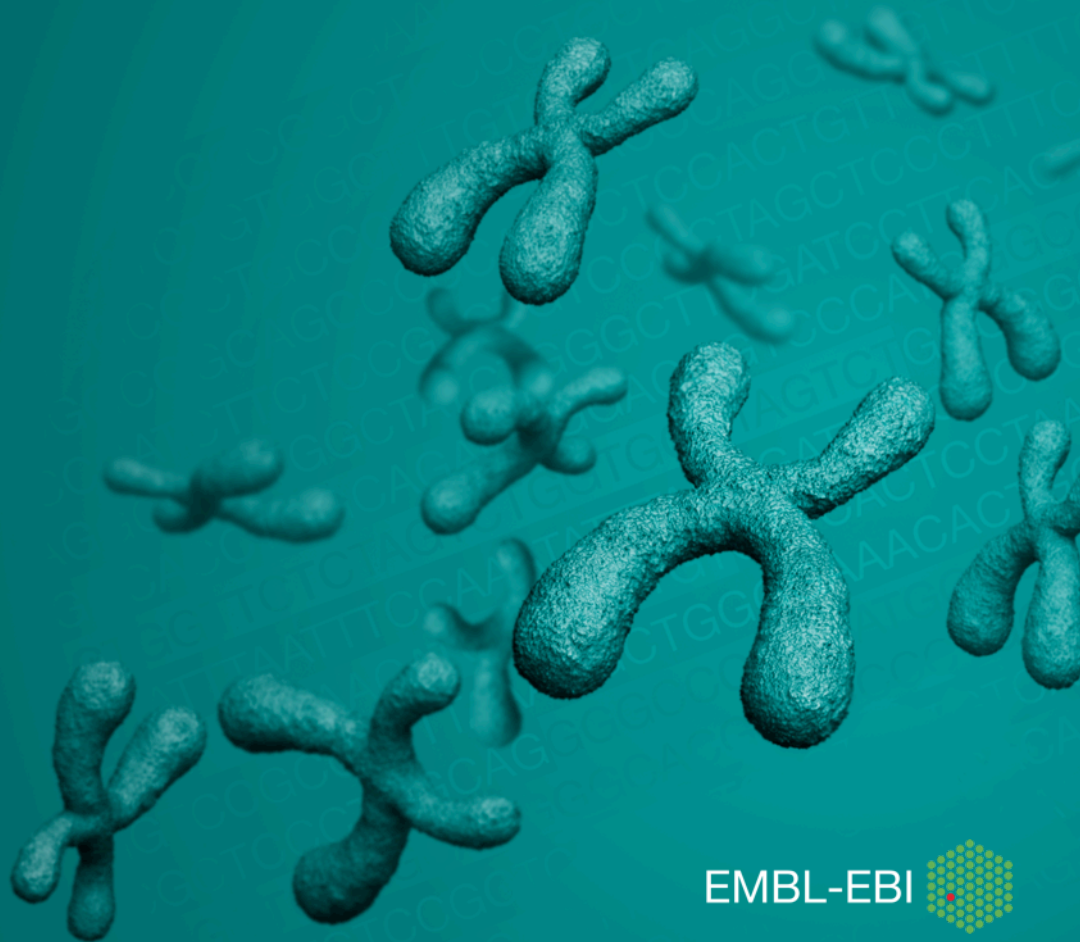


# The challenge of NGS data

Guy Cochrane



Data  
intensity



Availability of  
technologies



Broadening of  
applications



# Context

# EMBL European Bioinformatics Institute

## Genes, genomes & variation

European Nucleotide Archive  
1000 Genomes

Ensembl  
Ensembl Genomes

European Genome-phenome Archive  
Metagenomics portal

## Gene, protein & metabolite expression

ArrayExpress

Expression Atlas

Metabolights  
PRIDE

## Protein sequences, families & motifs

InterPro

Pfam

UniProt

## Molecular structures

Protein Data Bank in Europe  
Electron Microscopy Data Bank

## Chemical biology

ChEMBL

ChEBI

## Systems

BioModels  
Enzyme Portal

BioSamples

## Literature & ontologies

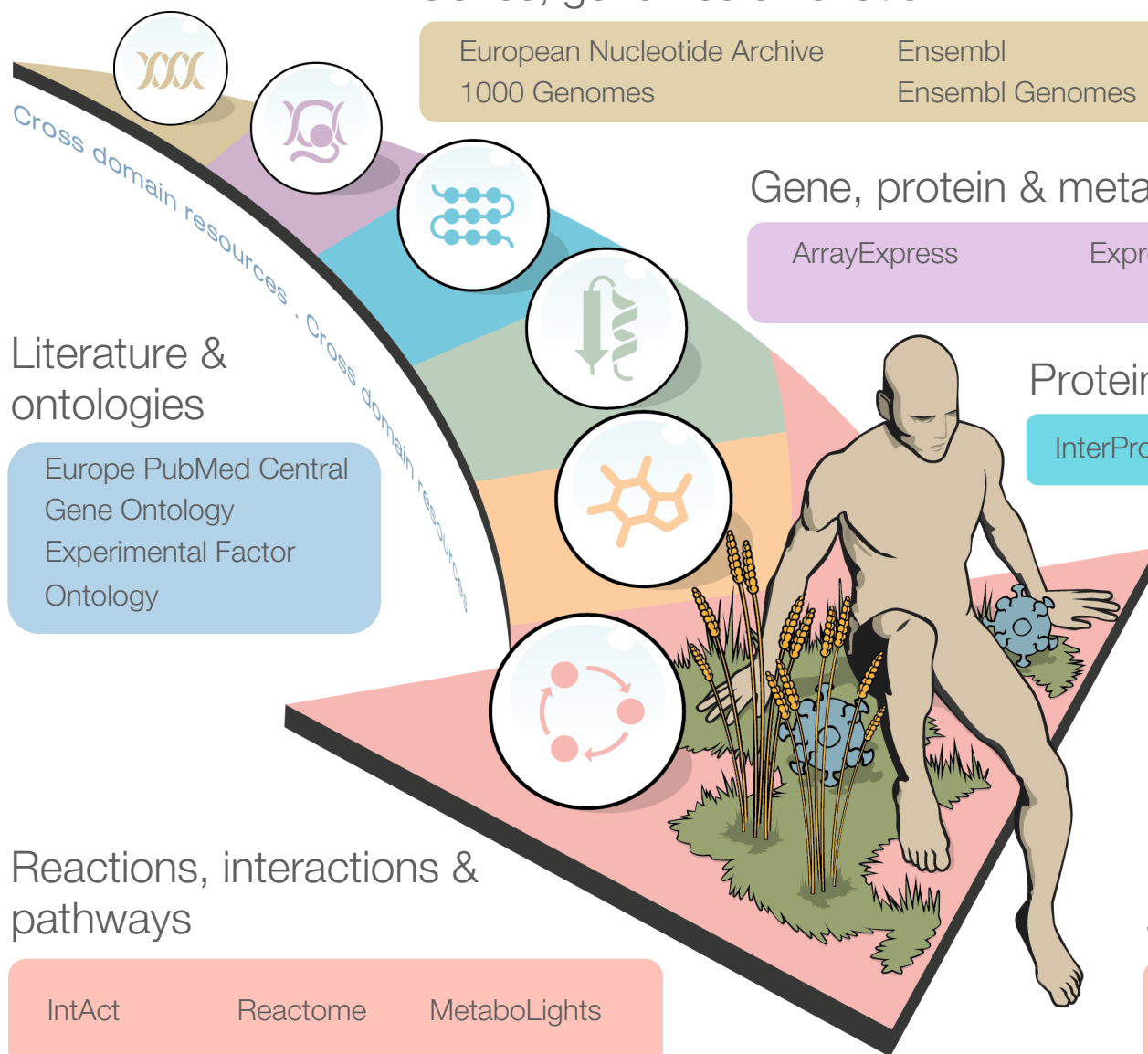
Europe PubMed Central  
Gene Ontology  
Experimental Factor  
Ontology

## Reactions, interactions & pathways

IntAct

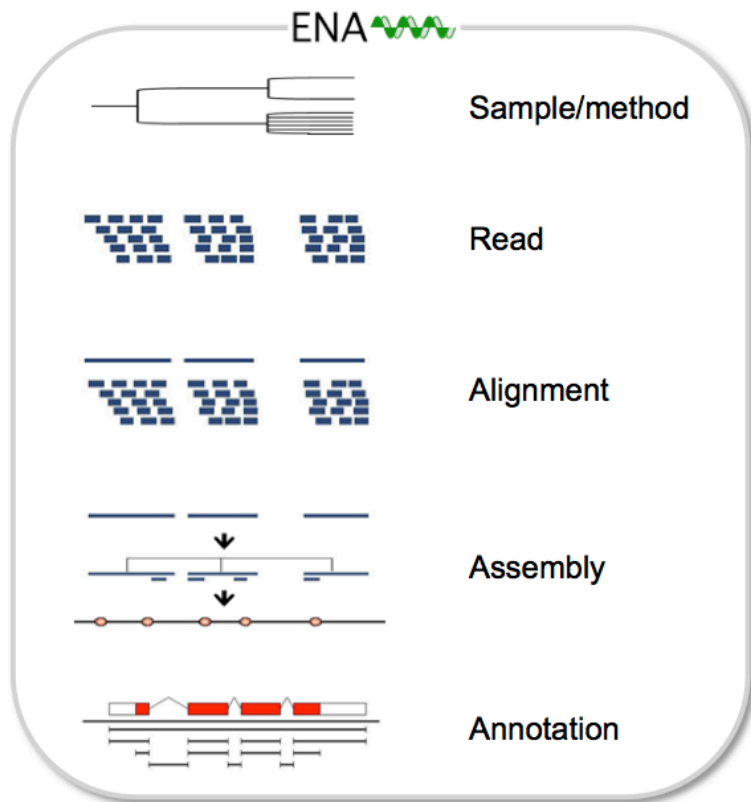
Reactome

MetaboLights





# European Nucleotide Archive (ENA)



ENA 

<http://www.ebi.ac.uk/ena/>

- A **broad platform** for the management, sharing, integration and dissemination of sequence data
- Established in the early 1980s, extended for **new technologies and applications**
- **Globally comprehensive scientific record** and European node of INSDC
- **Connectivity** with broader EMBL-EBI resources
- Sequence data **foundation**
- **Sustained** within EMBL-EBI under EMBL funding with additional support from EC, UK Research councils, Wellcome Trust, etc.
- **Substantial scale:** 1.3 petabase pairs across >1 million taxa, 2,000-5,000 active data providers, global consumer userbase
- Rich submission, discovery and retrieval **software, tools and services**

# Submissions

Home **New Submission** Studies Sample Groups Samples Experiments Runs Projects

Start >> **Sample** >> Finish

Please create new samples by uploading a spreadsheet or by following the instructions below.

Please select the checklist that you wish to use for your sample submission

If you already have a spreadsheet containing your data upload it here.

**Default Checklist**

[Upload Spreadsheet](#)

Book1 - Microsoft Excel

File Home Insert Page Layout Formulas Data Review View

From Access From Web From Text From Other Sources Get External Data Existing Connections Refresh All Properties Edit Links Connections Sort & Filter Filter Clear Reapply Advanced Text to Columns Remove Duplicates Data Validation Data Tools Consolidate What-If Analysis Group Ungroup Subtotal Outline

A1	#checklist_accession									
#checklist_accession	ERC000001									
#unique_name_prefix	mouse_dendrocyte_									
sample_alias	tax_id	scientific_name	common_name	anonymized_name	sample_title	sample_description	tissue_type	sex	collectio	
#template	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
#units										
mouse_dendrocyte_1	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
mouse_dendrocyte_2	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
mouse_dendrocyte_3	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
mouse_dendrocyte_4	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
mouse_dendrocyte_5	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
mouse_dendrocyte_6	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	
mouse_dendrocyte_7	10090	Mus musculus	house mouse		mouse dendrocyte	A dendrocyte brain sample was extracted from a mouse	brain	male	13/0	

Sheet1 Sheet2 Sheet3

Ready

85%

# Data discovery

*temperature* >=10 AND *temperature* <=25 AND *geo\_box1*(42, 17, 43, 18)

Accession	First public	Geographical location	Submitter's sample name	Depth (m)	Environment (Biome)	Temperature (C)	Sampling Site
<a href="#">SAMEA2591084</a>	2014-07-11	42.2038 N 17.715 E	TARA_E500000075	5.0	marine biome (ENVO:00000447)	17.32198	TARA_023
<a href="#">SAMEA2591093</a>	2014-06-23	42.2038 N 17.715 E	TARA_A100000551	5.0	marine biome (ENVO:00000447)	17.32198	TARA_023
<a href="#">SAMEA2591094</a>	2014-06-23	42.2038 N 17.715 E	TARA_A100000553	5.0	marine biome (ENVO:00000447)	17.32198	TARA_023
<a href="#">SAMEA2591095</a>	2014-06-26	42.2038 N 17.715 E	TARA_A100000552	5.0	marine biome (ENVO:00000447)	17.32198	TARA_023
<a href="#">SAMEA2591096</a>	2014-06-26	42.2038 N 17.715 E	TARA_A100000547	5.0	marine biome (ENVO:00000447)	17.32198	TARA_023
<a href="#">SAMEA2591097</a>	2014-07-18	42.2038 N 17.715 E	TARA_E500000056	5.0	marine biome (ENVO:00000447)	17.32198	TARA_023
<a href="#">SAMEA2591098</a>	2014-07-18	42.1735 N 17.7252 E	TARA_E500000081	55.0	marine biome (ENVO:00000447)	15.194062	TARA_023
<a href="#">SAMEA2591099</a>	2014-07-11	42.1735 N 17.7252 E	TARA_E500000080	55.0	marine biome (ENVO:00000447)	15.194062	TARA_023
<a href="#">SAMEA2591103</a>	2014-06-23	42.1735 N 17.7252 E	TARA_A100000559	55.0	marine biome (ENVO:00000447)	15.194062	TARA_023

*tax\_tree*(10090) AND *library\_source*="GENOMIC" AND *instrument\_platform*="ILLUMINA" AND *library\_strategy*="ChIP-Seq"

Study accession	Sample accession	Run accession	Scientific name	Fastq files (ftp)	Fastq files (galaxy)	Submitter's sample name
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604495</a>	<a href="#">ERR537823</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-WT-120m-R2
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604492</a>	<a href="#">ERR537824</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-KO-000m-R1
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604494</a>	<a href="#">ERR537825</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-input-120m-R1
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604493</a>	<a href="#">ERR537826</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-WT-000m-R1
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604496</a>	<a href="#">ERR537827</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-WT-120m-R1
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604491</a>	<a href="#">ERR537828</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-KO-120m-R2
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604500</a>	<a href="#">ERR537829</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-input-000m-R1
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604498</a>	<a href="#">ERR537830</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-KO-120m-R1
<a href="#">PRJEB6568</a>	<a href="#">SAMEA2604497</a>	<a href="#">ERR537831</a>	<a href="#">Mus musculus</a>	<a href="#">File 1</a> <a href="#">File 2</a>	<a href="#">File 1</a> <a href="#">File 2</a>	E-MTAB-2661:Exp2-Irf5-WT-000m-R1

# Cross-references and tagging

GUI: <http://www.ebi.ac.uk/ena/data/xref/search>



Search by Source    Search by ENA record

XREF Source: COMPARE-RefGenome    Accession:    Target:    Expanded:     Search    Clear

Download: 1 - 138 of 138 results in [TEXT](#)

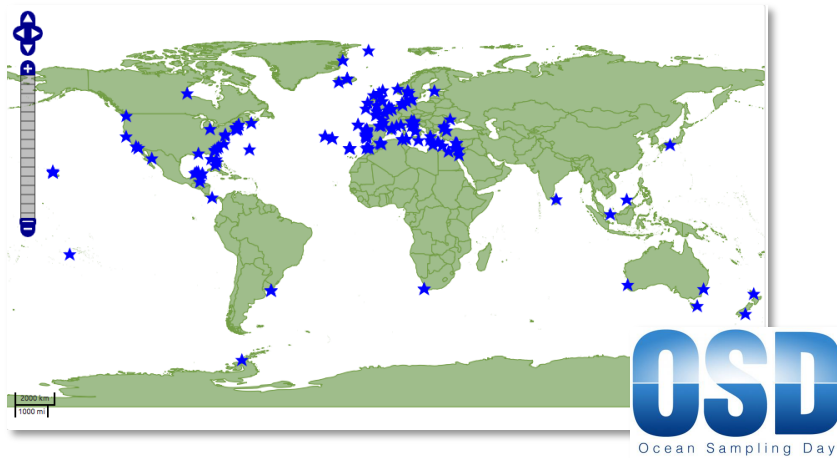
Showing results 1 - 10 of 138 results

Source Primary Accession	Source Secondary Accession	Target Primary Accession	Target Secondary Accession	Family	Genus	species	1st below-species level	2nd below-species level	3rd below-species level	Aggregated taxonomic name	genome
Picton/2003/AU		<a href="#">AY919139</a>		Caliciviridae	norovirus	GII	P21, 1			NoV/GII.P21/GII.1	partial
Pont de Roide 673/2004/France		<a href="#">AY682549</a>		Caliciviridae	norovirus	GII	P21, 2			NoV/GII.P21/GII.2	partial
NLV/Paris Island/2003/USA		<a href="#">AY652979</a>		Caliciviridae	norovirus	GII	P21, 3			NoV/GII.P21/GII.3	partial
OsakaNI/2004/JP		<a href="#">DQ366347</a>		Caliciviridae	norovirus	GII	P22, 2			NoV/GII.P22/GII.2	complete
Hokkaido/133/2003/JP		<a href="#">AB212306</a>		Caliciviridae	norovirus	GII	P22, 5			NoV/GII.P22/GII.5	partial
YURI		<a href="#">AB083780</a>		Caliciviridae	norovirus	GII	P22, 22			NoV/GII.P22/GII.22	complete
Arg320		<a href="#">AF190817</a>		Caliciviridae	norovirus	GII	Pa, 3			NoV/GII.Pa/GII.3	partial
SnowMountainvirus		<a href="#">AY134748</a>		Caliciviridae	norovirus	GII	Pc, 2			NoV/GII.Pc/GII.2	complete
GII/IPH2172-09VG4/2009/BE		<a href="#">JF697282</a>		Caliciviridae	norovirus	GII	Pe, 3			NoV/GII.Pe/GII.3	partial





# Ocean Sampling Day and Tara Oceans



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About us > News > Press releases > Tara Oceans: the data

## Tara Oceans: the data

Scientists from the *Tara Oceans* expedition have published a fleet of papers in a special issue of *Science*, offering a glimpse into the vast, uncharted biodiversity in the world's oceans: 40 million genes, interactions between as-yet-unknown species, and profound insights into the effects of changing water temperatures – and that's just the tip of the iceberg. Data from these studies, together with those from Ocean Sampling Day 2013, are now freely available through the *Pangea* sample registry at the University of Bremen and the European Nucleotide Archive (ENA) and Metagenomics Portal at EMBL-EBI.

Tara researchers from EMBL, CNRS and many other institutes have compiled genetic material from around 35,000 marine plankton samples, which included viruses, microbes, algae, fish larvae and many other complex microscopic organisms.

- Overview
- Press releases
- News archive
- Service news
- Brochures
- Photos of EMBL-EBI
- Blogs

**THE TARA OCEANS EXPEDITION - THE DATA**

**14,000 km**  
The distance sailed by the Tara Oceans expedition over 1140 days (2009-2013). The 140 scientists stopped at 210 stations to collect samples.

**361 million km<sup>2</sup>** - the surface area of the world's ocean

**35,000 samples**  
The 140 contributors to the Tara Oceans expedition helped collect, pack and ship around 35,000 samples of plankton and water.

**7012 datasets**  
As of May 2015, the ENA hosts data on 75% of the samples collected – 9349 are listed in the *PANGEA* sample registry. This is one of the richest and most consistently described molecular sample collection in the public domain, and it is freely available to everyone. More samples, from the Arctic expedition in 2013, will be submitted in 2015.

**11,535 gigabytes**  
The size of the Tara datasets in the European Nucleotide Archive is of May 2015. This represents 12,581 gigabases – roughly equivalent to 135 fully sequenced human genomes.

**40,000**  
Previously undescribed genetic sequences, which could represent thousands of new species and ecosystem interactions. Considering the size of the world's oceans, there is much, much more to discover.

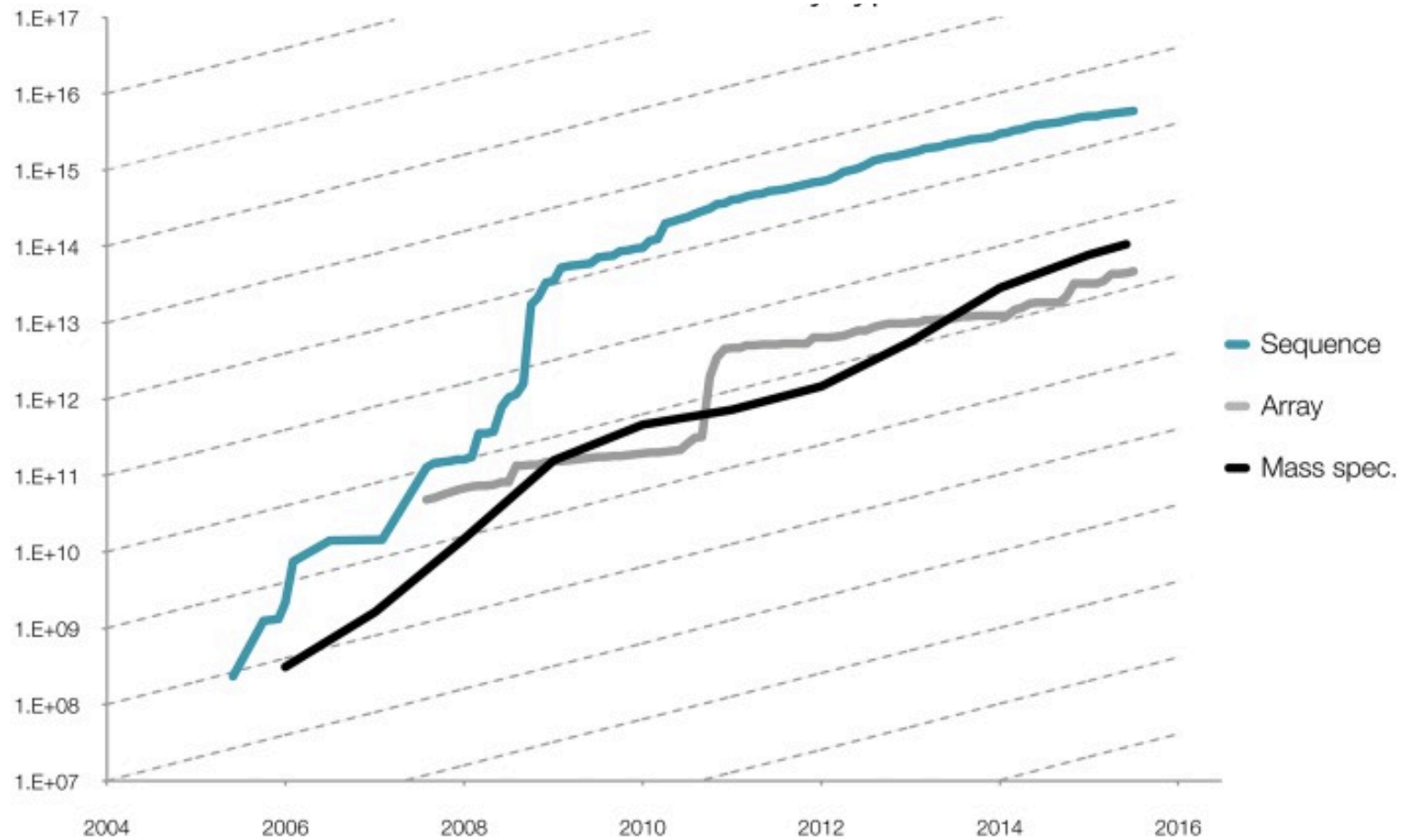
www.embl.ac.uk/services/tara-oceans-data

www.embl.de/tara-oceans/start

EMBL

Technology

# Data accumulation



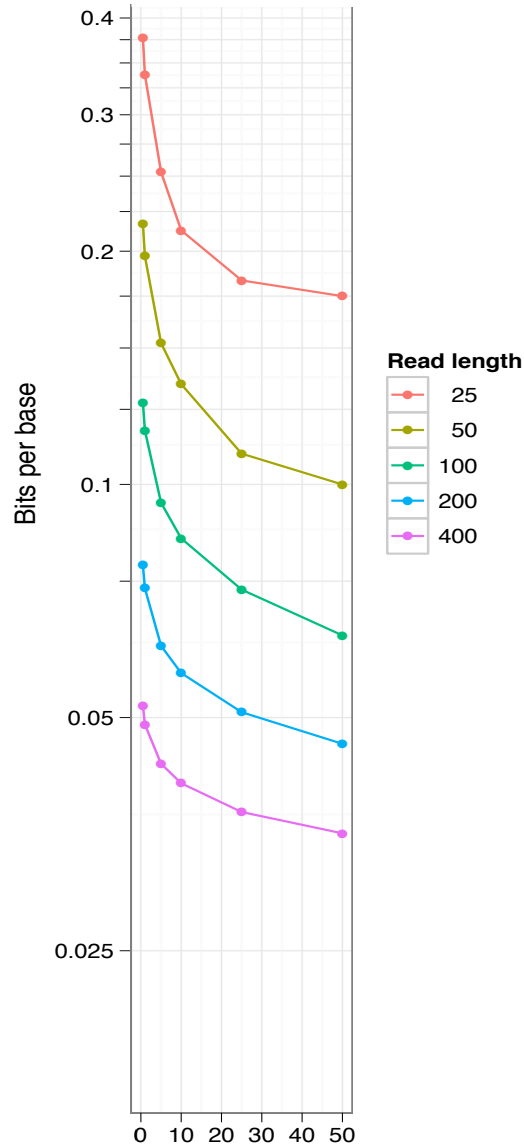
Cook CE, Bergman MT, Finn RD, Cochrane G, Birney E, Apweiler R. The European Bioinformatics Institute in 2016: Data growth and integration. *Nucleic Acids Res.* 2016 Jan;44(D1) D20-6. doi:10.1093/nar/gkv1352. PMID: 26673705; PMCID: PMC4702932.



# Starting point

TGAGCTCTAAGTACC  
329183050298757

# Sequence compression



- Encoding of read starts and differences
- 3.5x–100x compression over existing formats
- Scales favourably with increasing read length and density

Fritz, M.H. Leinonen, R., et al. (2011) Efficient storage of high throughput DNA sequencing data using reference-based compression. *Genome Res.* 21 (5), 734-40.

# Quality compression

TGAGCTCTAAGTACC  
329183050298757

Horizontal reduction



002020010022212

Vertical reduction



-2---30---9---7

# Quality compression: simple, horizontal reduction



Photograph from MichaelMaggs, [http://en.wikipedia.org/wiki/File:Amanita\\_muscaria\\_\(fly\\_agaric\).JPG](http://en.wikipedia.org/wiki/File:Amanita_muscaria_(fly_agaric).JPG)

# Models for data reduction

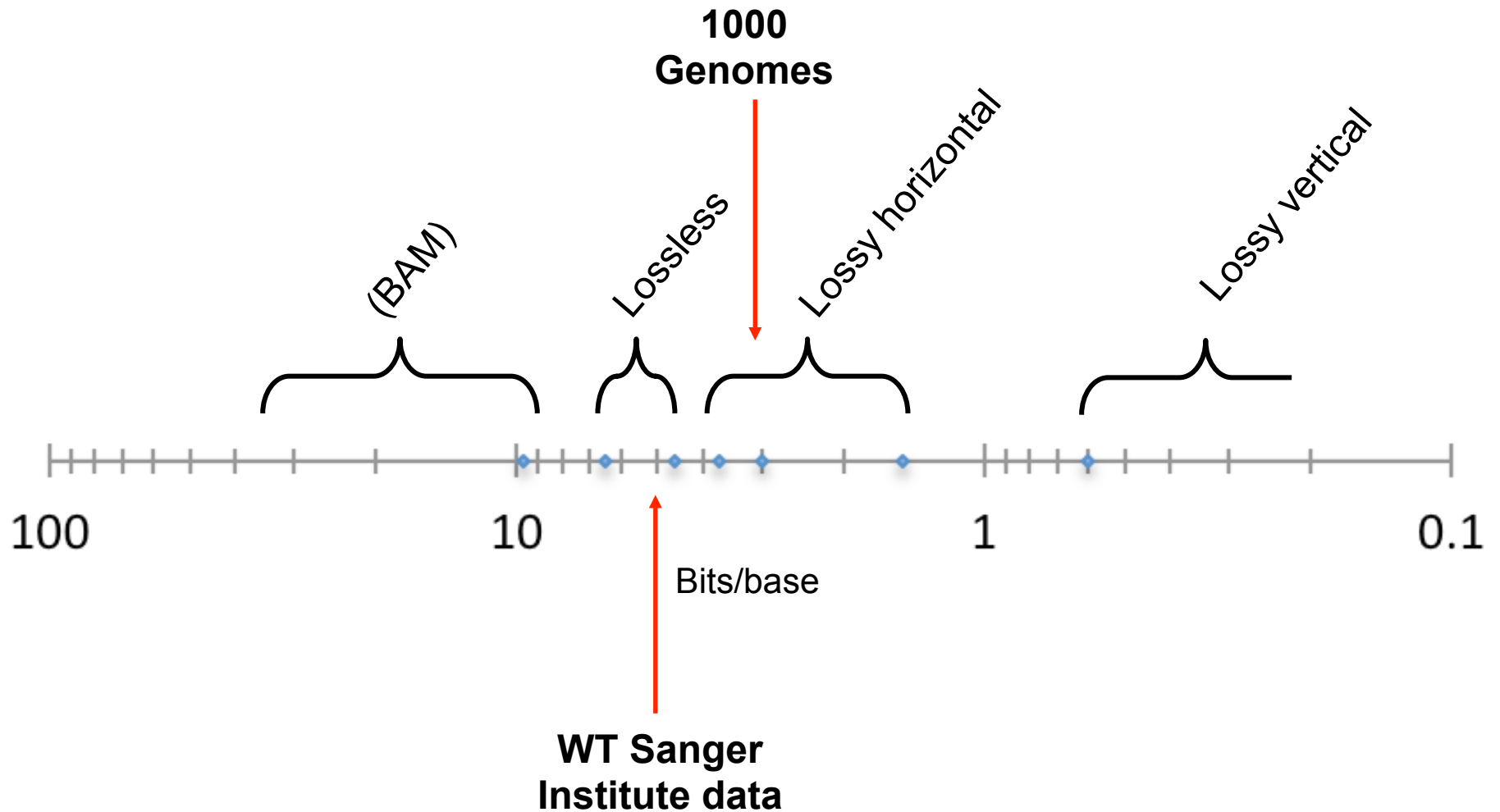


Jong-Seok Lee et al. (2009), [http://mmspg.epfl.ch/files/content/sites/mmspl/files/shared/lee\\_icme.pdf](http://mmspg.epfl.ch/files/content/sites/mmspl/files/shared/lee_icme.pdf)

# CRAM performance



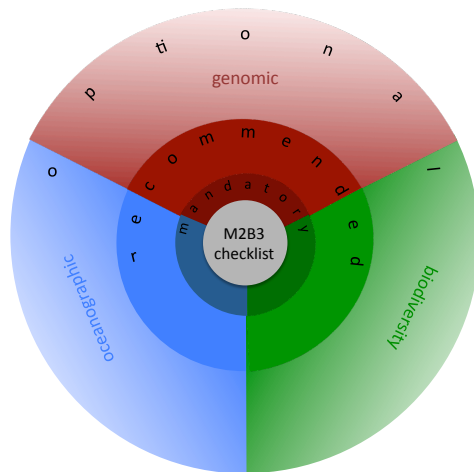
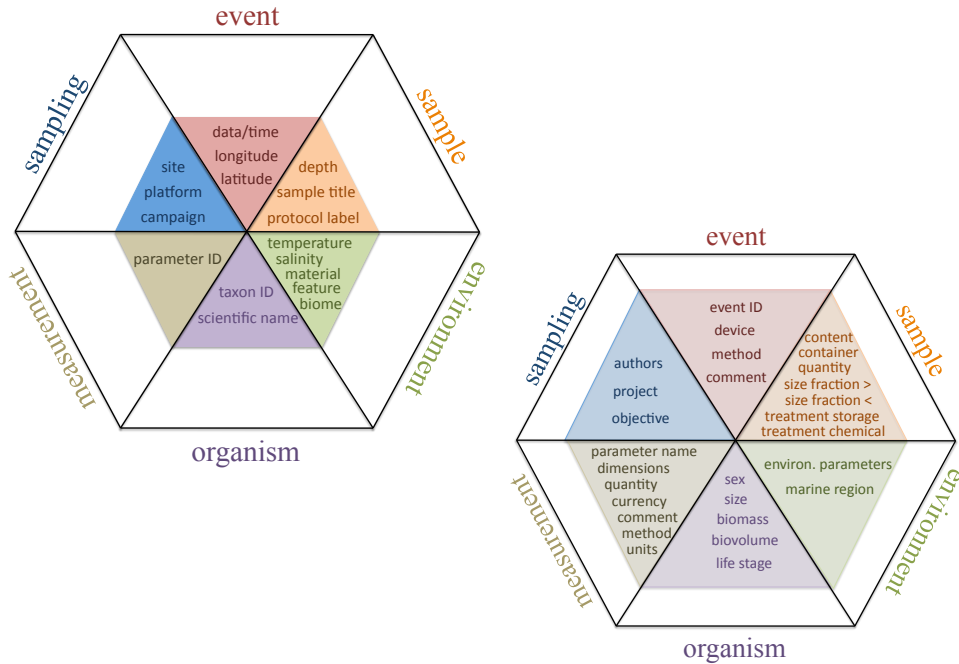
# CRAM performance



Human aspects



# Standards



**COMMENTARY**

**Open Access**

## Marine microbial biodiversity, bioinformatics and biotechnology (M2B3) data reporting and service standards

Petra ten Hoopen<sup>1</sup>, Stéphane Pesanti<sup>2</sup>, Renzo Kottmann<sup>3</sup>, Anna Kopf<sup>3,9</sup>, Mesude Bicak<sup>4</sup>, Simon Claus<sup>5</sup>, Klaas Deneudt<sup>5</sup>, Catherine Borremans<sup>5</sup>, Peter Thijssen<sup>7</sup>, Stefanie Dekeyser<sup>5</sup>, Dick MA Schaap<sup>7</sup>, Chris Bowler<sup>8</sup>, Frank Oliver Glöckner<sup>3,9</sup> and Guy Cochrane<sup>1\*</sup>

**Abstract**

Contextual data collected concurrently with molecular samples are critical to the use of metagenomics in the fields of marine biodiversity, bioinformatics and biotechnology. We present here Marine Microbial Biodiversity, Bioinformatics and Biotechnology (M2B3) standards for "Reporting" and "Serving" data. The M2B3 Reporting Standard (1) describes minimal mandatory and recommended contextual information for a marine microbial sample obtained in the epipelagic zone, (2) includes meaningful information for researchers in the oceanographic, biodiversity and molecular disciplines, and (3) can easily be adopted by any marine laboratory with minimum sampling resources. The M2B3 Service Standard defines a software interface through which these data can be discovered and explored in data repositories. The M2B3 Standards were developed by the European project Micro B3, funded under 7<sup>th</sup> Framework Programme "Ocean of Tomorrow", and were first used with the Ocean Sampling Day initiative. We believe that these standards have value in broader marine science.

**Keywords:** Data standard, Marine, Molecular, Biodiversity, Microbial, Bioinformatics, Reporting, Interoperability

**Background**

An immense wealth of genetic, functional and morphological diversity in marine ecosystems remains unexplored, offering the potential for substantial scientific and biotechnological discoveries. Indeed, significant interest in this area has led to large-scale initiatives, such as Tara Oceans [1], the Global Ocean Survey [2] and Malaspina [3], that target the exploration of marine biodiversity on planetary scales. While the shared goal of such initiatives is the development of an understanding of the compos-

biology will derive benefit. Prerequisite for the successful exploitation of acquired data are standards that enable interoperability in the data infrastructure.

Just as marine studies span many disciplines (e.g. biological, oceanographic, molecular), use of data from marine studies requires approaches that traverse the many disciplines, asking questions, for example, of species distribution, physical oceanographic parameters, molecular biology and data licensing. Each discipline has established infrastructure and best practice for the dissemination,

## ten Hoopen et al. *Standards in Genomic Sciences* (2015) 10:20

... explorations and will continue to build a powerful marine data infrastructure from which ecosystems biology and

positories, and analysis when data lack of a consistent environment for the discovery and retrieval of data.

The Marine Microbial Biodiversity, Bioinformatics, Biotechnology Project (Micro B3) [4] unites intensive oceanographic monitoring, thorough biodiversity studies

\* Correspondence: cochrane@ebi.ac.uk  
<sup>1</sup>European Nucleotide Archive, EMBL-EBI, Wellcome Trust Genome Campus Hinxton, Cambridge CB10 1SD, UK  
Full list of author information is available at the end of the article



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# Metagenomics

www.ebi.ac.uk/metagenomics/

Int GTLs ENA Search EuroPMC Cal edit EBI-docs ENA-docs Jira SUB UPD SAP WBV Conv >> +

# EBI Metagenomics

Home Submit data Projects Samples Comparison tool About Contact Not logged in Login

17558 data sets

4173 metagenomes  
888 metatranscriptomes  
12429 amplicons  
68 assemblies

Public 15129 runs  
14255 samples  
165 projects

Private 2429 runs  
2240 samples  
106 projects

## Browse projects

### By selected biomes

- Soil (20)
- Marine (35)
- Forest (4)
- Non-human host (42)
- Engineered (12)
- Freshwater (10)
- Grassland (5)
- Human gut (25)
- Air (1)
- Wastewater (2)

[View all biomes](#)

### Latest projects 165

- American Gut Project**  
The American Gut project is the largest crowdsourced citizen science project to date. Fecal, oral, skin, and other body site samples collected from thousands of participants represent the ...  
[View more - 8037 samples - compare](#)
- Brain meta-transcriptomics from harbor seals to infer the role of the microbiome and virome in a stranding event**  
Marine diseases are becoming more frequent, and tools for identifying pathogens and disease reservoirs are needed to help prevent and mitigate epizootics. Meta-transcriptomics provides ...  
[View more - 14 samples - compare](#)
- Study of the abundance of bacteria from human samples**

[View all projects](#)

## Spotlight

TARA ocean project

Plankton ecosystems contain a phenomenal reservoir of life: more than 10 billion organisms inhabit every litre of oceanic water, including viruses, prokaryotes, unicellular eukaryotes (protists), and metazoans. Plankton's importance for the earth's climate is at least equivalent to that of the rainforest. Yet only a small fraction of organisms that compose it have been classified and analysed.

**Display a menu** edition, led by EMBL senior scientist Eric Karsenti, has been travelling across the world (2009-10) collecting more than 200,000 samples containing millions of

## Tools

Functional sample comparison

Interested in comparing the functional profile of sequencing runs within a project? Now it is possible, using our comparison tool, which provides analysis based on a slimmed-down subset of Gene Ontology (GO) terms, specially developed to describe metagenomic data.

You can visualise the results using a range of interactive charts (bar charts, stacked columns, Principal Component Analysis and heatmaps). The different charts can be exported in PNG, PDF and SVG formats, so that they can easily be included in presentations and publications.

# Nucleic Acids Research

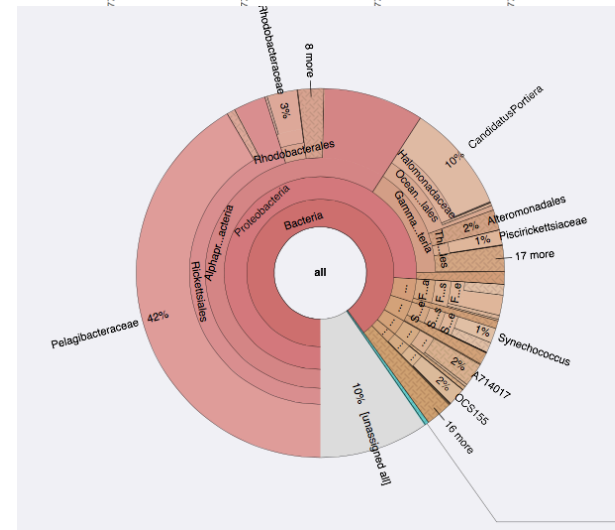
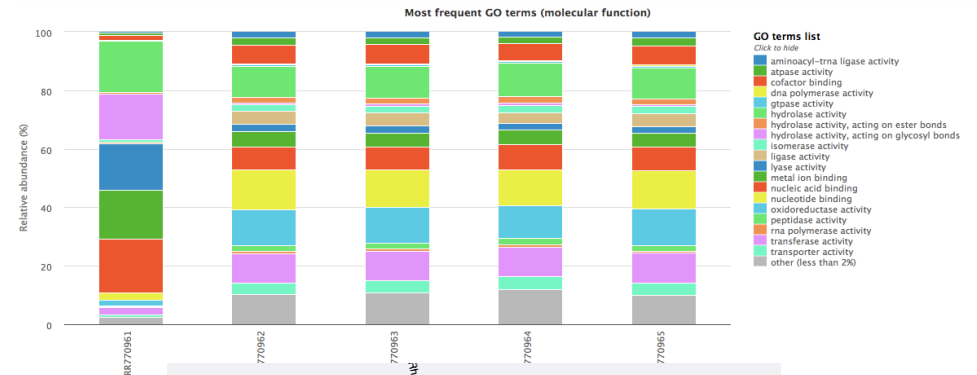
Nucleic Acids Res. 2016 January 4; 44(Database issue): D595–D603.  
Published online 2015 November 17. doi: [10.1093/nar/gkv1195](https://doi.org/10.1093/nar/gkv1195)

PMCID: PMC4702853

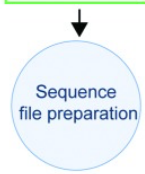
## EBI metagenomics in 2016 - an expanding and evolving resource for the analysis and archiving of metagenomic data

Alex Mitchell, Francois Bucchini, Guy Cochrane, Hubert Denise, Petra ten Hoopen, Matthew Fraser, Sebastien Pesseat, Simon Potter, Maxim Scheremetjev, Peter Sterk, and Robert D. Finn\*

Jump to: [Biological process](#) | [Molecular function](#) | [Cellular component](#)



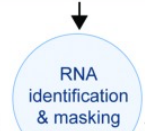
Raw reads



Initial reads



QC



Processed reads

RNA identification & masking

rRNAs

16S

Taxonomic analysis

Taxonomic assignments

Reads with rRNA masked

ORF predictions

Predicted CDS

Functional analysis

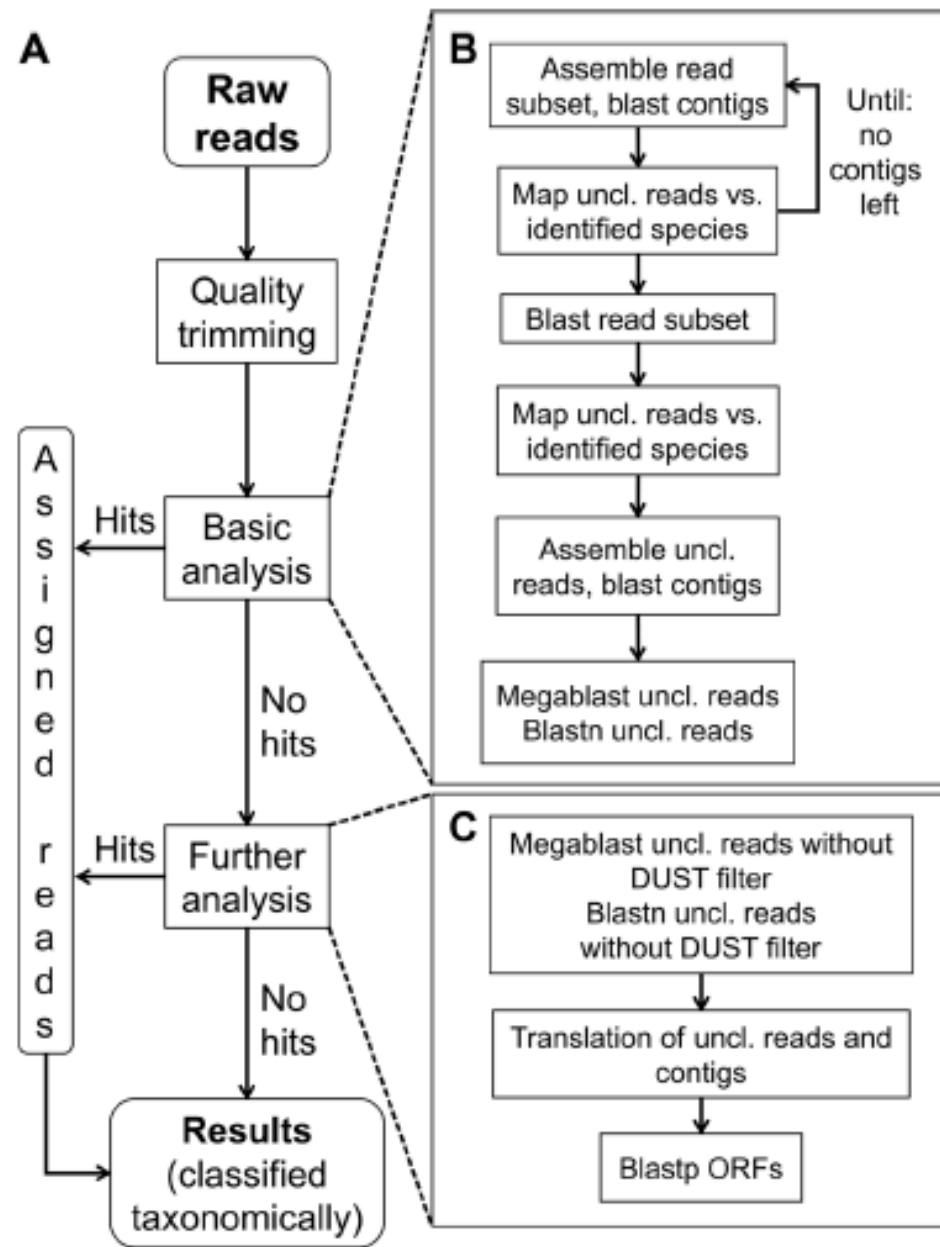
Functional assignments

- Download EBI metagenomics
- Download ENA
- Process/component
- Multiple files
- Not available for download

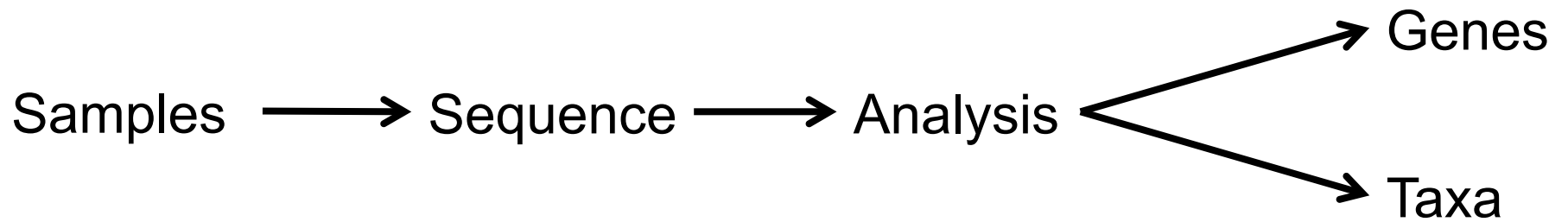
# RIEMS: a software pipeline for sensitive and comprehensive taxonomic classification of reads from metagenomics datasets

Matthias Scheuch<sup>†</sup>, Dirk Höper<sup>\*†</sup> and Martin Beer

Scheuch et al. (2015) BMC Bioinformatics 16:69



# Identification data



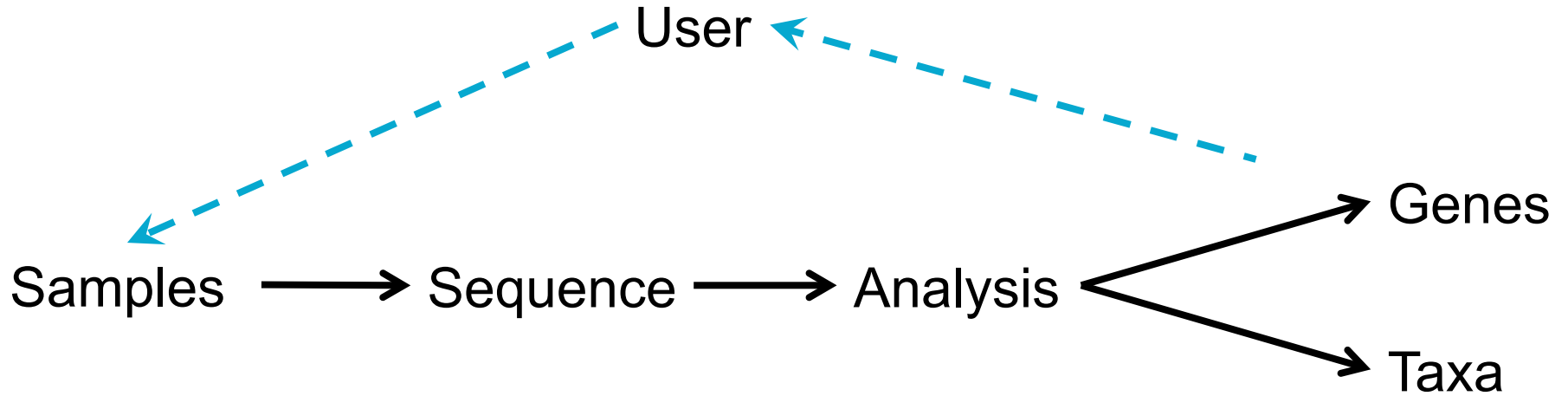


# Identification data accumulation

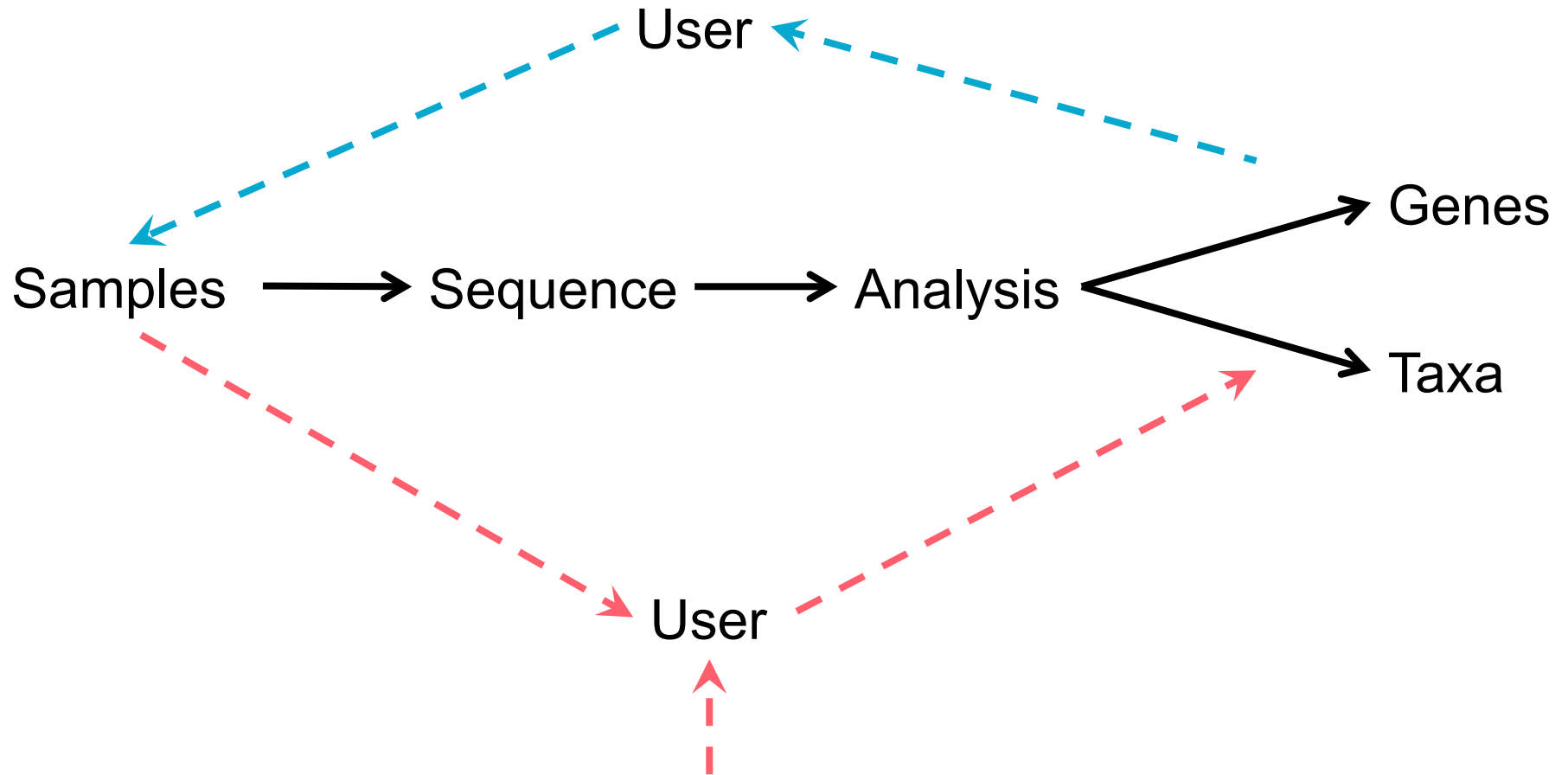
- Identification against context is informative
- (Molecular observations may be tentative or high confidence)
- Coincidental observations (time, place, virulence, host phenotypes)
- ‘Capture’ identifications to ‘connect’ coincidental observations



# Identification data



# Identification data



# Acknowledgements

## Standards & support

Ana Cerdeño-Tárraga, Ana Luisa Toribio, **Petra ten Hoopen**, Marc Rosello, Richard Gibson, Jeena Rajan, **Clara Amid**

## Compression technology

**Vadim Zalunin**, Ewan Birney, **James Bonfield**, Rasko Leinonen

## ENA technical services

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