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The BioMart community portal: an innovative alternative to large, centralized data repositories

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ABSTRACT

The BioMart Community Portal (www.biomart.org) is a community-driven effort to provide a unified interface to biomedical databases that are distributed worldwide. The portal provides access to numerous database projects supported by 30 scientific organizations. It includes over 800 different biological datasets spanning genomics, proteomics, model organisms, cancer data, ontology information and more. All resources available through the portal are independently administered and funded by their host organizations. The BioMart data federation technology provides a unified interface to all the available data. The latest version of the portal comes with many new databases that have been created by our ever-growing community. It also comes with better support and extensibility for data analysis and visualization tools. A new addition to our toolbox, the enrichment analysis tool is now accessible through graphical and web service interface. The BioMart

community portal averages over one million requests per day. Building on this level of service and the wealth of information that has become available, the BioMart Community Portal has introduced a new, more scalable and cheaper alternative to the large data stores maintained by specialized organizations.

INTRODUCTION

The methods of data generation and processing that are utilized in biomedical sciences have radically changed in recent years. With the advancement of new high-throughput technologies, data have grown in terms of quantity as well as complexity. However, the significance of the information that is hidden in the newly generated experimental data can only be deciphered by linking it to other types of biological data that have been accumulated previously. As a result there are already numerous bioinformatics resources and new ones are constantly being created. Typically, each resource comes with its own query interface. This poses a problem for the scientists who want to utilize such resources in their research. Even the simplest task such as compil-



Figure 1. BioMart USA community databases and their host countries.

ing results from a few existing resources is challenging due to the lack of a complete, up to date catalogue of already existing resources and the necessity of constantly learning how to navigate new query interfaces. A different challenge is faced by collaborating groups of scientists who independently generate or maintain their own data. Such collaborations are seriously hampered by the lack of a simple data management solution that would make it possible to connect their disparate, geographically distributed data sources and present them in a uniform way to other scientists. The BioMart project has been set up to address these challenges.

SOFTWARE

BioMart is an open source data management system, which is based on a data federation model (1). Under this model, each data source is managed, updated and released independently by their host organization while the BioMart software provides a unified view of these sources that are distributed worldwide. The data sources are presented to the user through a unified set of graphical and programmatic interfaces so that they appear to be a single integrated database. To navigate this database and compile a query the user does not have to learn the underlying structure of each data source but instead use a set of simple abstractions: datasets, filters and attributes. Once a user's input is provided, the software distributes parts of the query to individual data sources, collects the data and presents the user with the unified result set.

The BioMart software is data agnostic and its applications are not limited to biological data. It is cross-platform and supports many popular relational database management systems, including MySQL, Oracle, PostgreSQL. It also supports many third party packages such as Taverna

(2), Galaxy (3), Cytoscape (4) and biomaRt (5), which part of the Bioconductor (6) library.

The BioMart project currently maintains two independent code bases: one written in Java and one written in Perl. For more information about the architecture and capabilities of each of the packages please refer to previous publications (1,7). The latest version of the Java based BioMart software has been significantly enhanced with new additions to the existing collection of graphical user interfaces (GUIs). It has also been re-engineered to provide better support and extensibility for data analysis and visualization tools. The first of the BioMart tools based on this new framework has already been implemented and is accessible from the BioMart Community Portal.

The BioMart project adheres to the open source philosophy that promotes collaboration and code reuse. Two good examples of how this philosophy benefits the scientific community are provided by two independent research groups. The INRA group based in Toulouse, France has recently released a software package called RNABrowse (RNA-Seq De Novo Assembly Results Browser) (8). The Pfizer group based in La Jolla, USA has just announced the release of OASIS: A Web-based Platform for Exploratory Analysis of Cancer Genome and Transcriptome data (www.oasis-genomics.org). Both of these software packages are based on the BioMart software.

DATA

The BioMart community consists of a wide spectrum of different research groups that use the BioMart technology to provide access to their databases. It currently comprises 30 scientific organizations supporting 38 database projects that contain over 800 different biological datasets spanning ge-

Table 1. BioMart community databases and their host organizations

Database	Description	Host	Reference
Animal Genome databases ^{a,b}	Agriculturally important livestock genomes	Iowa State University, US	NA
Atlas of UTR Regulatory Activity (AURA) ^a	Meta-database centred on mapping post-transcriptional (PTR) interactions of trans-factors with human and mouse untranslated regions (UTRs) of mRNAs	University of Trento, Italy	(36)
BCCTB Bioinformatics Portal ^a	Portal for mining omics data on breast cancer from published literature and experimental datasets	Breast Cancer Campaign/Barts Cancer Institute UK	(37)
Cildb	Database for eukaryotic cilia and centriolar structures, integrating orthology relationships for 44 species with high-throughput studies and OMIM	Centre National de la Recherche Scientifique (CNRS), France	(38)
COSMIC	Somatic mutation information relating to human cancers	Wellcome Trust Sanger Institute (WTSI), UK	(39)
DAPPER ^a	Mass spec identified protein interaction networks in <i>Drosophila</i> cell cycle regulation	Department of Genetics, University of Cambridge, Cambridge, UK	NA
EMAGE	In situ gene expression data in the mouse embryo	Medical Research Council, Human Genetics Unit (MRC HGU), UK	(40)
Ensembl	Genome databases for vertebrates and other eukaryotic species	Wellcome Trust Sanger Institute (WTSI), UK	(41)
Ensembl Genomes	Ensembl Fungi, Metazoa, Plants and Protists	European Bioinformatics Institute (EBI), UK	(41)
Euraexpress	Transcriptome atlas database for mouse embryo	Medical Research Council, Human Genetics Unit (MRC HGU), UK	(42)
EuroPhenome	Mouse phenotyping data	Harwell Science and Innovation Campus (MRC Harwell), UK	(15)
FANTOM5 ^a	The FANTOM5 project mapped a promoter level expression atlas in human and mouse. The FANTOM5 BioMart instance provides the set of promoters along with annotation.	RIKEN Center for Life Science Technologies (CLST), Japan	(16)
GermOnLine	Cross-species microarray expression database focusing on germline development, meiosis, and gametogenesis as well as the mitotic cell cycle	Institut national de la santé et de la recherche médicale (Inserm), France	(17)
GnpIS ^a	Genetic and Genomic Information System (GnpIS)	Institut Nationale de Recherche Agronomique (INRA), Unité de Recherche en Génomique-Info (URGI), France	(18)
Gramene	Agriculturally important grass genomes	Cold Spring Harbor Laboratory (CSHL), US	(43)
GWAS Central ^a	GWAS Central provides a comprehensive curated collection of summary level findings from genetic association studies	University of Leicester, UK	(19)
HapMap	Multi-country effort to identify and catalog genetic similarities and differences in human beings	National Center for Biotechnology Information (NCBI), US	(20)
HGNC	Repository of human gene nomenclature and associated resources	European Bioinformatics Institute (EBI), UK	(21)
i-Pharm ^a	PharmDB-K is an integrated bio-pharmacological network databases for TKM (Traditional Korean Medicine)	Information Center for Bio-pharmacological Network (i-Pharm), South Korea	(22)
InterPro	Integrated database of predictive protein 'signatures' used for the classification and automatic annotation of proteins and genomes	European Bioinformatics Institute (EBI), UK	(44)
KazusaMart	Cyanobase, rhizobia, and plant genome databases	Kazusa DNA Research Institute (Kazusa), Japan	NA
MGI	Mouse genome features, locations, alleles, and orthologs	Jackson Laboratory, US	(23)
Pancreatic Expression Database	Results from published literature	Barts Cancer Institute UK	(24)
ParameciumDB	Paramecium genome database	Centre National de la Recherche Scientifique (CNRS), France	(25)
Phytozome	Comparative genomics of green plants	Joint Genome Institute (JGI)/Center for Integrative Genomics (CIG), US	(26)

Table 1. Continued

Database	Description	Host	Reference
Potato Database	Potato and sweetpotato phenotypic and genomic information	International Potato Center (CIP), Peru	NA
PRIDE	Repository for protein and peptide identifications	European Bioinformatics Institute (EBI), UK	(45)
Regulatory Genomics Group ^a	Predictive Models of Gene Regulation from High-Throughput Epigenomics Data	Universitat Pompeu Fabra (UPF), Spain	(27)
Rfam ^a	The Rfam database is a collection of RNA families, each represented by multiple sequence alignments, consensus secondary structures and covariance models (CMs).	Wellcome Trust Sanger Institute (WTSI), UK	(28)
RhesusBase ^a	A knowledgebase for the monkey research community	Peking University, China	(29)
Rice-Map	Rice (<i>japonica</i> and <i>indica</i>) genome annotation database	Peking University, China	(30)
SalmonDB	Genomic information for Atlantic salmon, rainbow trout, and related species	Center for Mathematical Modeling and Center for Genome Regulation (CMM), Chile	(31)
sigReannot	Aquaculture and farm animal species microarray probes re-annotation	INRA - French National Institute of Agricultural Research, France	(46)
UniProt	Protein sequence and functional information	European Bioinformatics Institute (EBI), UK	(32)
VectorBase	Genome information for invertebrate vectors of human pathogens	University of Notre Dame, US	(33)
VEGA	Manual annotation of vertebrate genome sequences	Wellcome Trust Sanger Institute (WTSI), UK	(34)
WormBase	<i>C. elegans</i> and related nematode genomic information	Cold Spring Harbor Laboratory (CSHL), US	(35)

^aDenotes new databases that have become available since last publication (9).

^bDenotes new databases that are not yet integrated into the portal.

http://central.biomart.org/martservice/attributes?datasets=btaurus_snp&config=snp_config

- filters available for a dataset:
http://central.biomart.org/martservice/filters?datasets=btaurus_snp&config=snp_config

Perl (www.biomart.org)

- registry information:
<http://www.biomart.org/biomart/martservice?type=registry>
- datasets available for a mart:
<http://www.biomart.org/biomart/martservice?type=datasets&mart=ensembl>
- attributes available for a dataset:
http://www.biomart.org/biomart/martservice?type=attributes&dataset=oanatinus_gene_ensembl
- filters available for a dataset:
http://www.biomart.org/biomart/martservice?type=filters&dataset=oanatinus_gene_ensembl
- configuration for a dataset:
http://www.biomart.org/biomart/martservice?type=configuration&dataset=oanatinus_gene_ensembl

Please note that the granularity between mart and dataset has been improved in the Java version through the introduction of multiple dataset configs. This facilitates the end-users to browse various views of the same dataset, which are presented through the portal either using a different GUI or subsets of data.

QUERY EXAMPLES

Given the coverage of the current BioMart datasets, many relevant biological questions can be answered. For example, a researcher who has detected potentially pathogenic variants in FGFR2 (ENSG00000066468) from exome sequencing patients may be interested if the same variants have been previously described and if they were associated with the same or similar diseases. To answer this, integrated data from Ensembl can be queried as shown in Table 2 to display all known variants annotated within FGFR2 that are predicted as pathogenic by SIFT (11) and Polyphen (12). The genomic position outputs can be compared to the researcher's variants and the phenotype data used to assess candidacy for their cases. For example, the first batch of results shows a C->G variant at position 121520160 on chromosome 10 that is associated with Apert syndrome (OMIM:176943).

Another common use case that BioMart is used for is to analyse a list of genes to establish whether they are associated with particular protein functions, pathways or diseases more often than would be expected by chance (enrichment analysis). For example, a researcher may have discovered that AURKA, AURKB, AURKC, PLK1, CDK1 and CDK4 are differentially expressed in their experiment and used BioMart's enrichment tool with its default settings to analyse these genes. The results show that these genes are enriched for involvement in the cell cycle, kinase activity and mitotic nuclear division amongst others. Many other real usage examples are documented in our previous paper (10)

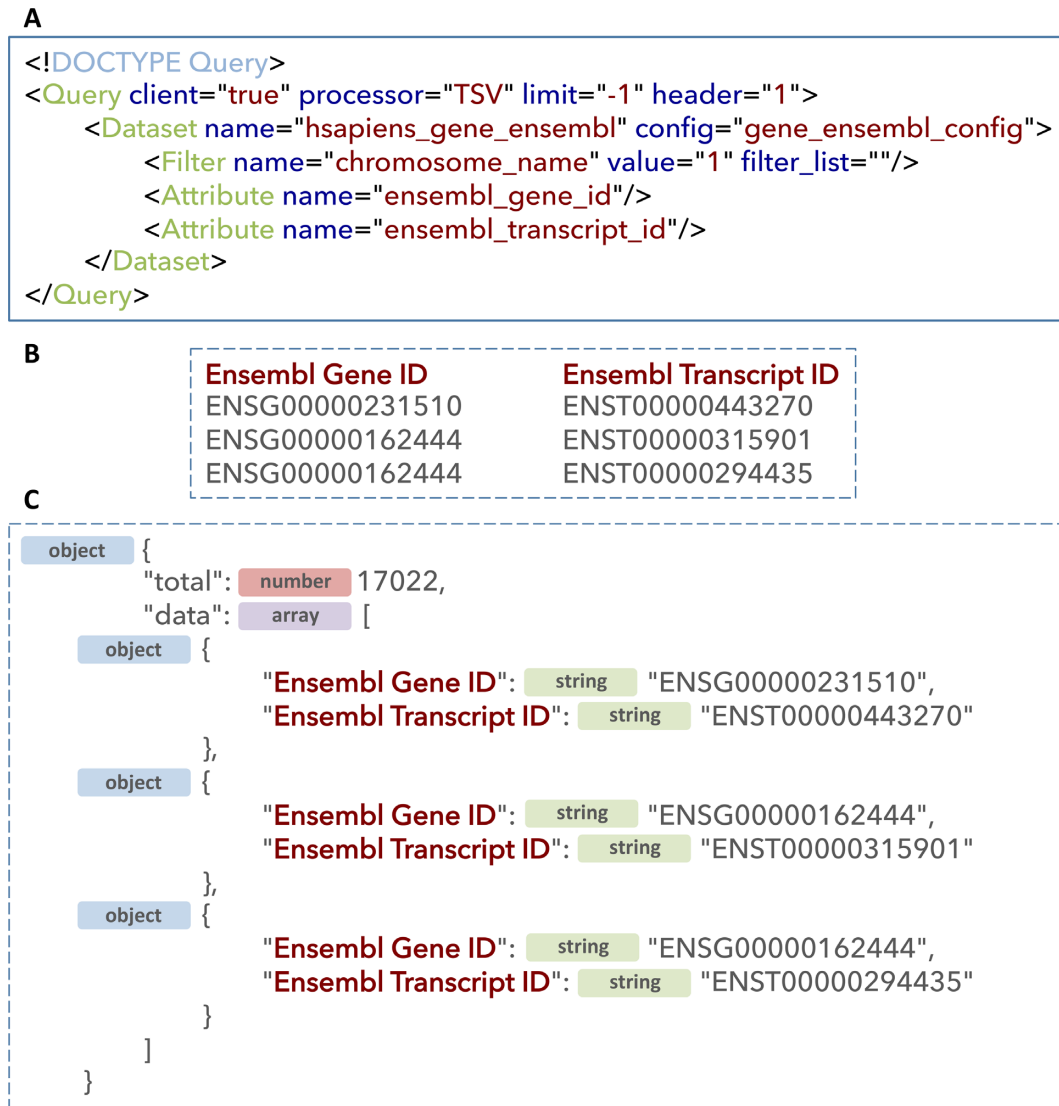


Figure 3. The XML web service query (A) and the corresponding two types of output: tab delimited following setting a processor to ‘TSV’ (B) and JSON following setting processor to ‘JSON’.

Table 2. Query to display phenotypic consequence for known, pathogenic variants in FGFR2

Database and dataset	Filters	Attributes
Ensembl 78 Short Variations (WTSI, UK)	Ensembl Gene ID(s): ENSG00000066468	Chromosome name Chromosome position start (bp) Chromosome position end (bp)
Homo sapiens Short Variation (SNPs and indels) (GRCh38)	SIFT Prediction: deleterious PolyPhen Prediction: probably damaging	Strand Variant Alleles Ensembl Gene ID Consequence to transcript Associated variation names Study External Reference Source name Associated gene with phenotype Phenotype description

and the BioMart special issue in Database: the journal of biological databases and biocuration (www.oxfordjournals.org/our_journals/databa/biomart_virtual_issue.html).

CONCLUSIONS

Since its conception as a data-mining interface for the Human Genome Project (13) BioMart has rapidly grown to become an international collaboration involving a large number of different groups and organizations both in academia and in industry (14). It has been successfully applied to many different types of data including genomics, proteomics, model organisms, cancer data, etc., proving that its generic data model is widely applicable (15–53). BioMart has also provided a first successful solution for the unprecedented data management needs of the International Cancer Genome Consortium proving that the federated model scales well with the amounts of data generated by Next Generation Sequencing (48).

There are a number of important factors that contributed to the BioMart's success and its adoption by many different types of projects around the world as their data management platform. BioMart's ability to quickly deploy a website hosting any type of data, user-friendly GUI, several programmatic interfaces and support for third party tools has proved to be an attractive solution for data managers who were in need of a rapid and reliable solution for their user community. BioMart has also proven to be a platform of choice for many smaller organizations that lack the necessary resources to embark on the development of their own data management solution. As a result, more and more database projects have become accessible through the BioMart interface. The arrival of these new resources coupled with the data federation technology provided by the BioMart software has galvanized the creation of the BioMart Community Portal. The federated model has proven to be very cost-effective since all development and maintenance of individual databases is left to the individual data providers. It also has proven to be very scalable as the internet and database traffic is handled by the local BioMart servers. As a result the BioMart Community Portal service has grown impressively not only in terms of available data but also the level of service. The BioMart community portal now averages over million requests per our services per day. Building on this level of service and the wealth of information that has become accessible through the BioMart interface, the BioMart Community Portal has effectively introduced a new, more scalable and much more cost-effective alternative to the large data stores maintained by specialized organizations.

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Conflict of interest statement. None declared.

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