SayaMatcher (狭山茶)

Saitama Medical School Research Center for Genomic Medicine



Hidemasa Bono

Division of Functional Genomics & Systems Medicine Research Center for Genomic Medicine

Saitama Medical School, JAPAN

http://kishoi.jp/SayaMatcher/

mailto: bono@saitama-med.ac.jp





Working both on bench and desk



I have studied tissue specific

RE/

genes, sat

Browse genes with tissue-specific expression

	<u>og spleen</u>	58 thymus	<u>os kidney</u>	10 heart				
	<u>07 brain</u>	15 cerebellum	12 lung	<u>13 liver</u>				
	65 cerebellum neonate10day	16 placenta	<u>17 testis</u>	83 uterus				
	18 pancreas	20 small_intestine	22 stomach	<u>90 colon</u>				
۱D	47 skin neonatotoday	98 <u>bone</u>	<u>xm muscle</u>	B4 adipose				
	Tissues colored by Tissue Ontology							
	Immune circulator neural germ digestive skin bone muscle adipose							

000

RikenExpression Array Database

READ: Log-transformed ratio data for in expression data

cloneid 09 58 06 10 12 13 07 15 65	5 16 17 83 18 20 22 90 47 98 xm 84			
1600013K070 0.10.10.20.20.10.60.40.3	3 201 0.2 0.6 0.4 0 0.2 0 0 0.3 0.8 0 TB5731 Mus musculus trophoblast specific protein alpha (Tpbpa), mRNA. [RTPS]			
5133400E0600.800.4000000.2	2 0.8 0.2 0.5 0.1 0.4 0 0.1 0 0.5 0.1 Unknown EST [phase1]			
1600012I11 0.2 1.4 0 0.1 0.5 0.1 0.2 0.1 0.2	2 0 0.10.10.3 0 0 0.20.2 0.7 0.2 Unknown EST [phase1]			
1600022H160.510.5000.30.200.1	1 0.3 1.7 0.5 0 0.3 0.1 0.2 0.3 0.2 0.5 TB6423 Mus musculus cathepsin J (Ctsj), mRNA. [RTPS]			
1600027L190.30.30.40.200.30.10.31.4	0.3 0.8 0.8 0.4 0 0.5 0.2 0.4 0.2 0 TB4512 Mus musculus prolactin-like protein C (Prlpc), mRNA. [RTPS]			
16000210090 0.11 0.10.20.40.10 0.2	2 0.1 0.3 1.2 0.4 0 0.1 0.1 0.4 0.3 0 TB5731 Mus musculus trophoblast specific protein alpha (Tpbpa), mRNA. [RTPS]			
1600019F100.20.20.20.400.10.30.20.7	7 0.3 0.2 0.4 0.1 0.3 0.3 0 0.2 0.3 0 0.2 0.3 0 similar to PROLACTIN-LIKE PROTEIN C BETA ISOFORM 1-A [Rattus norvegicus] [fantom]			
1600021M170 1.30.50 0 0.70.10 1.4	0.1 0.8 0.7 0.7 0 0.2 0 0.1 0 0.1 TF11211 similar to CEACAM11 [Mus musculus] [RTPS]			
1600019E220.40.60.90.3000.30.1000.2	2 0.3 1.4 0.5 0 0.6 0 0.3 0.7 0.9 0.3 Mm.43505 pregnancy-specific glycoprotein 28 [unigene] [phase1]			
1600027N170.60.60.30.20.60010.51.4	4 3 0.3 0.7 0.7 0.4 0.2 0.7 0 0 0.7 0.1 TF13031 hypothetical protein [RTPS]	_		
1600019K020 0.10.70 0.10.20 0.20.8	8			
1600025N01 0.2 1.3 0.3 0.5 0.1 0 0.2 0.2 0.8	8 0 0.70.90.4 0.6 0.2 0 0.3 0.6 0.1 pregnancy-specific glrcoprotein 21 [fantom]			
1620401J060.30.30 0.20.20 0.50 0.3	3 0 0 0.1 1.6 0.1 0.4 0.5 0.1 0 0.3 0.1 trophoblast specific protein alpha [fantom]			
1600017M180.2190.30.50.10.70.20.40.6	6 0 0.6 0.7 0.5 0.3 0 0.3 0.1 0 0.5 Mm.43505 pregnancy-specific glycoprotein 28 [unigene] [phase1]	Mm.43505 pregnancy-specific glycoprotein 28 [unigene] [phase1]		
1600025D070.40.20.60.200000.10.5	5 0 0.10.40.20.10 0.40.20.50 TB4361 Mus musculus chorionic somatomammotropin hormone 1 (Csh1), mRNA. [RTPS]	TB4361 Mus musculus chorionic somatomammotropin hormone 1 (Csh1), mRNA. [RTPS]		
16000290210.70.70.80.60.20 0.30.40.1	1 22 0.5 1.5 0.6 0 0.6 0.4 0 0.8 0.4 0.1 Mm.37203 chorionic somatomammotropin hormone 2 [unigene] [phase1]			
1600017E040.21.20.70.40.10.30.10.30.7	7 0.4 1.7 0.7 0.1 0.3 0.1 0 0.9 0.2 0 PROLACTIN-LIKE PROTEIN C [fantom]	PROLACTIN-LIKE PROTEIN C [fantom]		
1600029A130 1.20.70.70.70.50.20.30.6	6 - 0.4 1.3 0.6 0.6 0.7 0.3 0.2 0.5 0.3 0.1 Mm.46091 prolactin-like protein C 2 [unigene] [phase1]			
1600026F070.40.410.70.20.50.10.31.2	2 0.4 1.4 1 0.1 0.2 0.2 0.2 0.2 0 0.4 0 Unknown EST [phase1]	_		
3830418E20 1.1 0.1 0.8 0.9 0.4 0.6 0.3 0.2 0.3	3 0 0.6 1.1 0.2 0 0.1 0 0.3 1.2 0.3 trophoblast specific protein alpha [fantom]	_		
1620401H060.10.60.40.60.50.80.50.50.1	1 - 0.1 0.7 1.6 0.3 1 0.5 0.5 0.2 1.2 0.7 TB7972 Mus musculus cathepsin M (Ctsm), mRNA. [RTPS]			
1600019A080.71 0.20.40 0 0.10 1.3	3 0.5 0.6 0.9 0.1 0.3 0.4 0.3 0.3 1.2 0.2 TB5731 Mus musculus trophoblast specific protein alpha (Tpbpa), mRNA. [RTPS]			
1600021E160.400.50.200.30.10.30.6	6 0.5 0.7 0.2 0.1 0.2 0 0.1 0.7 0.5 TB6423 Mus musculus cathepsin J (Ctsj), mRNA. [RTPS]			
	1 0.3 0 0.2 0.1 0.1 0 0.1 0 1.2 0.1 TB9156 Mus musculus CE			
9430076M131 0.71.31 0.60.60.50.60.1		1		
1600016008 0.3 0.6 0 0 0.3 0.6 0.1 0.5 0	TB6362 Mouse pregnancy VICIC CXDICSSION DROILL OF			
1600016D150.30.21.2000.10.600.1	1 0 10 0 10 4 0 1 0 1 0 1 0 3 0 8 0 2 TB7972 Mus musculus cat			
1600020L08 0.8 0.2 0.4 0.2 0.3 0.2 0.2 0.2	0.1 0.4 1 0.2 0.3 0 0.1 0.1 0.6 0.2 prolactin-like protein F [f:			
1600029H120.10.91.20.40 0.30 0 1	10,70,10,10,10,00,00,00,00,00,00,00,00,00,00			
1600019K190.6 0.60.40.20.30 0.30.7				
1600027J170.60.81 0.30.10 0.20 0.2	2 0 0 1 0 1 0 1 0 1 0 1 0 1 0 1 0 1 0 1			
16000090200,10 0,30,40,10,50,40,10,5	5 0 0.50.50.5 0.40.407.0.70 similar to PROLACTIN-LIK b++ b++ b+ / / DEAD account loop in / fail			
1600021L090,6260,30,3000001006	A DID ZOZ DA DADIO A DE DE MM. 43505 pregnancy-spi IITTP://KEAD.GSC.FLKEN.JD/TQ	ITON		
5730550L030,40,50,90,20,50,20,50,30,5				



not so easy to interpret functional property of these to extract gene regulation network!



Depicting gene regulation networks from microarray data

- 1. Co-regulated genes
 - Any DNA motifs in upstream regions?
 - \Rightarrow hard to decipher them in mammal
- 2. Known transcription factor binding sites (TFBS)
 - Any features in gene expression profiles for genes with particular TFBS in upstream region?



What is SayaMatcher?

- System(pipeline) to get coordinates of transcription factor binding sites (TFBS) in the genome
 - TFBS pattern to be found is too short to blast/blat/ssaha, and have too many hits for genome sequence
 - Not interpretable only in the text output
 - export the annotation to genome browsers (Ensembl, UCSC)



Why is the pipeline needed?

- Genome sequences are updating
 - ⇒ not isolated task; searches need to be a 'pipeline '
- Too many iteration for one time job
 - All chromosome and its reverse complement
 - around 40 (around 20 x 2)
 - Species to be used (3-5)
 - Several kinds of NRE(around10)

Thousands of iterative operation needed for update



Methods for searching TFBS

- 1. Regular expression
 - When the 'consensus sequence' is reported
 - dreg, fuzznuc in EMBOSS
- 2. Position specific scoring matrix(PSSM)
 - 1 When consensus is weak, but multiple alignment is available
 - prophecy/profit in EMBOSS, HMMER

Saitama Medical School **Research Center for Genomic Medicine** Project Ensembl *e*! SayaMatcher Patterns of interest DAS (from biologist) → dreg · _DAS — candidate — HTML → fuzznuc Emboss BED Genome annotation profit Transcriptome prophecy annotation mfa UCSC Genome Browser (from papers)



Genes with predicted ERE in upstream region (-10kb)

000

http://10.53.95.9/~bono/ERE_human_10k.html

genes with ERE

Link to genome browser

x·18843340	-1	ENSG00000188310	\N
11:3399598	1	ENSG00000182139	N
11:5192437	-1	ENSG00000176742	Odorant receptor HOR3'beta1. [Source:SPTREMBL;Acc:Q9H2C8]
4:56728996	1	ENSG00000174799	\N
8:8132003	1	ENSG00000173295	\N
<u>8:11684731</u>	1	ENSG00000177907	\N
15:62106866	-1	ENSG00000166797	\N
<u>5:55043428</u>	1	ENSG00000152670	DEAD-box protein 4 (VASA homolog). [Source:SWISSPROT;Acc:Q9NQI0]
17:32838007	-1	ENSG00000108702	Small inducible cytokine Al precursor (CCL1) (T lymphocyte-secreted protein I-309). [Source:S
<u>15:73377925</u>	-1	ENSG00000140400	Alpha-mannosidase 2C1 (EC 3.2.1.24) (Alpha-D-mannoside mannohydrolase) (Mannosidase alpha cla
<u>15:76143692</u>	-1	ENSG00000136425	Kinase interacting protein 2 (KIP 2). [Source:SWISSPROT;Acc:075838]
11:128276628	-1	ENSG00000151704	ATP-sensitive inward rectifier potassium channel 1 (Potassium channel, inwardly rectifying, s
<u>16:30124321</u>	1	ENSG00000149923	Serine/threonine protein phosphatase 4 (EC 3.1.3.16) (Pp4) (Protein phosphatase X) (PP-X). [S
<u>17:67057071</u>	-1	ENSG00000070540	\N
<u>6:27875107</u>	1	ENSG00000124518	Histone H2A.c/d/i/n/p (H2A.1) (H2A/c) (H2A/d) (H2A/i) (H2A/n) (H2A/p) (H2A.1b). [Source:SWISS
<u>6:32026499</u>	1	ENSG00000166291	Helicase SKI2W (Helicase-like protein) (HLP). [Source:SWISSPROT;Acc:Q15477]
12:48484210	-1	ENSG00000167566	\N
12:53273432	-1	ENSG00000135447	Protein phosphatase inhibitor 1 (IPP-1) (I-1). [Source:SWISSPROT;Acc:Q13522]
12:54674057	1	ENSG00000139531	Sulfite oxidase, mitochondrial precursor (EC 1.8.3.1). [Source:SWISSPROT;Acc:P51687]
12:58484579	1	ENSG00000189370	\N
2:11686471	1	ENSG00000174934	GREB1 protein isoform a; gene regulated by estrogen in breast cancer protein. [Source:RefSeq;
12:94750169	1	ENSG00000139343	Small nuclear ribonucleoprotein F (snRNP-F) (Sm protein F) (Sm-F) (SmF). [Source:SWISSPROT;Ac
2:74014735	1	ENSG00000187833	\N
2:132402869	-1	ENSG00000182126	\N
20:10403543	1	ENSG00000149346	DJ1099D15.3.1 (Novel protein (Isoform 1)) (Fragment). [Source:SPTREMBL;Acc:Q9BR42]



.



Acknowledgement

- Grant-in-Aid for Scientific Research on Priority Areas (C) "Genome Science" from the Ministry of Education, Culture, Sports, Science and Technology of Japan
- Grant-in-Aid for Development of New Technology from The Promotion and Mutual Aid Corporation for Private Schools of Japan
 poster: D-4