

RNA-seq data in ArrayExpress and recent database improvements

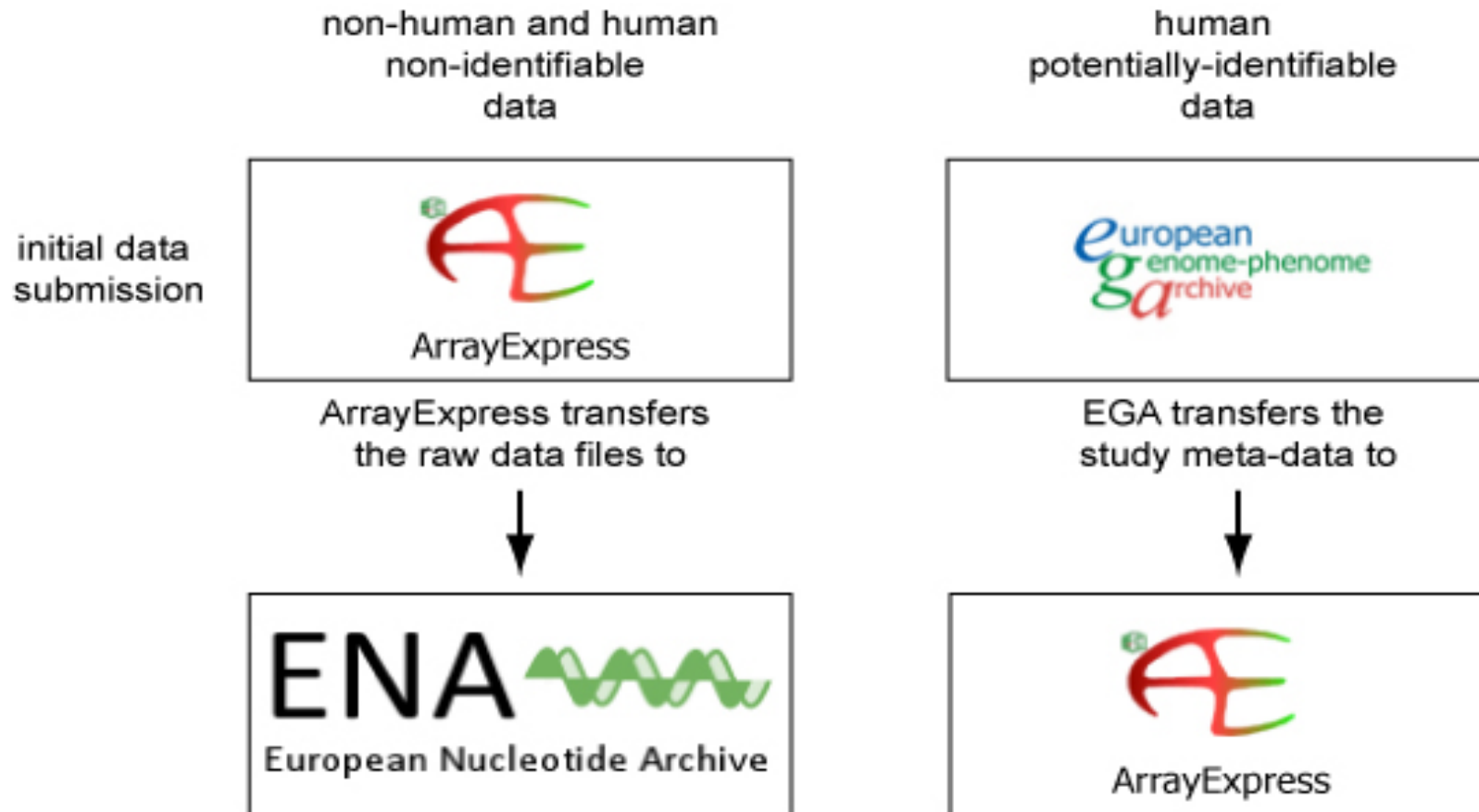
Gabriella Rustici
Functional Genomics Group



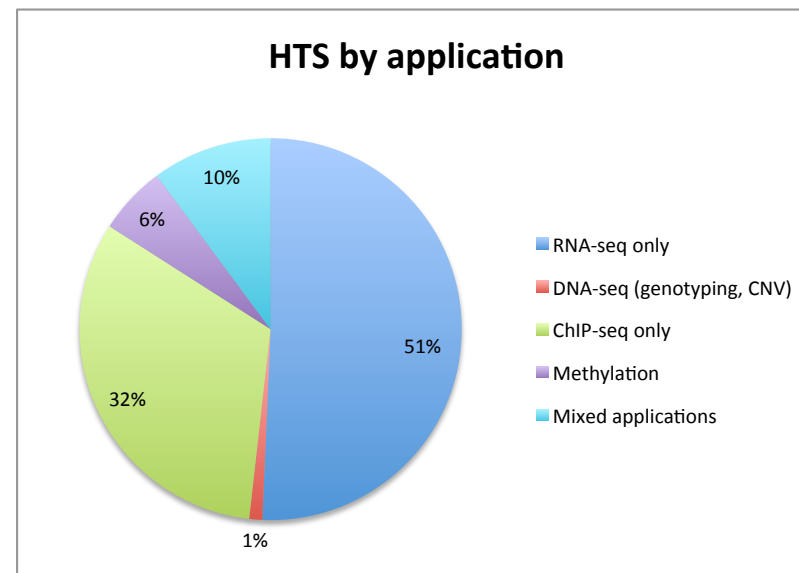
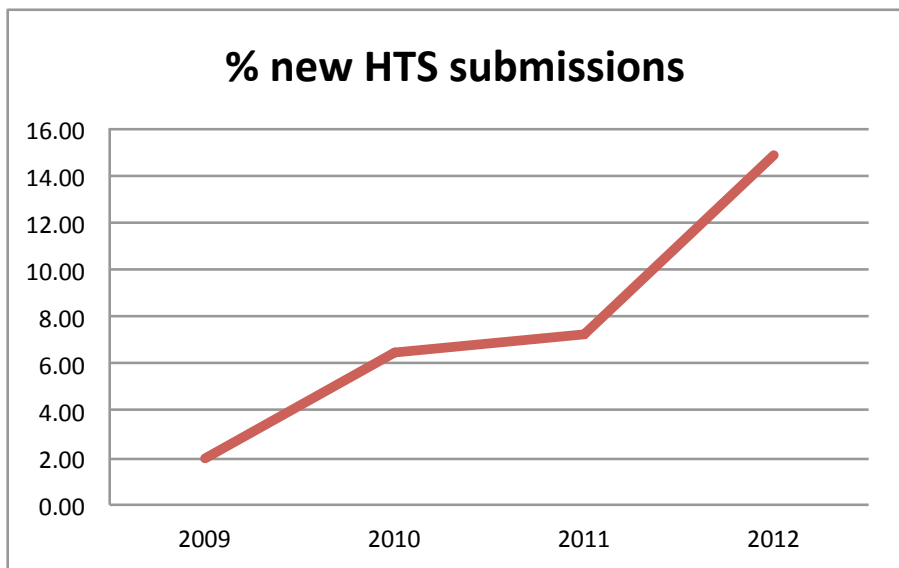
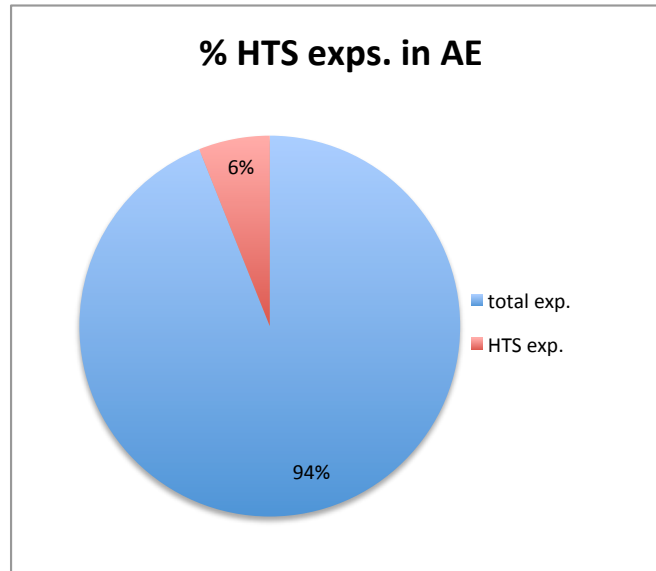
HTS data in ArrayExpress and Atlas

- We archive functional genomics data derived from HTS experiments in the [ArrayExpress Archive](#)
- Focus on RNA-seq and CHIP-seq experiments
- We are developing robust internal pipelines for RNA-seq data processing and quality assessment
- We display gene expression changes measured by RNA-seq in the [Expression Atlas](#)

Types of data that can be submitted to AE



Some stats



RNA-seq data in ArrayExpress

Experiment, citation, sample and factor annotations [clear] Filter on [reset] Display options [reset]

All species 25 experiments per page

ArrayExpress data only Advanced query syntax All arrays Detailed view

RNA assay by High-throughput sequencing Query

ID	Title	Assays	Species	Date	Processed	Raw	Atlas
E-MTAB-513	Illumina bodyMap2 transcriptome	48	Homo sapiens	2011-03-17	-		-
E-GEOD-27003	Deep Sequence Analysis of the Relationship between Gene Expression, CpG Island Methy	24	Homo sapiens	2011-03-16		-	-
E-MTAB-434	Transcription profiling by high throughput sequencing of zebra fish tissues and developm	7	Danio rerio	2011-03-11	-		-
E-GEOD-27837	Small RNA sequences from Schizosaccharomyces japonicus	2	Schizosaccharomyces japo	2011-03-09		-	-
E-GEOD-27699	Genome-wide discovery and analysis of microRNAs and other small RNAs from rice embr	2	Oryza sativa	2011-03-07			-
E-MTAB-533	GENCODE Targeted Sequencing Batch III	2	Caenorhabditis elegans, H	2011-03-01	-		-
E-MTAB-493	Transcription profiling by high throughput sequencing of Drosophila melanogaster tissues	4	Drosophila melanogaster	2011-02-28	-		-
E-MTAB-308	Zebrafish gene 3 prime end pull down for genome annotation	7	Danio rerio	2011-02-18	-		-
E-MTAB-475	Trypanosoma vivax Y486 bloodstream stage RNAseq	1	Trypanosoma vivax	2011-02-16	-		-
E-GEOD-27221	ENCODE Genome Institute of Singapore RNA-Seq	4	Homo sapiens	2011-02-14	-	-	-
E-MTAB-502	Citrobacter rodentium transcriptomics (Regulation)	2	Citrobacter rodentium	2011-02-12	-		-
E-GEOD-26230	Loqs-PD and R2D2 define independent pathways for RISC generation in Drosophila	4	Drosophila melanogaster	2011-02-10		-	-
E-MTAB-550	Toxoplasma gondii RNASeq	14	Toxoplasma gondii	2011-02-08	-		-
E-MTAB-549	Neospora caninum RNAseq	10	Neospora caninum	2011-02-08	-		-
E-GEOD-25132	[E-MTAB-75] Cryptic unstable transcripts in yeast	2	Saccharomyces cerevisiae	2011-02-05	-	-	-
E-GEOD-25003	[E-MTAB-5] Dynamic repertoire of a eukaryotic transcriptome surveyed at single nucleoti	31	Schizosaccharomyces pom	2011-02-05	-	-	-
E-GEOD-26136	Mammalian Dpy-30 regulates genomic H3K4 methylation and is essential for ES cell fate	10	Mus musculus	2011-02-01	-	-	-
E-GEOD-24938	Comparative transcriptomics of the model entomopathogenic fungi Metarhizium anisoplia	4	Metarhizium acridum, Met	2011-01-31		-	-
E-MTAB-460	Sanger_zebrafish_sequencing	8	Danio rerio	2011-01-28	-		-
E-GEOD-26444	Deep sequencing-based analysis of the anaerobic stimulon in Neisseria gonorrhoeae	4	Neisseria gonorrhoeae F62	2011-01-26		-	-
E-GEOD-26086	HIF-1synergizes with glucocorticoids to promote BFU-E progenitor self-renewal	9	Mus musculus	2011-01-22	-	-	-
E-GEOD-24788	Digital gene expression profiling for maize development - 2mm ear libraries, B73 vs. ram	7	Zea mays	2011-01-19		-	-
E-GEOD-24538	Ab initio identification of transcription start sites (TSSs) in the Rhesus macaque genome	3	Macaca mulatta	2011-01-19			-
E-GEOD-24284	Deep sequencing analysis of transcription-induced chimeras in human prostate adenocar	14	Homo sapiens	2011-01-19	-	-	-
E-GEOD-23776	Alternative expression analysis by RNA sequencing (comparison of 5-FU sensitive and res	2	Homo sapiens	2011-01-19			-

415 experiments, 5853 assays. Displaying experiments 1 to 25. Pages: 1 2 3 4 5 6 7 8 9 10 .. 17

MINSEQE score

Accession	Title	Assays	Species	Date	Processed	Raw	Atlas
E-MTAB-513	RNA-Seq of human individual tissues and mixture of 16 tissues (Illumina Body Map)	48	Homo sapiens	2011-05-05	-		-
Samples (19)	Click for detailed sample information and links to data						
Protocols (6)	Click for all experimental protocols						
Description	Illumina Human Body Map 2.0 Project. Transcription profiling by high throughput sequencing of individual and mixture of 16 human tissues RNA. Additional files available-E-MTAB-513.additional.zip						
Experiment types	transcription profiling by high throughput sequencing, co-expression, optimization, organism part comparison						
Contact	Gary P Schroth <gschroth@illumina.com>						
Citations	<p>Detection of Redundant Fusion Transcripts as Biomarkers or Disease-Specific Therapeutic Targets in Breast Cancer. Asmann, Yan W.; Necela, Brian M.; Kalari, Krishna R.; Hossain, Asif; Baker, Tiffany R.; Carr, Jennifer M.; Davis, Caroline; Getz, Julie E.; Hostetter, Galen; Li, Xing; McLaughlin, Sarah A.; Radisky, Derek C.; Schroth, Gary P.; Cunliffe, Heather E.; Perez, Edith A.; Thompson, E. Aubrey. <i>Cancer Res</i> 72(8):1921 (2012), UKPMC 22496456</p> <p>The GENCODE v7 catalog of human long noncoding RNAs: Analysis of their gene structure, evolution, and expression. Thomas Derrien; Rory Johnson; Giovanni Bussotti; Andrea Tanzer; Sarah Djebali; Hagen Tilgner; Gregory Guernec; David Martin; Angelika Merkel; David G. Knowles; Julien Lagarde; Lavanya Veeravalli; Xiaolan Ruan; Yijun Ruan; Timo Lassmann; Piero Carninci; James B. Brown; Leonard Lipovich; Jose M. Gonzalez; Mark Thomas; Carrie A. Davis; Ramin Shiekhattar; Thomas R. Gingeras; Tim J. Hubbard; Cedric Notredame; Jennifer Harrow; Roderic Guigó. <i>Genome Res</i> 22(9):1775 (2012). UKPMC 22955988</p>						
MINSEQE				-			
	Exp. design	Protocols	Factors	Processed	Seq. reads		
Files	Investigation Description		E-MTAB-513.idf.txt				
	Sample and Data Relationship		E-MTAB-513.sdrf.txt				
	Browse all available files						
Links	ENA - ERP000546 Send E-MTAB-513 data to GENOMESPACE						
Status	Released on 5 May 2011, last updated on 16 February 2012						

1 experiment, 48 assays.

MINSEQE guidelines

- Minimal Information about a high-throughput Nucleotide SEQuencing Experiment
- The proposed guidelines for MINSEQE are:
 1. General information about the experiment and sample-data relationships
 2. The description of the biological system, samples, and the experimental variables being studied
 3. Essential experimental and data processing protocols
 4. The sequence read data for each assay
 5. The processed data for the set of assays in the study
- For the last 2 years, MINSEQE scores are no worse than MIAME scores. There may be differences between HTS and microarray submissions in terms of what is absent (e.g., processed data for HTS).

Sample-data relationship file





































Accession	Title	Assays	Species	Date	Processed	Raw	Atlas
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MINSEQE				-			
	Exp. design	Protocols	Factors	Processed	Seq. reads		
Files	Investigation Description		E-MTAB-513.idf.txt				
	Sample and Data Relationship		E-MTAB-513.sdrf.txt				
	Browse all available files						
Links	ENA - ERP000546 Send E-MTAB-513 data to GENOMESPACE						
Status	Released on 5 May 2011, last updated on 16 February 2012						

1 experiment, 48 assays.

SDRF for RNA-seq experiment

Experiment E-MTAB-513

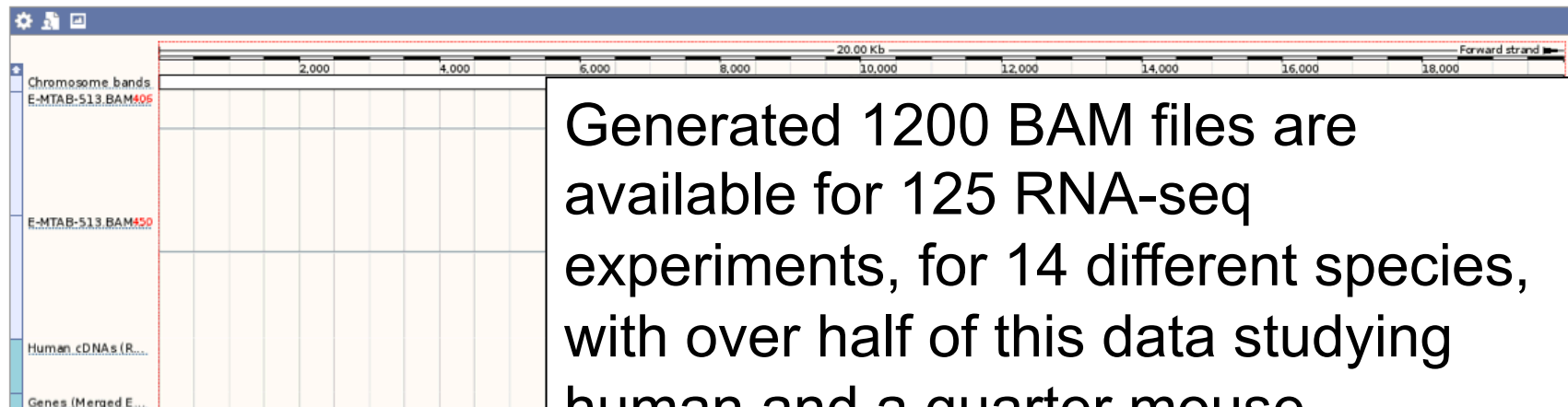
RNA-Seq of human individual tissues and mixture of 16 tissues (Illumina Body Map) (19 samples)

Source Name ^	Sample Characteristics					Factor Values		Links to Data
	Organism	Age (unit)	OrganismPart	Sex	ORGANISMPART	LIBRARYPREP	ENA	
HCT20142	Homo sapiens	60 years	kidney	female	kidney	mRNA-Seq	 	
HCT20142	Homo sapiens	60 years	kidney	female	kidney	mRNA-Seq	 	
HCT20142	Homo sapiens	60 years	kidney	female	kidney	mRNA-Seq	 	
HCT20143	Homo sapiens	77 years	heart	male	heart	mRNA-Seq	 	
HCT20143	Homo sapiens	77 years	heart	male	heart	mRNA-Seq	 	
HCT20143	Homo sapiens	77 years	heart	male	heart	mRNA-Seq	 	
HCT20144	Homo sapiens	37 years	liver	male	liver	mRNA-Seq	 	
HCT20144	Homo sapiens	37 years	liver	male	liver	mRNA-Seq	 	
HCT20144	Homo sapiens	37 years	liver	male	liver	mRNA-Seq	 	
HCT20145	Homo sapiens	65 years	lung	male	lung	mRNA-Seq	 	
HCT20145	Homo sapiens	65 years	lung	male	lung	mRNA-Seq	 	
HCT20145	Homo sapiens	65 years	lung	male	lung	mRNA-Seq	 	
HCT20146	Homo sapiens	86 years	lymph node	female	lymph node	mRNA-Seq	 	
HCT20146	Homo sapiens	86 years	lymph node	female	lymph node	mRNA-Seq	 	
HCT20146	Homo sapiens	86 years	lymph node	female	lymph node	mRNA-Seq	 	
HCT20147	Homo sapiens	73 years	prostate	male	prostate	mRNA-Seq	 	
HCT20147	Homo sapiens	73 years	prostate	male	prostate	mRNA-Seq	 	
HCT20147	Homo sapiens	73 years	prostate	male	prostate	mRNA-Seq	 	

BAM files visualization in Ensembl

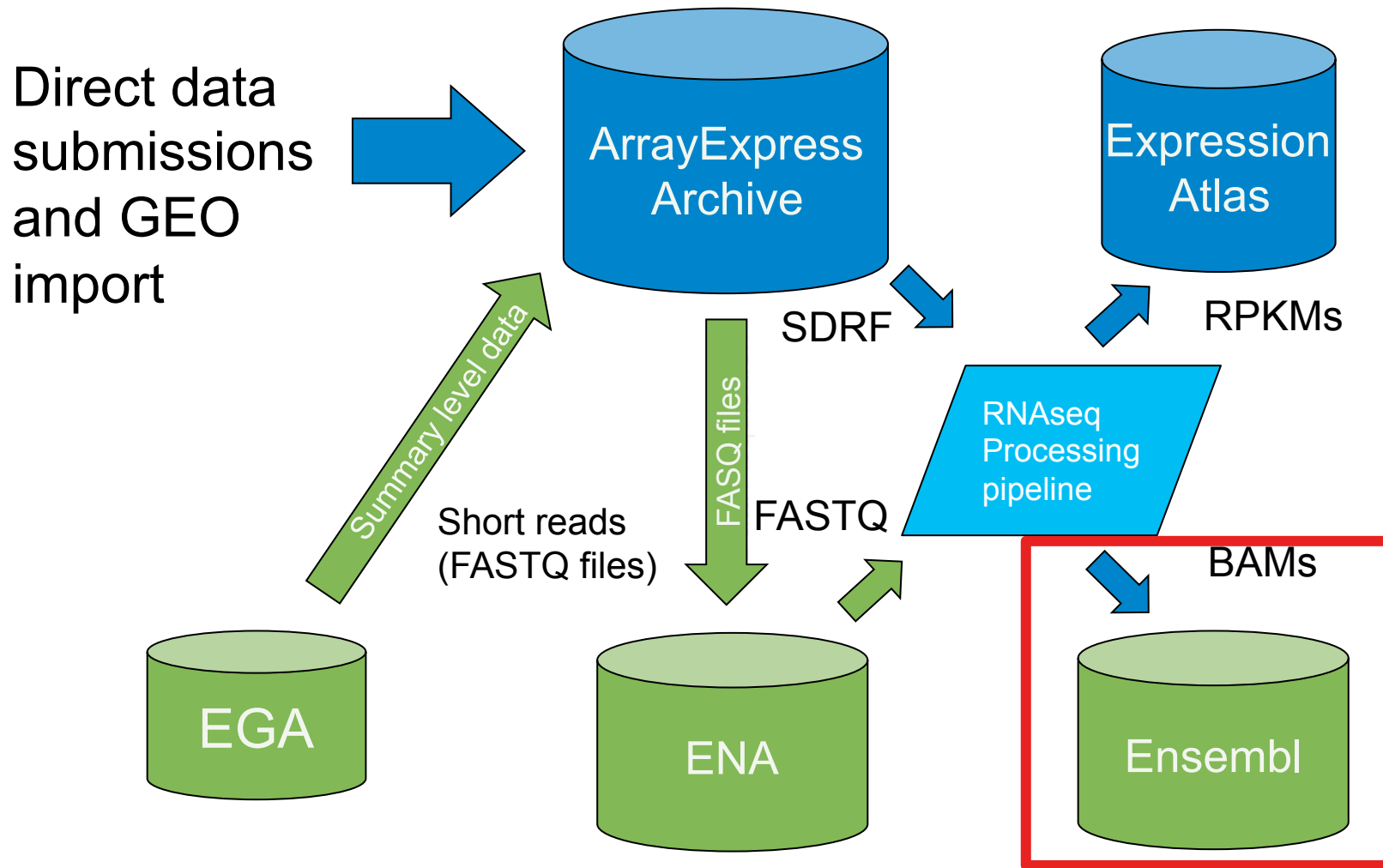


Location: 1:1-19999 Gene:

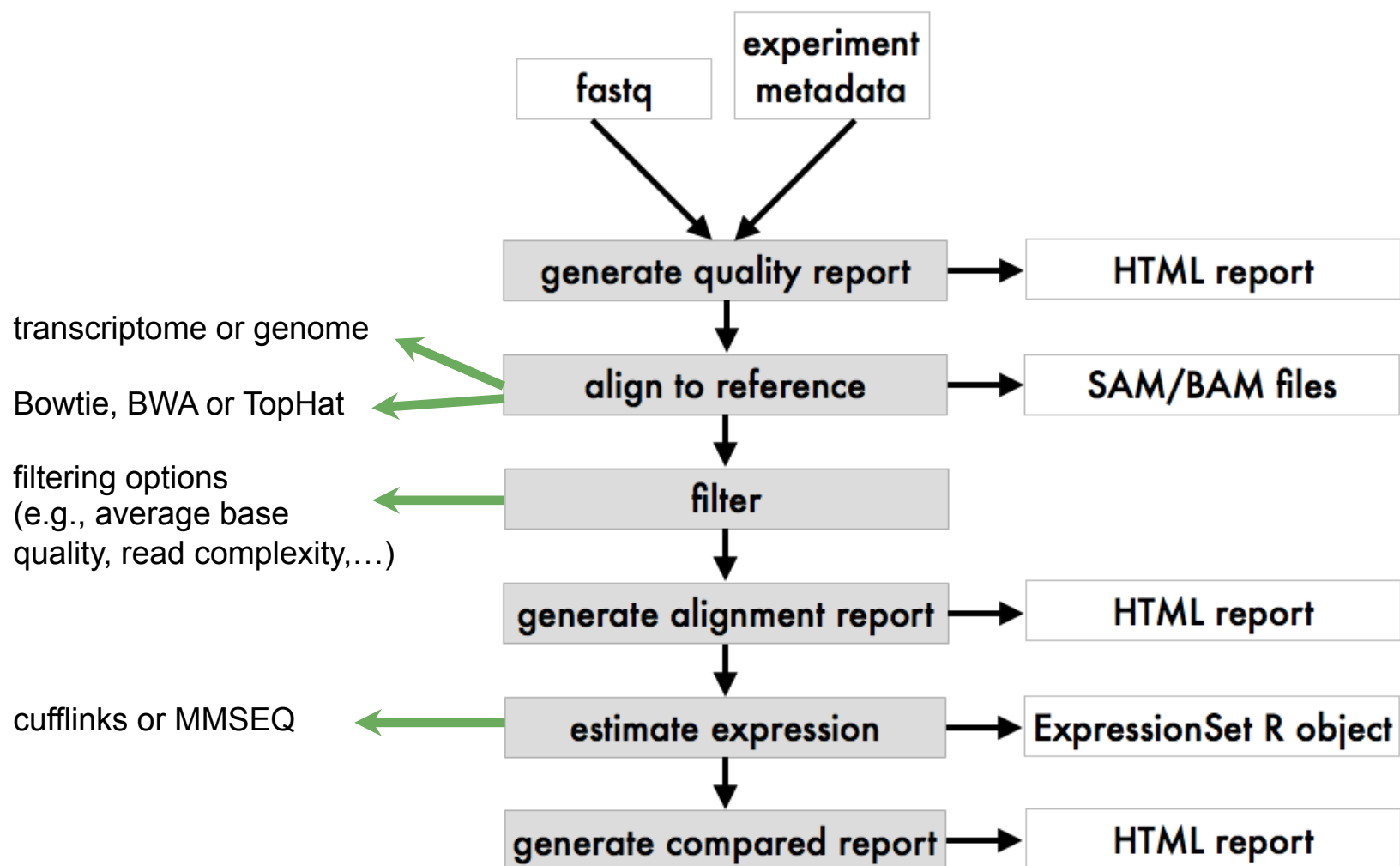


Generated 1200 BAM files are available for 125 RNA-seq experiments, for 14 different species, with over half of this data studying human and a quarter mouse

RNA-seq processing pipeline



ArrayExpressHTS pipeline



Links with data analysis tools

Accession	Title	Assays	Species	Date	Processed	Raw	Atlas
E-MEXP-1422	RNAi knock down using siRNAs targeting PROX1 or GFP in human colon carcinoma cells	6	Homo sapiens	2008-09-03			
Samples (6) Click for detailed sample information and links to data							
Platform (1) A-AFFY-37 - Affymetrix GeneChip Human Genome U133A 2.0 [HG-U133A_2]							
Protocols (7) Click for all experimental protocols							
Description siRNA transfection using either two different siRNAs targeting PROX1 or GFP							
Experiment types RNAi profiling by array, cellular modification, co-expression, in vitro							
Contact Antti Nykänen <antti.z.nykanen@helsinki.fi>							
Citation Independent filtering increases detection power for high-throughput experiments. Bourgon, Richard; Gentleman, Robert; Huber, Wolfgang. <i>Proc Natl Acad Sci U S A</i> , UKPMC 20460310							
MIAME							
		-					
	Platforms	Protocols	Factors	Processed	Raw		
Files							
	Data Archives	E-MEXP-1422.processed.1.zip, E-MEXP-1422.raw.1.zip					
	Investigation Description	E-MEXP-1422.idf.txt					
	Sample and Data Relationship	E-MEXP-1422.sdrf.txt					
	R ExpressionSet	E-MEXP-1422.eSet.r					
	Browse all available files						
Links							
	Send E-MEXP-1422 data to GENOMESPACE						
Status Submitted on 27 August 2008, released on 3 September 2008, last updated on 27 March 2012							

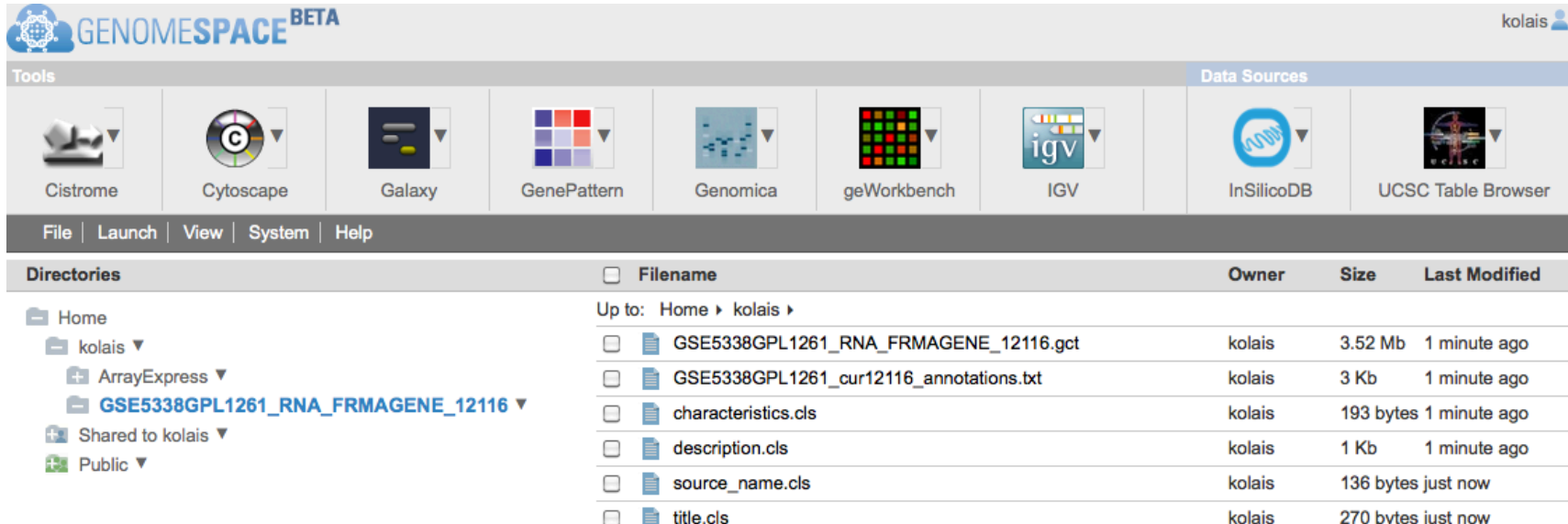
1 experiment, 6 assays.

Link with GenomeSpace

EBI > ArrayExpress > Experiments > E-MEXP-31 > Send to GenomeSpace

SEND TO  GENOMESPACE

Please login to (or register an account at) GenomeSpace first.



The screenshot shows the GenomeSpace interface. At the top, the user is logged in as 'kolais'. Below the navigation bar, there are two sections: 'Tools' and 'Data Sources'. The 'Tools' section includes icons for Cistrome, Cytoscape, Galaxy, GenePattern, Genomica, geWorkbench, and IGV. The 'Data Sources' section includes icons for InSilicoDB and UCSC Table Browser. Below these sections is a menu bar with 'File', 'Launch', 'View', 'System', and 'Help'. The main content area displays a file list under the 'Directories' section. The file list has columns for 'Filename', 'Owner', 'Size', and 'Last Modified'. The files listed are:

Filename	Owner	Size	Last Modified
GSE5338GPL1261_RNA_FRMAGENE_12116.gct	kolais	3.52 Mb	1 minute ago
GSE5338GPL1261_cur12116_annotations.txt	kolais	3 Kb	1 minute ago
characteristics.cls	kolais	193 bytes	1 minute ago
description.cls	kolais	1 Kb	1 minute ago
source_name.cls	kolais	136 bytes	just now
title.cls	kolais	270 bytes	just now

Link with Bioconductor for microarray data

Assay	Title	Assays	Species	Date	Processed	Raw	Atlas
EXP-1422	RNAi knock down using siRNAs targeting PROX1 or GFP in human colon carcinoma cells	6	Homo sapiens	2008-09-03			

Samples (6) [Click for detailed sample information and links to data](#)

Platform (1) A-AFFY-37 - Affymetrix GeneChip Human Genome U133A 2.0 [HG-U133A_2]

Protocols (7) [Click for all experimental protocols](#)

Description siRNA transfection using either two different siRNAs targeting PROX1 or GFP

Experiment types RNAi pro

Contact Antti Nyk

Citation Independ
Sci U S A

MIAME Platform

Files Data Arc
Investiga

R Express

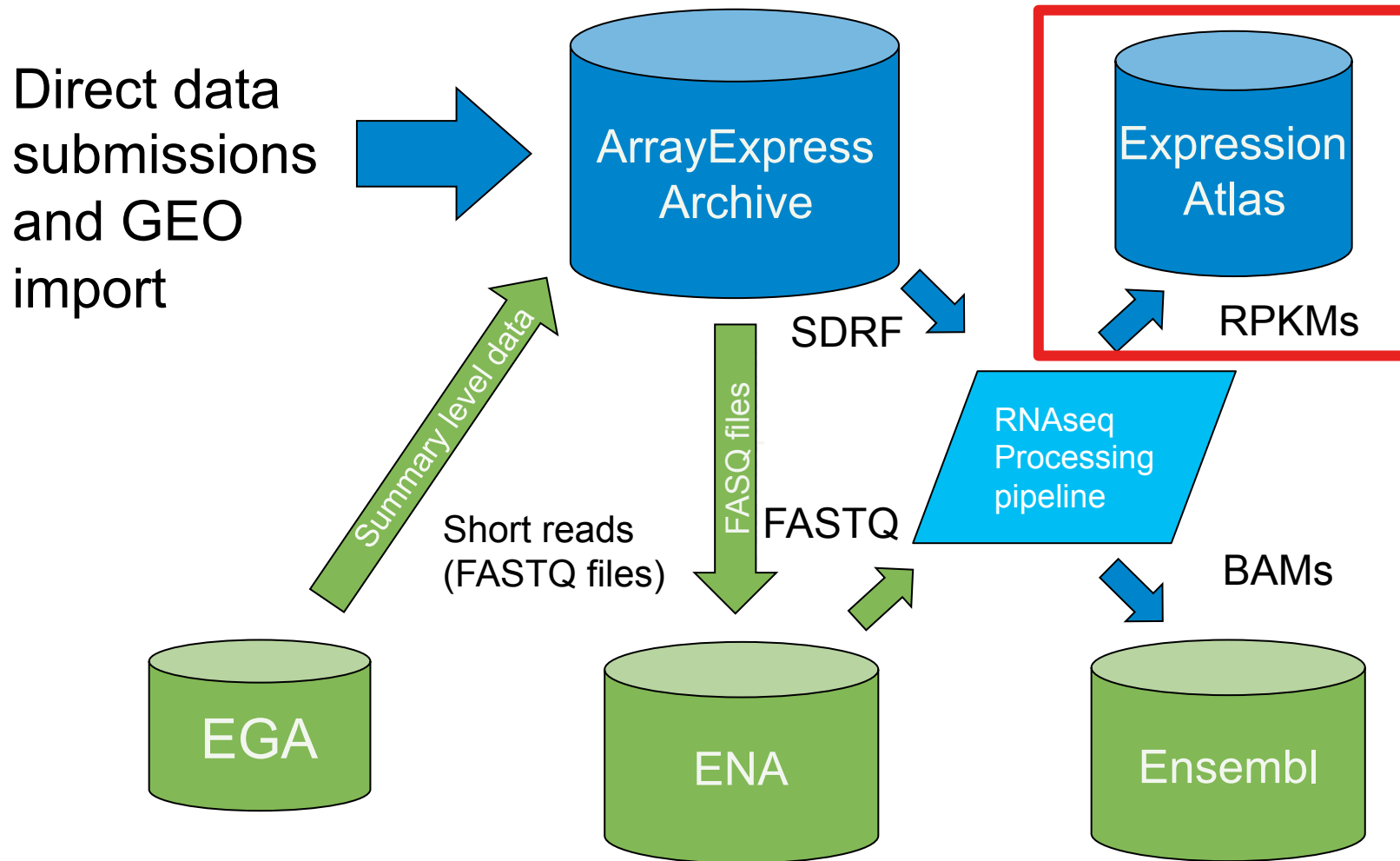
Links Gene Exp
Send E-M

Status Submitte

1 experiment, e

- Pre-generated R objects for 16250 out of 25,000 gene expression microarray experiments with raw data files available, using ArrayExpress package
- R objects available for > 85% of Affymetrix data
- Older submissions, other technologies and experiments with only processed data available can still be loaded in R, but require user-specified settings

RNA-seq processing pipeline



Expression Atlas

SERPINE2 Homo sapiens

[JSON](#) [XML](#)

SERPINE2 is differentially expressed in 282 experiments: 135 organism parts: liver, bone marrow, ... (133 more);, 164 disease states: normal, glioblastoma, ... (162 more);, 11 cell types 2 cell types 62 cell types and 87 other conditions.

Synonyms

PI7, PN1, SERPINE2, nexin, GDN, PNI

Orthologs

SRPN6 (Anopheles gambiae) SRPN16 (Anopheles gambiae) SRPN4 (Anopheles gambiae) srp-6 (Caenorhabditis elegans) srp-2 (Caenorhabditis elegans) srp-1 (Caenorhabditis elegans) NP_777094.1 (Bos taurus) ENSCING00000009089 (Ciona intestinalis) serpine2 (Danio rerio) SERPINE2 (Gallus gallus) Serpine2 (Mus musculus) GDN_RAT (Rattus norvegicus) srp-8 (Caenorhabditis elegans) NP_001023824.1 (Caenorhabditis elegans) Spn5 (Drosophila melanogaster) CG7219 (Drosophila melanogaster) CG6687 (Drosophila melanogaster) srp-3 (Caenorhabditis elegans) (Compare orthologs)

UniProt Accession

P07093, B4DIF2, B4DMR3, C9JN98, C9JRK5, C9JYC8, C9K031

Gene Ontology Term

cell differentiation, extracellular space, glycosaminoglycan binding, heparin binding, nervous system development...

InterPro Term

Protease inhibitor I4, serpin, Protease_inhib_I4_serpin

Search EB-eye

ENSG00000135919

Show more properties

Experimental Factors

Organism part

studied in E-AFMX-5, E-GEOD-803, E-GEOD-1577, E-MEXP-1251, E-MEXP-433, ... (31 experiments)

Number of published studies where the gene is over/under expressed compared to the gene's overall mean expression level in the study.



show expression data for all values of this factor>>

Cell line

studied in E-GEOD-17482, E-GEOD-3860, E-GEOD-5258, E-GEOD-21750, E-MEXP-1006, ... (37 experiments)

Factor Value	U/D	Experiments
MCF-7	4	E-GEOD-20081, E-GEOD-5823, E-GEOD-3529, E-GEOD-19639
A549	3	E-GEOD-3202, E-GEOD-6013, E-MTAB-62
Mcf7	3	E-MTAB-62, E-MTAB-37, E-GEOD-5258
PC3	3	E-GEOD-5258, E-MEXP-2034, E-GEOD-14464
SW480	3	E-MEXP-1014, E-MTAB-37, E-GEOD-18005
BT474	1/2	E-MTAB-37, E-MTAB-62, E-MEXP-440

368 more value(s).
show this factor only>>

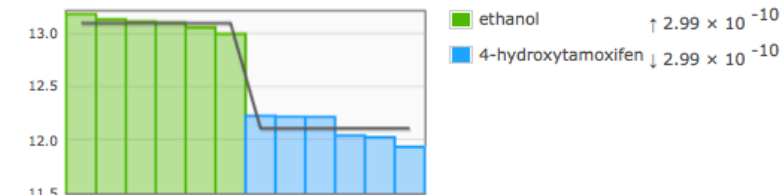
Expression Profiles

[1](#) [2](#) [3](#) [4](#) [5](#) ... [56](#) [57](#)

282 experiments showing differential expression

E-MEXP-2241: Transcription profiling of human cells transfected with a miR-34a LNA inhibitor to investigate its role in oncogene-induced senescence

Experimental Factors



Array Design: A-AFFY-44 Affymetrix GeneChip Human Genome U133 Plus 2.0 [HG-U133_Plus_2]

Show expression profile / experiment details

E-MEXP-2590: Transcription profiling of human colon cancer cell line SW620 treated with tetradecylthioacetic acid (TTA) or control (NaOH).

Experimental Factors

RNA-seq data in Atlas

EMBL-EBI [EB-eye Search](#) All Databases [Go](#) [Reset](#) [Advanced Search](#) [Give us feedback](#)

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ATLAS [home](#) | [about the project](#) | [faq](#) [feedback](#) | [blog](#) | [das](#) | [api](#) **new** | [help](#)

Transcription profiling by high throughput sequencing of 60 HapMap CEU population individuals

[JSON](#) [XML](#)

Data shown for array design: A-ENST-1

Individual **Population**

■ ARF5:ENST00000467281
 ■ C2orf56:ENST00000412815
 ■ C2orf56:ENST00000443335

[box plot](#) [line plot](#)

Gene	Design Element	Genome View	Experimental Factor	Factor Value	UP/DOWN	T-Statistic	P-Value	
			All factor values			All expressions		
ARF5	ENST00000467281	Genome View	Individual	NA12043	UP	1375.959	<1 × 10 ⁻¹⁰	
C2orf56	ENST00000412815	Genome View	Individual	NA12872	UP	1248.913	<1 × 10 ⁻¹⁰	
C2orf56	ENST00000443335	Genome View	Individual	NA12760	UP	1222.672	<1 × 10 ⁻¹⁰	
ARF5	ENST00000459680	Genome View	View in the Ensembl Genome Browser (new window)		11994	UP	1160.970	<1 × 10 ⁻¹⁰
SEMA3F	ENST00000002829	Genome View	Individual	NA11829	UP	1006.333	<1 × 10 ⁻¹⁰	
FUCA2	ENST00000451668	Genome View	Individual	NA11894	UP	992.569	<1 × 10 ⁻¹⁰	
C2orf56	ENST00000432075	Genome View	Individual	NA12044	UP	927.148	<1 × 10 ⁻¹⁰	
CYP26B1	ENST00000001146	Genome View	Individual	NA12044	UP	889.018	<1 × 10 ⁻¹⁰	
SEMA3F	ENST00000450338	Genome View	Individual	NA11830	UP	794.103	<1 × 10 ⁻¹⁰	

E-MTAB-197

Platform: A-ENST-1
 Organism: homo sapiens
 Samples: 73
 Individuals:
 Study type:

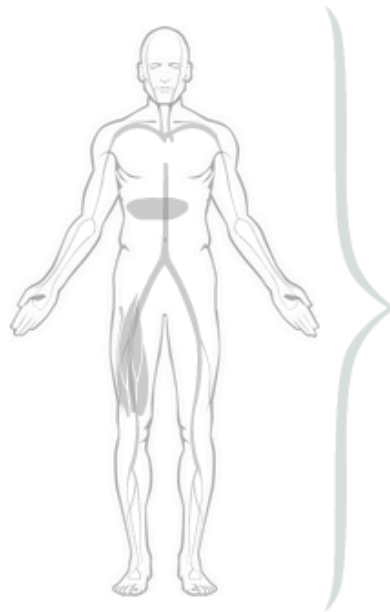
- experiment design
- experiment analysis

New baseline Atlas prototype

Gene IDs (max 10) Organism parts Expression level cutoff

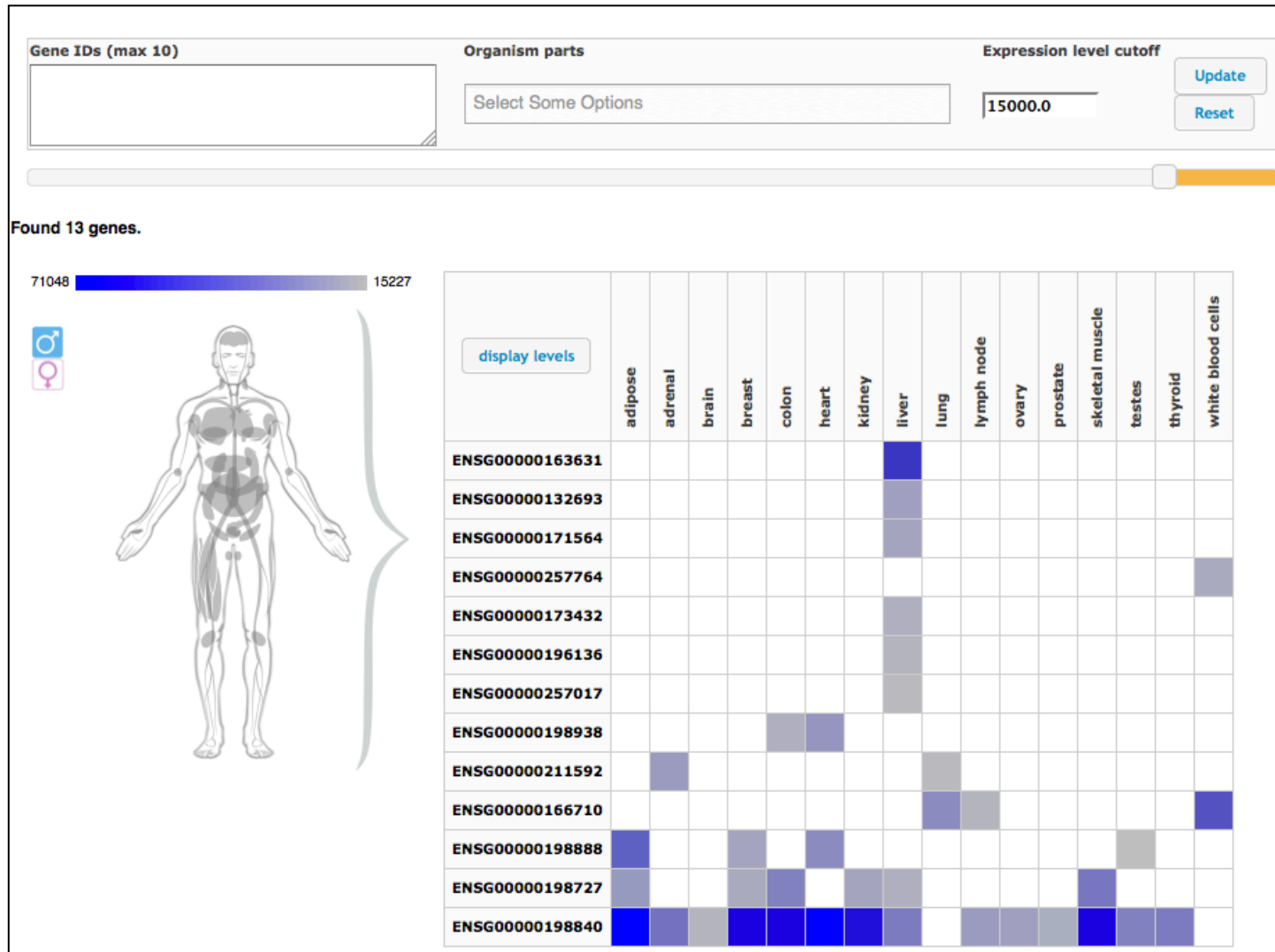
Found 18 genes.

48053 10787



	liver	skeletal muscle	white blood cells
ALB	48053		
CRP	23110		
FGB	21521		
RP11-1143G9.4			20135
SAA1	19355		
SERPINA3	17995		
HP	16767		
FGG	12953		
CKM		11919	
RBP4	10787		

New baseline Atlas prototype



How to improve our current pipeline?

- We are working towards understanding the effects of analyzing RNA-seq data using different tools
- Survey of different short read mapping methods: Fonseca et al, Tools for mapping high-throughput sequencing data, Bioinformatics (2012)
- This will be followed by comparison of different quantitation methods run on top of different mappers, on a selected datasets

GenePattern

Modules & Pipelines Suites Job Results Resources Downloads Help GenomeSpace

Modules & Pipelines

category suite all
open all | close all

- Recently Used
- Annotation
- caBIG
- Clustering
- Data Format Conversion
- Flow Cytometry
- Gene List Selection
- IGV
- Image Creators
- Missing Value Imputation
- Pathway Analysis
- Prediction
- Preprocess & Utilities
- Projection
- Proteomics
- RNA-seq
- Sequence Analysis
- SNP Analysis
- Statistical Methods
- Survival Analysis
- Uncategorized
- Visualizer
- pipeline

Welcome to the GenePattern / GenomeSpace public beta integration server. [hide]

Show parameter descriptions

MAGETABImportViewer version 1.1

* required field [properties](#) | [export](#) | [help](#)

sdrf file* Batch Parameter
 Specify URL Basic Upload
 The sdrf file - .txt

data files* no file selected
 Select a single file under 2GB to upload.
 Specify URL Basic Upload
 A zip file containing MAGE-TAB data matrix files - .zip

[properties](#) | [export](#) | [help](#)

View code to call MAGETABImportViewer:

Recent Jobs Uploads GenomeSpace

No jobs to display

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