SAM-uniprot|UPI0000D9C6B0|UPI0000D9C6B0/1-67 PREDICTED:
 similar to l(3)mbt-like 3 [Macaca mulatta]
 MSSLVSKVFGVQTLTGCEDQARLFKDEMIDGEAFLLTQADIVKIMSVKLGPALKIYNAILMF
 KNA
 1-67_UPI0000D9C6B0 1.929
 >smart|SAM-uniprot|UPI0000E8015B|UPI0000E8015B/684-745
 PREDICTED: similar to KIAA1798 protein isoform 2 [Gallus gallus]
 VSKWSTDEVDEHQSLPLCVFLLKQIDGEAFLLMQSDIVKIMSIKLGPALKIFNSILMFKA
 A
 684-745_UPI0000E8015B 1.950
 >smart|SAM-uniprot|UPI0000EB4740|UPI0000EB4740/815-883
 Lethal(3)malignant brain tumor-like 3 protein (L(3)mbt-like 3
 protein) (H-l(3)mbt-like protein). [Canis lupus familiaris]
 AGAACMKRVDRFVISLEKCEAEGKCVFSFSQIDGEAFLLMQTDIVKIMSIKLGPALKIFNSIL
 MFKA
 815-883_UPI0000EB4740 1.952
 >smart|SAM-uniprot|Q1RLH6|Q1RLH6_CIOIN/446-513 Zinc finger
 protein [Ciona intestinalis]
 VKTWSIEKVAEFINVLTGKTECGTFIDHEIDGESLLLLTQEDILKILKIKLGPAVKIYNAILM
 FKT
 446-513_Q1RLH6 1.965
 >smart|SAM-uniprot|Q5KJP5|Q5KJP5_CRYNE/17-83 Protein kinase
 regulator, putative [Filobasidiella neoformans]
 VLQWDEDAAVSYLSSIGLQYEAAIHEHGIAGDVLVALDHETLQDMGMSSVGHLMLRAVYEL
 KME
 17-83_Q5KJP5 2.022
 >smart|SAM-uniprot|Q4H2U6|Q4H2U6_CIOIN/636-704 Sex comb on
 midleg like protein [Ciona intestinalis]
 PTKWSVNDVVSYIGEREPAQPHLHLFNKHEIDGAALLLNnnnnIVKFMSLKLGP TLKLC
 SLVE
 GLKEK
 636-704_Q4H2U6 2.322
 >smart|SAM-uniprot|UPI0000E7FAC4|UPI0000E7FAC4/635-704
 PREDICTED: hypothetical protein [Gallus gallus]
 PSTWSIEEVMCVKEADPQALAPHAELFRRHEIDGKALLLRSDMIMKYMGLKLGPALKLCYHI
 ERLRQG
 635-704_UPI0000E7FAC4 2.338

>smart|SAM-uniprot|UPI0000EBC71D|UPI0000EBC71D/583-651
PREDICTED: similar to sex comb on midleg homolog 1 (*Drosophila*)
isoform 7 [*Bos taurus*]
PSSWTVEDVMQFVREADPQFGPHADLFRKHEIDGKALLLRSDTMMKYMGLKLGPALKLCYHID
RLKQG
583-651_UPI0000EBC71D 2.341
>smart|SAM-uniprot|UPI0000F2C14C|UPI0000F2C14C/348-417
PREDICTED: similar to sex comb on midleg-like 4 (*Drosophila*),
isoform 2 [*Monodelphis domestica*]
PSTWTVEDVVVFVKDADPQALGPHVELFRKHEIDGNALLLKSDMIMKYLGLKLGPALKLCYHI
DKLKQS
348-417_UPI0000F2C14C 2.343
>smart|SAM-uniprot|UPI0000F2032D|UPI0000F2032D/540-608
PREDICTED: similar to sex comb on midleg homolog 1 (*Drosophila*),
[*Danio rerio*]
PNLWTVEDVMQFIRDIDPQLGPHADLFRKHEIDGKALLLRSDVMMKYMGLKLGPALKLSFHID
RLKEG
540-608_UPI0000F2032D 2.347
>smart|SAM-uniprot|Q0IHT6|Q0IHT6_XENTR/635-704 Sex comb on
midleg-like 2 [*Xenopus tropicalis*]
PATWSVDDVMRFVKEADPQALAPHAELFRRHEIDGTALLLRSDMIMKYLGLKLGPALKLCYHI
ERLKQG
635-704_Q0IHT6 2.351
>smart|SAM-uniprot|UPI00004BBD7D|UPI00004BBD7D/343-412
PREDICTED: similar to sex comb on midleg-like 4 [*Canis lupus*
familiaris]
PSTWTVEDVVRFVKDADPQALGPHVELFRKHEIDGNALLLKSDMIMKYLGLKLGPALKLCYHI
DKLKQA
343-412_UPI00004BBD7D 2.353
>smart|SAM-uniprot|UPI00006A0C5F|UPI00006A0C5F/605-673 Sex comb
on midleg homolog 1. [*Xenopus tropicalis*]
PSLWNTEDEVMQFVRDADPQLGFHAELFRKHEIDGKALLLRSDVMMKYMGLKLGPALKLSHHID
KLKQG
605-673_UPI00006A0C5F 2.355
>smart|SAM-uniprot|UPI000155C8CF|UPI000155C8CF/243-312
PREDICTED: similar to sex comb on midleg-like 4 (*Drosophila*)
[*Ornithorhynchus anatinus*]
PSAWTVDDVVCFVEGVDPRAFGPHAALFRKHEIDGSALLLKSDVMKYLGLKLGPALKLCYHI
DKLKQT
243-312_UPI000155C8CF 2.357
>smart|SAM-uniprot|UPI0000F209EB|UPI0000F209EB/338-407
PREDICTED: hypothetical protein [*Danio rerio*]

PSRWSVDEVVWFIFKDADPQALGPHVELFRKHEIDGDALLLKSDMIMKYLGLKLGPALKLCYHI
 DKLQK
 338-407_UPI0000F209EB 2.358
 >smart|SAM-uniprot|Q9VHA0|SCM_DROME/803-871 Polycomb protein Scm
 [Drosophila melanogaster]
 PIDWTIEEVIQYIESNDNSLAVHGDLFRKHEIDGKALLLNSEMMMKYMGALKLGPALKICNLVN
 KVNGR
 803-871_Q9VHA0 2.361
 >smart|SAM-uniprot|Q172L2|Q172L2_AEDAE/808-876
 Lethal(3)malignant brain tumor [Aedes aegypti]
 TSDWTIEDVIRFIAVQDPALAVHADLFRKHEIDGKALLLNSDMMMKYMGALKLGPALKICNLVS
 RAKGR
 808-876_Q172L2 2.362
 >smart|SAM-ensembl|ENSPTRP00000049763|ENSPTRP00000049763/101-170
 sex comb on midleg-like 4 [Source:RefSeq_peptide;Acc:NP_932347]
 [Pan troglodytes]
 PSAWTVEDVVWFVKDADPQALGPHVELFRKHEIDGNALLLKSDMVMKYLGLKLGPALKLCYHI
 DKLQK
 101-170_ENSPTRP00000049763 2.363
 >smart|SAM-uniprot|Q4RRA4|Q4RRA4_TETNG/186-255 Chromosome 14
 SCAF15003, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PSSWGVEEVVSFIKDADPQALGPHADTFRRHEIDGDALLLKSEMMMKYLGALKLGPALKLSYHI
 DNLKQS
 186-255_Q4RRA4 2.365
 >smart|SAM-uniprot|Q4T6D4|Q4T6D4_TETNG/11-78 Chromosome
 undetermined SCAF8815, whole genome shotgun sequence [Tetraodon
 nigroviridis]
 PNLTVDNVMQYIRDIDPLAPHADLFRKHEIDGKALLLRSDTMMKYMGALKLTFHID
 KLKR
 11-78_Q4T6D4 2.366
 >smart|SAM-uniprot|UPI000065D460|UPI000065D460/182-251 Homolog
 of Homo sapiens "OTTHUMP00000040508 [Takifugu rubripes]
 PSSWGVEEVVSFIKDADPQALGPHDAFRKHEIDGDALLLKSEMMMKYLGALKLGPALKLSYHI
 DKLQK
 182-251_UPI000065D460 2.373
 >smart|SAM-uniprot|UPI0000D572FF|UPI0000D572FF/527-604
 PREDICTED: similar to CG9495-PA [Tribolium castaneum]
 TQEWGIEEVIQFIESADSCLGVHADLFRKHSLVNWELLQEIDGKALLLNSDMMMKYMGALKLGP
 ALKICNLVSRLKGR
 527-604_UPI0000D572FF 2.401

>smart|SAM-uniprot|Q6C5C5|Q6C5C5_YARLI/3-69 Similar to
tr|AAM08676 Aspergillus fumigatus AAM08676 Ste50p [Yarrowia
lipolytica]
SQAWNTDQVGQWISSLGFPQYSKSFIDNNITGDVLVHLDHDDISDIGVT
KEK
3-69_Q6C5C5 2.724

Chapter 2

TMKink: a method to predict transmembrane helix kinks

TMKink: A method to predict transmembrane helix kinks

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Abstract: A hallmark of membrane protein structure is the large number of distorted transmembrane helices. Because of the prevalence of bends, it is important to not only understand how they are generated but also to learn how to predict their occurrence. Here, we find that there are local sequence preferences in kinked helices, most notably a higher abundance of proline, which can be exploited to identify bends from local sequence information. A neural network predictor identifies over two-thirds of all bends (sensitivity 0.70) with high reliability (specificity 0.89). It is likely that more structural data will allow for better helix distortion predictors with increased coverage in the future. The kink predictor, TMKink, is available at <http://tmkinkpredictor.mbi.ucla.edu/>.

Keywords: membrane protein; protein structure; structure prediction; protein folding

Introduction

Roughly half of all transmembrane helices contain bends or other deviations from ideality.^{1,2} Distortions in helix geometry can facilitate conformational changes required for protein function by providing sites of flexibility^{3,4} and can be important for positioning key residues precisely in the structure.⁵ Kinks that open the polar backbone to alternative hydrogen bonds are often wedged by water, thereby providing a polar region within the hydrophobic core.⁶ Proline kinks can also prevent off-pathway events during the folding of membrane proteins, thereby serving as a negative design feature.⁷

Because of the common occurrence of helix breaks in membrane proteins, predicting where they occur could be an important tool for membrane protein structure prediction.^{8,9} Existing structure prediction efforts have typically started with the prediction of transmembrane helices that are then packed together in a separate step.^{8,10–13} Clearly, knowing where helix deviations are likely to occur would be useful information for packing together transmembrane segments, but this requires that local sequence at least partially encodes the distortion. An early indication that local sequence can provide predictive information about helix deviations was the work of Rigoutsos *et al.*, who found predictive patterns in transmembrane sequences, although the database at the time was too small to perform rigorous cross validation.¹⁴ Langelaan *et al.* developed a kink prediction method exploiting the recent dramatic improvement in database size, but the performance is hard to assess because the database did not exclude homologous proteins.²

One sequence signature that is clearly a powerful indicator of helix kinking is the presence of a proline, an amino acid that is incompatible with a

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helix.^{1,2,5,15,16} Yohannan *et al.* found that kinks can often be identified by looking for prolines in the aligned sequences of homologs.⁵ For 36 of 39 kinks examined, Pro occurred near the kink in at least 10% of homologous family members. More recent work with a larger database suggests that the Pro signature may much be less common than originally seen,¹ however (also see below). Nevertheless, it is clear that Pro in either the protein itself or in a homolog provides strong predictive information.

There are indications that nonproline residues can also provide information about kink formation, although the picture is much less clear. Hall *et al.* found that Ser, Thr, and Gly are common in kinked helices.¹ Ser and Thr may collaborate with Pro to modulate bend angle.¹⁷ Langelaan *et al.* did not observe the enhancement in Ser, Thr, and Gly frequencies but did find changes in the prevalence of other polar residues.² Clearly as our database expands, our understanding of residue preferences is evolving.

Here, we examine kinked helices and find that there are distinct residue preferences in kinked versus nonkinked helices in a nonredundant database. We exploit these differences and residue conservation to predict kinked helices using a neural network algorithm.

Results and Discussion

Kinked helix search space

To identify possible sequence differences between kinked and nonkinked helices, we first constructed a library of kinked and nonkinked regions from a database of 41 nonhomologous, high-resolution membrane protein structures. Although there are now many more unique structures available, we believe it is important to reduce biases as much as possible by only using unrelated proteins.

We examined nine-residue segments of transmembrane helices and identified 323 kinked and 567 nonkinked segments, defined by strict bend angle criteria as described in Methods. Using only bend angle criteria has the disadvantage of lumping together many different helix anomalies (π -helix, 3_{10} helix, etc.).¹⁶ Moreover, some helix distortions do not lead to a change in bend angle and are ignored by this criterion. Nevertheless, we are restricted by the small data set of nonhomologous high-resolution membrane protein structures currently available, so we opted not to refine the kink type categories further.

We bolstered the limited sequence data available by adding information from homologous sequences to each segment. We counted the amino acids found at each position in the kinked and nonkinked segments, but we reduced counting biases by (1) weighting the counts by sequence divergence so as

not to overcount close homologs to the protein of known structures and (2) using only 100 randomly chosen homologs per segment so that all segments were roughly equally weighted (see Methods). We were unable to find 100 homologs for 75 of the kinked segments, and these were eliminated from the residue preference analysis.

Amino acid preferences in kinked helices

The observed amino acid abundance ratios for kinked versus nonkinked helices are shown in Figure 1 (histograms corresponding to this data can be found in Supporting Information Fig. S1); the kink center was defined as position 5. As expected from prior work,^{1,2,5} Pro is highly overrepresented at positions 5 through 9 of the kinks. The bias of Pro toward the C-terminus of kinks makes sense because the loss of the hydrogen bond and steric clashes occur at residues preceding the proline. The spread of Pro over many kink positions at least partly reflects the difficulty of defining the center of a kink as well as diversity of kink structures. The overall occurrence of Pro in kinks is lower than we had observed previously.⁵ In particular, we found that in a smaller, less diverse database, ~90% of kinks contained Pro in 10% of homologs. In the current database, the percentage decreased to 56%. Although Pro had the most pronounced change, other residues also exhibited significant biases. Other than Pro, the residues that were at least twofold overrepresented in kinked helices were as follows: Trp at position 1, Asn at position 4, Trp at position 6, and Glu at position 8. Residues at least twofold underrepresented were Glu at position 1, Asn at position 2, Gln at position 3, Thr, Lys, and Arg at position 4, Asn at position 6, His at position 7, Gln at position 8, and Arg at position 9. The relative dearth of strongly polar residues in kinked helical regions was also observed by Langelaan *et al.*² This result is perhaps surprising as polar side chains might be expected to help satisfy any broken backbone hydrogen bonds¹ or support hydrogen bonding to water molecules that often wedge kinks.⁶ We do not see the preferences for Gly, Ser, and Thr observed by Hall *et al.*¹ nor the dramatic enhancement of Asp noted by Langelaan *et al.*,² perhaps because of differences in our database construction. Because of the variety of kinks and the likely variety of kinking mechanisms, however, understanding the reason for the residue preferences in kinks is not straightforward. Nevertheless, the results indicate that there are differences in amino acid composition in kink positions that could be exploited for kink prediction.

A kink predictor

We developed a neural network analogous to secondary structure prediction algorithms.^{19–21} A feedforward network consisting of an input, hidden, and

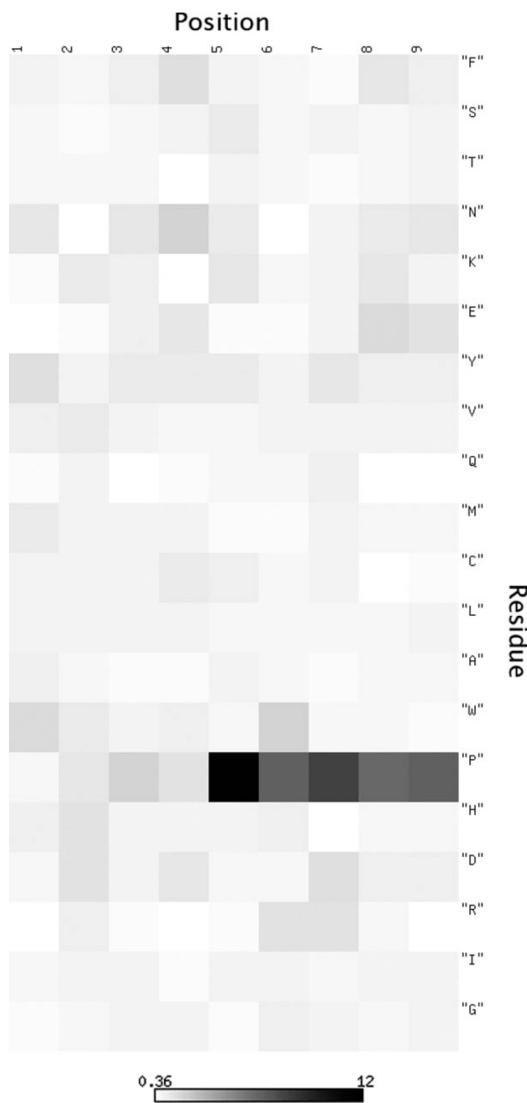


Figure 1. Amino acid composition differences in kinked and nonkinked transmembrane segments. The ratio of residue frequencies in kinked versus nonkinked nine-residue segments from the strict database is shown. Darker regions signify residues that are overrepresented when comparing kinked to nonkinked structures. The position in the segment is labeled along the horizontal axis, and residue type is labeled along the vertical axis. A numerical scale has been provided in the bottom half of the figure. A more quantitative view of this data is provided by histograms in the Supporting Information. This heatmap was generated using Matrix2png.¹⁸

output layer was constructed as shown in Figure 2. The network inputs were amino acid sequence composition at each position and a measure of sequence conservation as described under Methods. The weights were adjusted using back propagation,²¹ and we employed early stopping to prevent over-

training of the network.²² Network performance was assessed using the leave-one-out method. We found that five hidden nodes provided the best performance on the strict set of training kinks and non-kinks. We also tested performance with 7, 9, 15, and 19 residue window sizes and found that a nine-residue window resulted in the best network performance.

The performance of the network can be seen in the receiver-operator characteristic plot shown in Figure 3. We defined a network output threshold that maximized the Matthews correlation coefficient for the strict and relaxed kink databases. The threshold point for the strict database is shown in Figure 3 and yields a Matthews correlation coefficient of 0.40. At this threshold, the sensitivity is 0.46 and the specificity is 0.99. Thus, about half of all kinks are predicted and those that are predicted are almost always correct.

We next tested the kink predictor with more realistic criteria. The training data set employed strict criteria to rigorously separate kinks and nonkinks, but it omits visually obvious kinks. We therefore tested our kink predictor using more relaxed criteria for kink identification that more accurately reflect what is seen by eye (see Methods). We also assessed prediction performance more accurately in practice by performing predictions across all possible protein windows rather than just selected nine-residue windows in our strict database. Using the relaxed criteria, the sensitivity increased to 0.70 and the specificity diminished somewhat to 0.89. The statistics described in Table I indicate that the majority of kinks are predicted, and when a kink is predicted, it is almost always a correct prediction. As there were no known freely available TM kink prediction algorithms with which to compare our method, we compared our prediction algorithm to a well-known secondary structure prediction algorithm developed for soluble proteins PSIPRED.²³ We defined predicted kinks as a predicted coil or strand and nonkinks as predicted helices. The results using the relaxed kink criteria are shown in Table I. Considering the very different physicochemical basis of helix formation in soluble and membrane proteins, PSIPRED does surprisingly well. Nevertheless, our method clearly outperforms PSIPRED.

The performance for our neural network predictions is illustrated for two known structures in Figure 4. These examples were chosen to simply highlight the types of correct and incorrect predictions possible. To give a fuller picture, predictions for all structures in the database are given in the supplement. The predicted segments were always excluded during network training (see Methods). For the structure 1OTS [Fig. 4(A)], seven kinks were correctly identified (highlighted in red; true positives), four were missed (dark green; false negatives), three

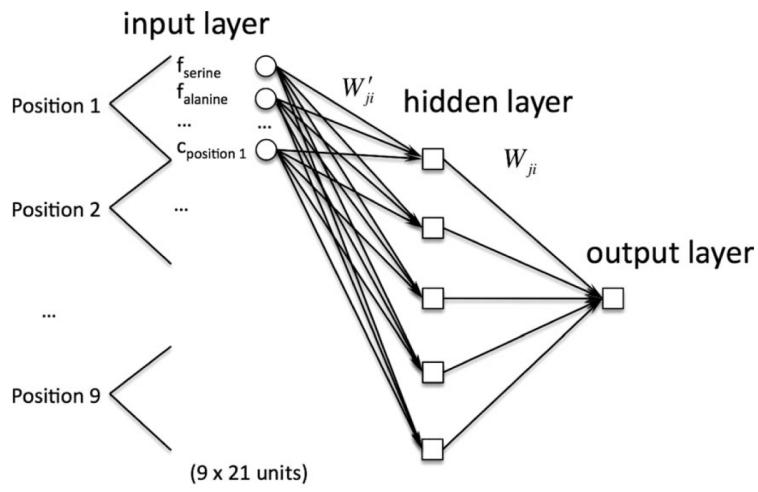


Figure 2. Neural network design. The neural network consisted of three layers: input, hidden, and output. The inputs are the weighted fractions of each residue type and a conservation score for each position in a set of homologous sequences (see Methods). The total number of inputs was 189 because of 9 positions \times 20 residues for each candidate, and nine conservation score inputs. Two sets of weight matrices were evolved through learning: one between the input and hidden layers and the other between the hidden and output layers. For the output layer, the corresponding expressions are highlighted below and have been left general so as to also apply to the hidden layer. I_i denotes input (i.e., output from hidden node i) and W_{ji} its corresponding weight. i denotes the hidden node number and j the output node number into which it goes. Y_j denotes the output from a given output node. Sigma is the threshold of a neuron. The expressions are similar for the hidden layer, with weights W'_{ji} used in place of the previous weights.

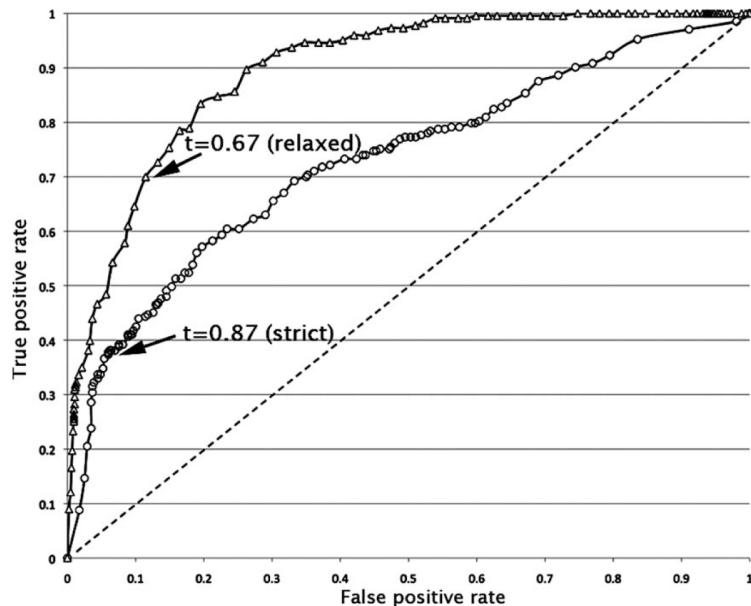


Figure 3. Receiver-operator characteristic plots for prediction performance. The black solid lines represent the receiver-operator curves for leave-one-out validation on the strict and relaxed databases. Statistics are provided in Table I.

Table I. Statistical Summary of Prediction Performance

Structures	Sensitivity	Specificity	Jack-knife correlation coefficient	Training correlation coefficient	Kink prediction	Nonkink prediction
Strict	0.46	0.99	0.40	0.57	96%	78%
Relaxed	0.70	0.89	0.56	—	62%	92%
PSIPRED relaxed	0.64	0.81	0.42	—	54%	86%

Prediction performance was assessed using the leave-one-out method for the strict set of kink and nonkink structures as well as for the relaxed set of kink and nonkink structures. The PSIPRED results are considered equivalent to a jack-knifed procedure because none of the proteins used in the current database were included in the original soluble protein training database.

nonkinked helices were correctly predicted as nonkinked (true negatives), and there were no false positives. Looking at the four kinks that were missed, it appears from Figure 4(A) that these are relatively subtle bends and may have less distinctive sequence signatures. For the structure 1H2S [Fig. 4(B)], two kinks were correctly identified, one was missed, and

the algorithm predicts kinks at five positions that are not kinked. The seventh helix illustrates an error in kink identification. We predict a kink at a point where the helix clearly breaks, but the bend angle does not change. Thus, we would argue that the algorithm actually predicted this deviation correctly, but our method of identifying true kinks in

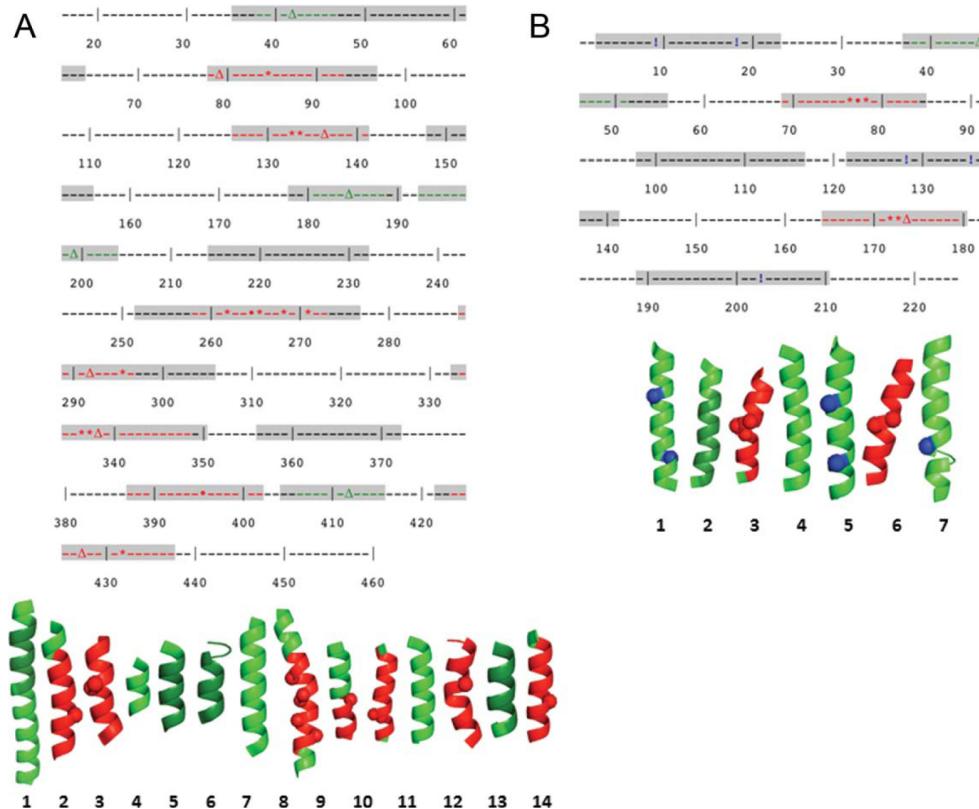


Figure 4. Examples of predictions. Illustration of kink prediction results using the relaxed database. (A) PDB code 1OTS, chain A and (B) PDB code 1H2S, chain A. The upper section of each figure illustrates the prediction results as a one-dimensional sequence. Transmembrane regions have been highlighted in gray. Regions highlighted in red correspond to correctly predicted regions (true positives), and blue exclamation points to incorrectly predicted positions (false positives). The maximum bend angle in each kinked region has been indicated by a Δ . The center of each kink prediction has been marked with a *, and overlap of a prediction center and the maximum bend angle has been marked with a ●. Kinked regions not predicted have been highlighted in green (false negatives). The bottom of the figure shows the individual transmembrane helices. The coloring scheme is the same as for the upper figure. The correctly predicted kinks are highlighted in red, and the center of each kink prediction is indicated by a red ball at the $C\alpha$ position. False-negative predictions are highlighted in dark green. False-positive predictions are indicated by a blue ball at the $C\alpha$ position.

structures is not foolproof. The one missed kink and five mispredicted kinks that are not in fact kinked may be attributed to the fact that 1H2S chain A had only 118 homologs compared to other structures that often had thousands of homologs.

Conclusion

Our results indicate that there are significant sequence differences between kinked and nonkinked transmembrane helices and that these differences can be exploited to predict kinks. This does not mean that kinks are generated only by local sequence, however, as we cannot necessarily link cause and effect. For example, certain residue types may provide stronger long-range contacts that can distort helices. Alternatively, if kink occurrence is more frequent at functional sites, the residue preferences could reflect the likelihood of being in functional sites rather than kinks. It is likely that prediction results could be greatly improved with more structures. Sequence preferences other than proline are much more subtle and seem to vary with database construction. Some of the more subtle sequence pattern information is likely masked because of the limited number of kink classifications we can use. Clearly not all helix distortions are the same, but because of the small dataset, we used a simple binary classification of kinked or nonkinked based solely on bend angle. Ideally, one would like to parse helix bends into distinct classes as the sequence signatures may be quite different. In spite of these limitations, we have been able to develop a prediction method that provides useful information that can be incorporated into helix packing algorithms.^{8,11,24} Kink prediction will only improve in accuracy and refinement as more structures become available.

Methods

Structure database

A nonredundant database of membrane proteins was used as the source from which all kinks and nonkinks were identified. None of the sequences shared more than 30% sequence identity, and only structures solved at a resolution of 2.5 Å or better resolution were retained. This resulted in a total of 41 membrane protein structures: 1C3W, 1EHK, 1H2S, 1JB0, 1K4C, 1KB9, 1KQF, 1NKZ, 1OKC, 1OTS, 1Q16, 1SU4, 1U19, 1V54, 1XIO, 1XQF, 1Z98, 2A65, 2AHY, 2BHW, 2BL2, 2BS2, 2F2B, 2FBW, 2J58, 2J7A, 2J8C, 2NQ2, 2RH1, 2UUI, 2VPZ, 2W2E, 2WGM, 2Z73, 2ZZE, 3B45, 3B9W, 3BKD, 3C02, 3DDL, and 3KCU.

Kinked and nonkinked transmembrane helix identification

Kink and nonkink identification was performed differently for training and for validation purposes. To ensure that we only trained on the clearest, most well-

defined kinks, we were particularly strict in our training set definitions. Subsequently, to more accurately assess our prediction performance, we relaxed the criteria to identify all genuine kinks and nonkinks. The strict and relaxed criteria are described below.

Kinked and straight helices were identified by measuring bend angles of nine-residue stretches of transmembrane helices. Only unique chains for each structure were used. For each chain, the transmembrane regions defined in the Orientations of Proteins in Membranes (OPM) database²⁵ were used as a starting point. The transmembrane region was then extended to the end of the helix if the region was defined as helical by DSSP.²⁶ These extended helices constituted the template from which the nine-residue segments were extracted, although only segments with a center residue within at least one residue of the transmembrane region (as defined by OPM) were retained. Kinks and nonkinks for transmembrane helical regions were identified using bend angles measured using the ProKink plug-in available for Simulaid.²⁷

To extract kinked segments using strict criteria for training purposes, the nine-residue segments were filtered using several steps. First, the bend angle of all segments was sorted from largest to smallest. Beginning with the largest bend angle segment, nearby segments were eliminated if they overlapped. This process was repeated for every remaining structure in order of decreasing bend angle. Those segments that remained were then filtered using two criteria. The average bend angle over four neighboring sliding windows was required to be greater than (or equal to) 13°, or the bend angle for any individual window was required to be greater than 24°. If only three or fewer neighboring windows were available because they were at a helix end, then any candidate with a bend angle of 13° or less was thrown out. Ultimately, using the strict criteria, 323 kinked structures were obtained.

To identify clear nonkinked segments, we also employed strict criteria. First, the bend angle of all segments was sorted from smallest to largest. Any window exceeding 20° in bend angle was immediately discarded. Beginning with the smallest bend angle segment, nearby segments were eliminated if they overlapped. This process was repeated for every remaining structure in order of increasing bend angle. Those structures that remained were then filtered using two criteria: the average bend angle over four neighboring segments was required to be less than 8°, and the bend angle for every individual segment had to be less than 20°. In this manner, 567 nonkinked segments were obtained.

For prediction assessment we employed relaxed criteria that identified more visually obvious bends. A kink was identified as an authentic kink if the average bend angle over three neighboring windows

was greater than (or equal to) 10° or the bend angle of an individual window was greater than 18°. A total of 2048 relaxed kink windows were extracted. This does not mean that there are 2048 kinks because overlapping windows can contain the same kink. We wanted to identify all kink windows for assessment of predictions because all possible windows are tested in kink prediction. Nonkink regions were identified as all regions other than the identified kink regions above.

Family building

Similar sequences in RefSeq's nonredundant protein sequence database²⁸ were identified using PSIBLAST with one iteration, and a *P*-value cutoff of 1e – 10 to ensure close similarity between the known structure and its sequence homologs. The number of sequences identified was limited to 30,000. Any aligned sequences with an alignment length less than 70% of the length of the original sequence were discarded. All nonnative gapped regions were discarded.

To count amino acid frequencies at each position, the counts were weighted by (1-% identity) to incorporate some information about the likelihood a residue might have changed in the homologous sequence (% identity refers to the whole protein). Thus, when a homolog sequence is 100% identical to the sequence of the known structure, the new amino acid is not counted. In this manner, close homologs do not overweight the counting statistics.

For input to the neural network, each residue at a given position of the original candidate sequence was added at full weight a fixed number of times per homolog. The number of times was optimized to maximize network performance; this value was 2.

This weighting scheme is described by the following equation for the 180 inputs corresponding to the 9 positions × 20 amino acids:

$$I(\text{res}_i, \text{pos}_j) = \frac{\sum_{\text{all homologs}} [(1 - \% \text{ identity})S(\text{res}_i) + \lambda]}{\sum_{N_{\text{pos}_j}} [(1 - \% \text{ identity}) + \lambda]}$$

$$S(\text{res}_i) = \begin{cases} 1, & \text{if } \text{res}_i = \text{res}_i \\ 0, & \text{if } \text{res}_i \neq \text{res}_i \end{cases}$$

$$\lambda = \begin{cases} 2, & \text{if } \text{res}_i = \text{res}_i \text{ in known structure} \\ 0, & \text{if } \text{res}_i \neq \text{res}_i \text{ in known structure} \end{cases}$$

$$N_{\text{pos}_j} = \# \text{ residues at position } j, \text{ excluding gaps.}$$

Use of sequence conservation

Sequence conservation scores were used as inputs to the neural network in addition to the primary sequence. Conservation scores were calculated for the nine-residue segment belonging to each candidate sequence window. This was accomplished by using the PSIBLAST data produced by the family

building step described above. From all pairwise alignments for each PDB/chain, 3000 were randomly selected and conservation scores calculated using SCORECONS with the Trident scoring method.²⁹ For this purpose, ClustalW-formatted pseudo-alignments were constructed.³⁰ Positions with excessive gaps were assigned a score of 0.

Neural network design

A feedforward neural network consisting of an input, hidden, and output layer was constructed. There were 9 (positions) × 20 (amino acids) = 180 inputs to the network related to the fractional occurrence of all possible residue/position combinations (as described by the input equation above). In addition, we used nine inputs corresponding to the conservation scores of the nine residues in the native sequence. Two sets of weight matrices were evolved through learning: one between the input and hidden layers and the other between the hidden and output layers. Weights were randomly assigned to values between –0.5 and 0.5 to start. The number of hidden nodes was optimized according to network performance, and the best performance was found to occur using five hidden nodes. The output layer is governed by the following equations, which have been left general so as to also apply to the hidden layer. I_i denotes the input (i.e., output from hidden node i) and W_{ji} its corresponding weight. i denotes the hidden node number (e.g., 3) and j the output node number into which it goes. Y_j denotes the output from a given output node. Sigma is the threshold of a neuron and was set to 0.5²¹.

$$X_j = \sum_i W_{ji} I_i$$

$$Y_j = \frac{1}{1 + e^{-(X_j - \sigma)}}.$$

The expressions are similar for the hidden layer, with weights W_{ji} used in place of the previous weights. For each cycle, the weight matrices were adjusted once for each example in the training set. Training was performed over 1000 cycles. The order of training examples during weight adjustments was shuffled every cycle to reduce any noise due to ordering.

Weight adjustment was performed in the following fashion. The output layer consisted of a single node predicting whether a candidate sequence was a kink (maximum output value of 1) or a nonkink (minimum output value of 0). The error in this output value for each training example was computed as follows:

$$\delta = D - Y,$$

where D is the correct value and Y is the predicted value.²¹ Adjustments to the weight matrix between

the hidden and output layers were made according to the expression:

$$W_{ji}(t+1) = W_{ji}(t) + \eta \delta_j I_i,$$

$$\delta_j = \delta,$$

where η is the learning rate. The optimal learning rate was found to be 0.1 (according to how far training progressed). The weight matrix between the input and hidden layers was adjusted by first back propagating the error according to the expression²¹:

$$\delta'_i = \sum_j W_{ji} \delta.$$

The weight layer adjustments were then made using the $W_{ji}(t+1)$ expression above but with weights W'_{ji} used in place of the previous weights, error δ'_i in place of the previous error, and inputs to the appropriate layer.

Early stopping was used to prevent overtraining. To find the point at which overtraining occurs, we evaluated the network every 10th cycle (of 1000 cycles) for its ability to predict structures in a test set that was not used for training. The test set consisted of 50 kinked and 50 nonkinked structures. The total error over all test set examples was computed as follows:

$$\delta_T = \sum_{\text{all examples in test set}} \delta.$$

The training network for which the lowest total error occurred on the test set was taken as the best network.²²

Training and test set creation

To ensure no bias in the datasets used for training and early stopping, kink and nonkink selection was randomized. Test set kink selection was done by selecting 50 structures at random from the total number of identified strict kinks. Similarly, test set nonkink selection was done by selecting 50 structures at random from the total number of identified strict nonkinks. These two groups became the test set used for early stopping. The remaining strict structures (273 kinks and 517 nonkinks) were retained for training of the neural network.

Evaluation

Because of limited data for membrane proteins, we chose to use the jack-knife or leave-one-out method to evaluate our strict and relaxed predictions. The threshold for each database (strict or relaxed) was chosen using a receiver-operator plot where the Matthews correlation coefficient was maximized (see Fig. 3). Relaxed predictions were done across all possible protein windows in transmembrane segments.

Relaxed evaluation was done using the jack-knife approach for trained examples from the strict database and the best network (i.e., one with lowest total error) for all other structures. Assessment of kinks and nonkinks using the relaxed criteria was done separately. Because it is difficult to unambiguously separate the transition region between kinked and nonkinked regions, a buffer region of four residues to the left and right of the original kinked region centers was created. We only evaluated nonkinked region centers that were more than four intervening residues from this unambiguously defined transition kink area. These residue separations were arrived at after visual evaluation of many structures.

Kinked regions that passed the relaxed criteria were merged together to form contiguous regions when separated by one residue or less. These merged regions defined the contiguous kink regions that would be checked to see if they contained predicted kinks or not. If a contiguous kinked region contained a single kink prediction, it was recorded as a single true positive. If a contiguous kinked region did not contain a single kink prediction, it was recorded as a single false negative.

Nonkinks were evaluated differently. When evaluating a nonkink region, every single nine-residue window contained within the contiguous region was evaluated individually (rather than as a continuous region). If a given window was correctly predicted as a nonkink, it was recorded as a single true negative. If it was incorrectly predicted as a kink, it was recorded as a single false positive. All of these true and false negatives and positives were summed over all predictions. Predictions using the secondary structure prediction algorithm PSIPRED²³ were evaluated in the same way except that predicted coils or strands were defined as kinks. As with kink prediction, a nine-residue window around the predicted coil or strand was defined as the kink region.

Website predictor

De novo prediction by our web application available online is done solely using the primary amino acid sequence of entire proteins. The transmembrane helical regions can be input manually or determined automatically by Proteus²⁴.

We ran three trials of our neural network using five hidden nodes. This resulted in jack-knife networks with correlation coefficients of 0.39, 0.38, and 0.40. We chose to use the network with the highest correlation coefficient. We looked at all networks making up this jack-knife network and retained the network with the lowest total error when early stopped using the test set. This network was installed for use in online predictions.

The network threshold for online predictions was chosen to be that determined by the receiver-operator curve analysis for the relaxed database ($t =$

0.67). This was done because the relaxed database most closely mirrors practical usage of the prediction software on entire proteins.

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Chapter 3

Structural differences between mesophilic and thermophilic membrane proteins

Introduction

Protein thermostability remains a focal point of interest for protein scientists¹.

Thermophiles, in contrast to mesophiles, evolved to survive at higher temperatures, ranging from 45 to 122° C². As a result of their greater thermostability, thermophiles are of great interest to those studying how to improve protein thermostability. Knowledge gained can be applied towards protein engineering.

The differences in thermostability between mesophilic and thermophilic soluble proteins have been extensively studied. Properties studied include packing density, burial fraction, hydrogen bonding, ion pairs, and length of surface loops^{1,3–7}. The only general rule has been that the number of ion pairs increases with rising, optimal growth temperature⁸.

Several groups argue that protein thermostabilization is achieved by different mechanisms for different groups of proteins^{8,9}. Berezovsky and Shakhnovich argue that for soluble proteins, there are two major physical mechanisms of thermostabilization: structure-based and sequence-based⁹. Some hyperthermophilic proteins are described as being more structurally compact than their mesophilic homologues. Other thermophilic proteins retain similar structure but differ in a small number of strong, sequence-based interactions. The strategy adopted by a given protein is thought to depend on its evolutionary origin. Organisms that began life in an extreme environment such as hyperthermophilic archaea have proteins that are structurally more compact.

Organisms that began as mesophiles but migrated into a warmer environment adopted a sequence-based evolutionary strategy based on a few strong interactions⁹.

The vast majority of thermostability studies comparing mesophilic and thermophilic proteins have been conducted on soluble proteins. This has largely been because of an abundance of structural data for these proteins. While no comparable number of structures exist for membrane proteins, approximately 60% of approved drugs targets are membrane proteins¹¹. Understanding the thermostability of membrane proteins is thus of critical importance to lowering their crystallization barrier and increasing the amount of structural data available for study¹².

Membrane proteins differ from soluble proteins in many ways. Soluble proteins exist in an aqueous environment whereas membrane proteins are surrounded by phospholipid bilayers. To interface with such an environment, membrane proteins have adapted a much greater abundance of hydrophobic residues and far fewer polar or charged residues. As a result of these differences, the same questions investigated to study thermostability in soluble proteins may lead to different answers when looking at membrane proteins.

While there are many factors that may contribute to thermostability in membrane proteins, I investigated burial and packing differences between mesophiles and thermophiles. Packing differences may explain the thermostability difference between mesophilic and thermophilic membrane proteins. Large scale studies using non-

redundant soluble proteins have shown that there is no difference in the packing of soluble proteins⁶. Membrane protein packing is different from soluble protein packing^{13–16}, so thermophilic adaptation may be different. Some groups have found there to be greater packing in membrane proteins than in soluble proteins^{13,14}. Faham *et al.* has argued that packing is the dominant force driving membrane protein folding¹⁶. And Hildebrand *et al.* found lesser packing in membrane proteins than in soluble proteins¹⁵; Adamian and Liang have found more cavities in membrane proteins to explain this¹⁷.

To study thermostability in membrane proteins, a non-redundant database was created consisting of alpha-helical membrane proteins to examine the differences between mesophiles and thermophiles. Structurally analogous pairs of mesophiles and thermophiles were identified using structure and sequence information. Using these pairs and the database, burial and packing density were evaluated to see if they helped explain the difference in thermostability between thermophiles and mesophiles.

Results and Discussion

Mesophile-thermophile database construction

To explore the structural differences between mesophiles and thermophiles that may contribute to thermostability of membrane proteins, I constructed a library of mesophiles and thermophiles from a database of 126 non-homologous, high-resolution alpha-helical membrane protein structures. Using optimal growth temperature data, these structures were divided into 101 mesophiles and 25 thermophiles (list provided in Methods). To reduce biases further in looking at properties of membrane proteins, only unique chains (as described in Methods) were used. Prosthetic groups were carefully included and author-recommended oligomer forms were used. Burial and packing were studied within the context of this database. In addition, a database of structurally similar pairs of mesophiles and thermophiles was created (as described in Methods). Differences found using the unpaired structure database may be due to variation in function among the proteins studied. Structurally similar pairs are more likely to share function. Thus, differences found using the paired structure database are more likely to explain differences in thermostability. A difference observed in both the unpaired and paired structure databases is more likely to explain a difference in thermostability. Lastly, those proteins with a greater number of residues were weighed more heavily as has been described in the Methods. The rationale was that a protein with more residues contributes more data to analyze than one with fewer, and thus should have greater weight.

Burial in mesophiles and thermophiles

Burial characterizes how exposed a residue to what is outside of a protein such as solvents and substrates. I divided analysis of burial values into analysis of soluble regions and analysis of transmembrane regions. Soluble regions of membrane proteins are the closest relative to regions of soluble proteins while transmembrane regions are a region that might be expected to differ due to the environment in which membrane proteins reside.

Burial values were calculated for mesophiles and thermophiles in the unpaired and paired structure databases, and tabulated in Table 3-1 below. Little or no difference in burial values was found in four comparisons:

- (1) soluble regions of unpaired mesophiles and thermophiles
- (2) transmembrane regions of unpaired mesophiles and thermophiles
- (3) soluble regions of paired mesophiles and thermophiles
- (4) transmembrane regions of paired mesophiles and thermophiles

The distributions of these values were all found to be approximately normal, with no outliers.

Table 3-1: Burial and packing in mesophiles and thermophiles

Measurement		Soluble	Transmembrane
Burial	Unpaired ratio	1.02±0.03 ¹	1.1±0.1 ¹
	Paired difference	0.003±0.004 ²	0.007±0.008 ²
Packing	Unpaired ratio	1.006±0.005 ¹	1.008±0.005 ¹
	Paired difference	0.004±0.001 ²	0.001±0.002 ²

¹ mean burial or packing ratio (mesophiles/thermophiles) with standard error

² mean burial or packing difference (mesophile-thermophile) with standard error

An unpaired ratio of 1 indicates no burial or packing preference when mesophiles are compared to thermophiles. A paired difference of 0 indicates no burial or packing difference between paired mesophiles and thermophiles.

Packing in mesophiles and thermophiles

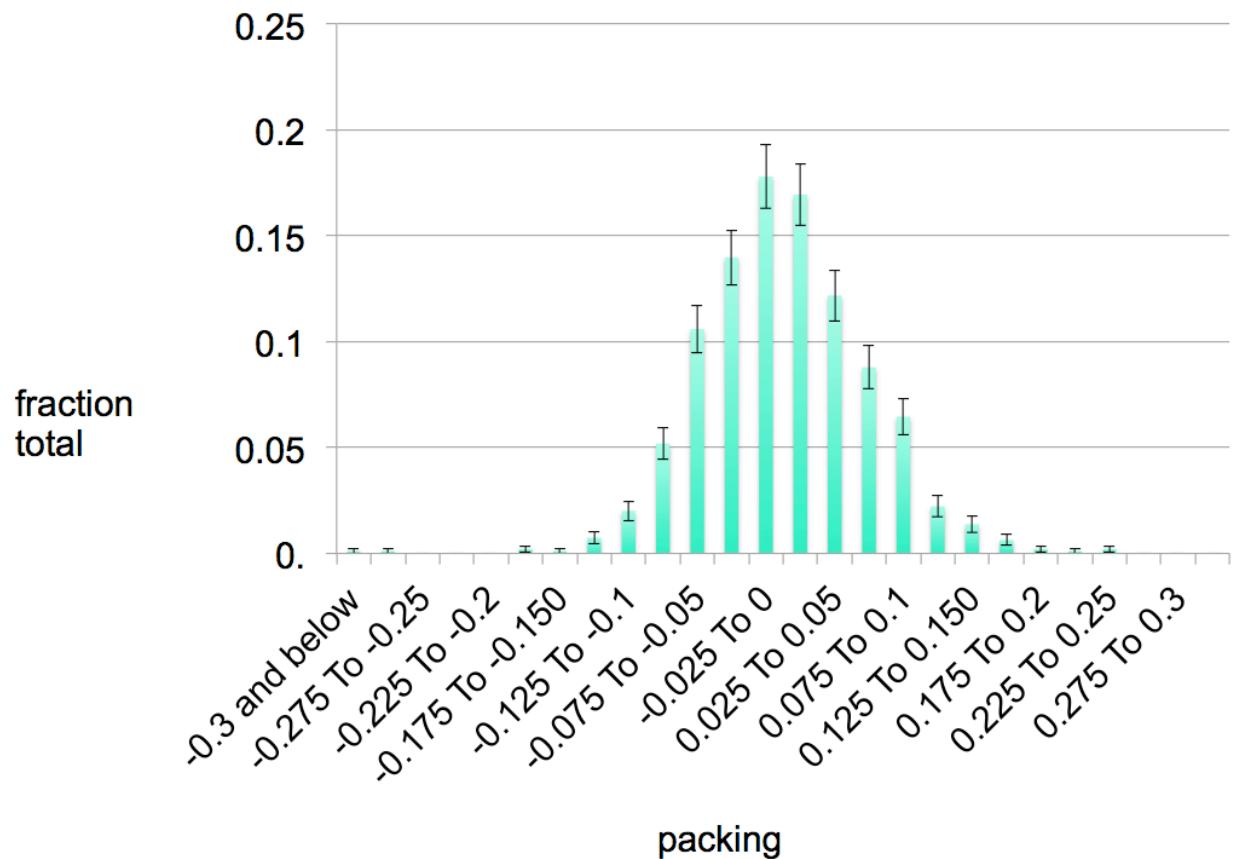
Similar to the analysis for burial values, the analysis of packing values was divided into soluble region and transmembrane region analyses. Packing describes how many atoms fit into a fixed volume. Packing values were calculated for mesophiles and thermophiles in the unpaired and paired structure databases, and tabulated in Table 3-1 above. Little or no difference in packing values was found in four comparisons:

- (1) soluble regions of unpaired mesophiles and thermophiles
- (2) transmembrane regions of unpaired mesophiles and thermophiles
- (3) soluble regions of paired mesophiles and thermophiles
- (4) transmembrane regions of paired mesophiles and thermophiles

The distributions of these values were all found to be approximately normal. The transmembrane pair packing distribution, in addition to having a normal distribution, also had 2 outliers (shown in Figure 3-1 below). These outliers were explained by too short alignment lengths (31 and 28 residues). With such short alignment lengths, the

likelihood of alignment of one protein to any other protein becomes extremely high (e.g., one alpha helix from one protein to the alpha helix of another protein) and packing differences lose significance. As a result, these outliers were eliminated from analysis.

Figure 3-1: Average packing difference distribution for transmembrane regions of mesophile-thermophile pairs



The ‘packing’ axis describes the distribution of the difference in average packing density for every mesophile-thermophile protein pair. The ‘fraction total’ axis describes the fraction of protein pairs found in each packing value difference bin.

It is surprising that both membrane and soluble proteins have no packing or burial differences despite their vastly different environments⁶ and the differing role of packing in membrane proteins^{13–16}. Looking solely at burial and packing values, soluble and

membrane proteins appear closer cousins than previously thought^{13–16}. However, these results are in agreement with previous work by Joh *et al.*, who found little difference in packing energetics of membrane and water soluble proteins¹⁸.

With no burial or packing differences, the question arises as to what the thermostability differences can be attributed to. The structure of membrane proteins is dominated by alpha helices and hydrogen bonding to stabilize such helices. As a result, one might expect differences in thermostability to arise from differences in interhelical hydrogen bonding or other stability adaptations such as kinking of helices. Ions pairs may be another contributor to thermostability as these have also been found in membrane proteins^{19–21}. In addition, differences within soluble regions of membrane proteins such as surface loop length may be distinguishing features of thermophiles from mesophiles. Lastly, structural pairs may reveal sequence determinants of stability differences that have not previously been apparent from pure sequence analysis. These parameters have not been studied for mesophiles and thermophiles in membrane proteins and are an active area of exploration. As additional membrane protein structures are deposited, it may become clear that several different mechanisms may have evolved to explain differences in thermostability for different groups of membrane proteins as has been the case for soluble proteins^{8,9}. For example, the thermostability of hyperthermophiles may be explained by structural differences between hyperthermophiles and mesophiles. And the thermostability of thermophiles may be explained by sequence differences between thermophiles and mesophiles.

Conclusion

Even though membrane proteins differ from soluble proteins in many fundamental ways such as an abundance of hydrophobic residues and the role of packing^{13–16}, burial and packing values appear to be shared between mesophiles and thermophiles in both types of proteins. The addition of new non-redundant structures to the existing membrane protein database in the future may modify this picture, but the present data unambiguously suggest this conclusion. Little or no difference in burial or packing values exist between mesophiles and thermophiles in membrane and soluble proteins. One question this observation raises is what can explain the thermostability differences between mesophiles and thermophiles in membrane proteins. The only general pattern for soluble proteins relates the number of ion pairs and rising growth temperature⁸. For membrane proteins, many properties such as surface loop length, interhelical hydrogen bonding, kinking of helices, ion pairs, and sequence determinants remain to be explored. Ultimately, as for soluble proteins, a combination of different mechanisms may have evolved and will have to be investigated in the future^{8,9}.

Methods

Structure database

A nonredundant database of mesophiles and thermophiles was created using alpha-helical membrane proteins from Stephen White's database (<http://blanco.biomol.uci.edu/mpstruc/listAll/list>)¹⁰. Growth temperatures were obtained from the American Type Culture Collection (<http://www.atcc.org/ATCCAdvancedCatalogSearch/AllCollectionSearch/tabcid/454/Default.aspx>) and additional papers from the literature. Structures were divided into mesophiles and thermophiles based on a 50 °C cutoff; those greater than or equal to this cutoff were considered thermophiles.

Each group of structures (mesophiles, thermophiles) was made non-redundant separately. To reduce each set of structures to a non-redundant set, a random structure was chosen and its sequence was blasted against that of all other structures. If the random structure's sequence had a similarity expectation value of less than 1e-10, the structure was removed. This was repeated for randomly chosen structures until the number of member structures did not change for 20 iterations.

For each structure, the author-recommended oligomer was chosen as described by REMARK 350 in each Protein Data Bank (PDB) file. To further reduce the dataset to a meaningful one for burial and packing analysis, a number of additional requirements were enforced for member structures. Only unique chains (i.e., non-identical) were used for analysis. Nuclear magnetic resonance (NMR) and electron microscope (EM)

structures were eliminated. For each oligomer, three or more transmembrane helices were required to retain it in the structure database; those with fewer transmembrane helices were thought to contain insufficient data to perform transmembrane packing and burial analysis. Structures lacking membrane protein topology when examined by eye were removed. The prosthetic groups for each structure were carefully examined and retained in the oligomeric structures so that they would be considered for burial and packing analysis. Lipids and other non-intrinsic components were removed from each structure for consistency in analysis. The members making up the structure database were as follows.

Mesophiles (101): 1AIJ, 1BCC, 1BGY, 1COW, 1EFR, 1FFT, 1KPL, 1KQF, 1L0V, 1LGH, 1NEK, 1NKZ, 1PW4, 1Q16, 1Q90, 1QO1, 1U7G, 1V54, 1YEW, 1ZCD, 1ZOY, 1ZRT, 2A79, 2ACZ, 2BHW, 2BL2, 2BS2, 2C3E, 2CK3, 2E74, 2EI4, 2EXW, 2GMH, 2GSM, 2H8A, 2I37, 2J58, 2JLN, 2NUU, 2O01, 2O7L, 2OAR, 2OAU, 2Q7M, 2QCU, 2QI9, 2QKS, 2QTS, 2RCR, 2RDD, 2V8N, 2VQG, 2W5J, 2WCD, 2WSS, 2WSW, 2X2V, 2XUT, 2YFY, 2ZUQ, 2ZW3, 2ZXE, 3B8E, 3B9W, 3B9Y, 3BF0, 3BKD, 3CHX, 3CK6, 3CX5, 3D31, 3DQB, 3EAR, 3EHZ, 3FH6, 3G5U, 3H90, 3HD6, 3JQO, 3K07, 3KCU, 3KG2, 3KP9, 3LIM, 3LRB, 3LUT, 3M71, 3MK7, 3MKT, 3NAF, 3NVO, 3O7Q, 3OAA, 3P0G, 3PJZ, 3QAK, 3QE7, 3QNQ, 3RLB, 3RVY, and 3S0X

Thermophiles (25): 1EHK, 1JB0, 1ORQ, 2A0L, 2B2F, 2F2B, 2HN2, 2ONK, 2QJU, 2VPZ, 2ZY9, 3A5C, 3AQP, 3ARC, 3B4R, 3BVD, 3DIN, 3GIA, 3HYW, 3K5B, 3KBC, 3KDS, 3M9S, 3MP7, and 3ORG

Transmembrane region assignment

Transmembrane regions used for analysis were determined using the unpublished work of Seong-Kyu Han described here: Transmembrane regions were obtained from a consensus database constructed from the Protein Data Bank of Transmembrane Proteins (PDBTM) and Orientations of Proteins in Membranes (OPM) database, or using TMDET to identify transmembrane regions for those proteins not present in these databases^{22–24}.

Mesophile-thermophile structural pair assignment

Mesophile-thermophile structural pairs were identified by Seong-Kyu Han as has been described here: Structurally homologous pairs were identified using the structural alignment tool TM-align²⁵. Aligned residues were defined by those that showed greater than 1Å root-mean-square deviations of Cα atoms (Cα RMSD) since these were considered areas of interesting change between proteins with similar function. Pairs that did not match transmembrane (or soluble) regions between proteins were excluded from packing and burial analysis.

Burial calculation

Burial data was obtained by calculating relative solvent accessibilities per residue using a program called surfarea produced using the Ezprot library by Frank Pettit (<http://www.doe-mbi.ucla.edu/local/software/ezprot>); burial values resulted from subtracting the accessibility from 100. Results were cross-checked with another popular

program for burial calculations called naccess

(<http://www.bioinf.manchester.ac.uk/naccess/>). Burial values were calculated separately for soluble and transmembrane regions of mesophiles and thermophiles belonging to the database, as well as for the mesophile-thermophile structural pairs. For the database, values were averaged over all mesophiles and thermophiles separately. In computing the average, burial values were weighted by the number of residues in a given protein to obtain a weighted burial average over all proteins; this ensured that burial values from smaller proteins did not unnecessarily bias the results calculated over all proteins. For the structural pairs, difference values between each mesophile-thermophile pair were averaged over all mesophile-thermophile pairs.

Packing calculation

Packing values were calculated using Rother *et al.*'s Voronoia program for computing packing densities²⁶. Surface residues were excluded from the calculation since they are known to pack far less efficiently than buried residues¹⁵. A 0.20 Å grid distance and the ProtOr radii were chosen as parameters based on similar ones chosen by Hildebrand *et al.*¹⁵; the ProtOr radii was chosen since I was interested in an experimental rather than molecular dynamics application.

Packing values were calculated separately for soluble and transmembrane regions of mesophiles and thermophiles belonging to the database, as well as for the mesophile-thermophile structural pairs (as was done above for burial values). For the database, values were averaged over all mesophiles and thermophiles separately. In computing

the average, packing values were weighted by the number of residues in a given protein to obtain a weighted packing average over all proteins. For the structural pairs, difference values between each mesophile-thermophile pair were averaged over all mesophile-thermophile pairs; if either packing value for a pair was zero, this pair's data was excluded from the packing analysis.

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