# The BeeBiome Data Portal provides easy access to bee microbiome information

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# 16 Abstract

- 17 Bees can be colonized by a large diversity of microbes, including beneficial gut symbionts and
- 18 detrimental pathogens, with implications for bee health. Over the last few years, researchers
- 19 around the world have collected a huge amount of genomic and transcriptomic data about the
- 20 composition, genomic content, and gene expression of bee-associated microbial communities.
- 21 While each of these datasets by itself has provided important insights, the integration of such
- 22 datasets provides an unprecedented opportunity to obtain a global picture of the microbes
- associated with bees and their link to bee health. The challenge of such an approach is that
- 24 datasets are difficult to find within large generalist repositories and are often not readily
- 25 accessible, which hinders integrative analyses.
- 26 Here we present a publicly-available online resource, the BeeBiome data portal
- 27 (https://www.beebiome.org), which provides an overview of and easy access to currently
- available metagenomic datasets involving bee-associated microbes. Currently the data portal
- 29 contains 33,678 Sequence Read Archive (SRA) experiments for 278 Apoidae hosts. We present
- 30 the content and functionalities of this portal.

# 31 Importance

- 32 By providing access to all bee microbiomes in a single place, with easy filtering on relevant
- 33 criteria, Beebiome will allow faster progress of applied and fundamental research on bee biology
- and health. It should be a useful tool for researchers, academics, funding agencies, and
- 35 governments, with beneficial impacts for stakeholders.

#### Introduction 36

37 A multitude of factors contribute to bee declines worldwide, but microbes have been identified to 38 play a major role. From pathogens, like viruses or fungi causing severe diseases, to beneficial gut bacteria important for protection against pathogens or digestion of nutrients, microbes play a 39 40 key role in bee fitness and survival. Sequencing efforts around the world have contributed to a 41 better understanding of the genetic and functional diversity of microbes associated with bees 42 (1). These datasets are publicly available from dedicated databases, and notably include 43 amplicon sequences, isolated genomes, shotgun metagenomes, and transcriptomes. The rapid 44 accumulation of such datasets offers new opportunities for data integration and cross-study 45 analyses (2). However, such approaches are hampered by the lack of standardization in dataset 46 annotation which makes it difficult to systematically (and automatically) search for all sequence 47 resources of one data type.

48

49 There is agreement in the community that a centralized bioinformatics tool which would 50 systematically catalog and provide access to sequence datasets from bee-associated microbes 51 would be a useful resource for both fundamental and applied research (1). One way how this 52 can be achieved is via a data portal. A data portal is an online platform which provides access to 53 data, not by storing the data directly, but by systematically cataloging and linking datasets 54 deposited elsewhere. Many large-scale, data-driven projects in biology have dedicated portals, 55 for example the TARA Oceans data portal (3). There are also portals which provide access to 56 data unified by a common theme, even though they were generated by multiple projects with no 57 prior coordination. For example HumanMetagenomeDB (4) provides access to public human

- 58 metagenomes, by organizing the relevant metadata. As raw data is deposited in the open
- 59 databases of NCBI, EBI, and DDBJ (5–7), access to metadata linked to that raw data can be
- 60 sufficient to empower users.
- 61 Here, we present the BeeBiome data portal, which automatically collects and systematically
- 62 stores metadata of publicly available DNA and RNA sequence datasets of bee microbiome
- 63 projects and hence makes them readily accessible to the growing research community working
- 64 on bee-associated microbes. This portal will facilitate data integration and cross-study analyses
- with the ultimate goal to understand the ecology and evolution of bee-associated microbes and 65
- 66 viruses, and advance our understanding of their impact on bees and bee health, from managed
- 67 honey bees to solitary wild bees.

#### **Results** 68

#### Content 69

70 The Beebiome portal integrates metadata from all NCBI(7) available genomic or transcriptomic

71 microbiome data for which the host is identified as Apoidea. Thanks to INSDC data sharing, this

72 also includes all data submitted to ENA(8) or DDBJ(5). As of 5 January 2025, Beebiome

73 contains 30427 BioSamples (unique entries), encompassing 453 Bioprojects and 33678 SRA

74 experiments (Table 1). This represents data from 278 Apoidea host species, which includes honey bees, bumble bees, stingless bees, sweat bees, and carpenter bees (among others). All

data is automatically updated every month (see Data and methods). This offers the advantage
 that newly deposited datasets are integrated into the Beebiome portal on a regular basis.

However, datasets with ambiguous taxonomic annotations can be missing. For example, NCBI

79 Biosamples which are only annotated as "Bombus" without species identification are not

80 included in the Beebiome portal, because "Bombus" refers both to a genus and to a subgenus

81 and hence matches two NCBI TaxIDs: 28641 (genus) and 144708 (subgenus). Such

82 ambiguities need to be avoided to begin with, or corrected at the level of the database to have

83 these samples integrated into the Bebiome portal.

84

85 **Table 1**: High level Beebiome content

86

GSC MIxS or NCBI package name	BioSamples
Metagenome or environmental; version 1.0	12016
MIMARKS: survey, host-associated; version 6.0	7947
MIMS: metagenome/environmental, host-associated; version 6.0	4181
MIMARKS: survey, air; version 6.0	2351
Microbe; version 1.0	1171
Virus; version 1.0	867
MIMARKS: specimen, host-associated; version 6.0	745
MIMARKS: survey, plant-associated; version 6.0	281
MIMS: metagenome/environmental, air; version 6.0	182
MIGS: cultured bacteria/archaea, host-associated; version 6.0	179
Invertebrate; version 1.0	166
Pathogen: clinical or host-associated; version 1.0	92
MIMS: metagenome/environmental, plant-associated; version 6.0	53
MIMS: metagenome/environmental, agriculture; version 6.0	45
MIMARKS: survey, microbial; version 6.0	33
MIGS: cultured bacteria/archaea; version 6.0	32
MIUVIG: uncultivated virus genome, host-associated; version 6.0	29

MIMARKS: specimen; version 6.0	15
Pathogen: environmental/food/other; version 1.0	14
Generic	13
MIGS: eukaryote, host-associated; version 6.0	11
MIGS: cultured bacteria/archaea, agriculture; version 6.0	1
MIGS: cultured bacteria/archaea, human-associated; version 6.0	1
MIGS: cultured bacteria/archaea, miscellaneous; version 6.0	1
MIMS: metagenome/environmental, miscellaneous; version 6.0	1

87

88 For each sample, BeeBiome collects metadata which facilitates searching for relevant datasets

by key words in different categories such as 'Organism' (e.g., *Snodgrassella alvi*) 'Host' (e.g.

90 Apis mellifera), 'Library strategies' (e.g. amplicon or WGS), 'Library sources' (e.g.

91 metagenomic), or 'Collection locality' and 'Collection date'. For example, to identify all amplicon

92 sequence datasets, a user would search for Library source 'Genomic' and Library strategy

93 'Amplicon'. Of note, it is not yet possible to automatically filter the gene amplified, for example to

94 search only 16S rRNA gene amplicons; this information is rarely available. Filtered and sorted

95 data can then be recovered from primary databases through BioProject, BioSample, SRA

96 experiment, or NCBI Nucleotide identifiers, which are all linked back to the source databases.

97 We also store assay type, center name and instrument used (e.g. "Illumina HiSeq 2000"), to

98 allow filtering when relevant.

### 99 Access

100 The primary access to BeeBiome data is through our webpage, at <u>beebiome.org</u>. The

101 homepage provides direct access to a 'basic search', as well as menus to navigate towards an

102 'advanced search', a map, and a wiki. The map simply shows the geographical location of

103 collection for all samples for which this information is available, while 'Browse table' allows to

see the complete table of all data in BeeBiome (**Figure 1**).

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**Figure 1**: Main views of BeeBiome data. (A) "Browse Table" view; (B) Map view.

108

109 The 'basic search' is a query on all metadata. Thus a query for 'Lactobacillus' will return 511 110 entries (as of 5 January 2025). The 'advanced search' allows guerying by specific metadata 111 fields, joined by 'and' (Figure 2). There is also an option to 'browse' all metadata. All primary 112 views are tables which can be filtered by terms; filtering the browse view reproduces the same 113 results as a basic search. All tables can also be sorted by clicking on column names. Which 114 metadata is shown adapts dynamically to the window size, while all other metadata remains 115 accessible by unfolding each row. Whatever is shown, all metadata is used for filtering by terms. 116 The order of metadata columns was established following a poll of the bee microbiome 117 community, to make sure that the relevant information is visible even when window size is 118 limited. As of writing, the first columns are thus: BioProject accession, BioSample accession, 119 SRA experiment entries, NCBI Nucleotide entries, Host, and Organism. It should be noted that 120 for the microbiome of e.g. the honey bee, Apis mellifera is the Host, while the Organism is the 121 microbe or pest (e.g. Snodgrassella alvi) or the type of microbial community (e.g. insect gut 122 metagenome). All search or filtering results, as well as the complete contents of Beebiome, can 123 be downloaded in TSV or copied to clipboard for easy re-use. For example, the results of a 124 guery can be downloaded as a TSV and imported to a spreadsheet software such as MS Excel. 125 from which the Biosample IDs can be copied then used to query NCBI simply by pasting them 126 into the NCBI search. Then they can be simply batch downloaded from e.g. NCBI SRA. The 127 results of advanced search can also be shown on the map (Figure 2C). 128

A			В	Results are Clicking or	ordered by 1 (the '+' sign a	BoProject acc.; the	n 'BioSample acc. ation for each sa	. The order could	i be changed by c	iding on one col	lumn, then pres	s shift and click on	nother column.			
		Advanced search		Show 10	+ entries			9	towing 1 to 10 of 1	13 entries					Files	
		mbine them using a 'AND' boolean logic. More details on each field are available in our			aProject *	BioSample -	SRA experiment	NCBI Nucleotide			Collection	Geo. location	Library	Library	Library	
	BioProject accession	BioSample accession	Host			A00.	entries	entries	Host	Organism	date	name	source(s)	layout(s)	strategy(ies)	instrument(s)
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#### 129

- 130 **Figure 2**: Example of advanced search in BeeBiome. (A) Search for pollen metagenomes in
- 131 host Lasioglossum; (B) table of results; (C) map of results.
- 132 The Beebiome wiki currently contains a comprehensive catalog of Apis and non-Apis diseases
- 133 and microbes, providing an overview of most of the microorganisms known to date (1): Apis bee
- 134 diseases, including known hosts and known effects on hosts; non-Apis bee diseases, including
- 135 known hosts and known effects on hosts; Apis bee microbes; and non-Apis bee
- 136 microorganisms.

# 137 Discussion

138 While generalist microbiome databases can be very powerful (9), the volume and diversity of

- 139 data can be daunting, and make it difficult for small teams and researchers from diverse
- 140 backgrounds to find what they need (2). Thus we also need dedicated database portals to
- 141 organize and access relevant metadata, and allow researchers to easily find datasets of
- 142 interest. For example, HumanMetagenomeDB provides access to standardized metadata for
- 143 human metagenomes (4), and TerrestrialMetagenomeDB to terrestrial metagenomes (10).
- 144 Unlike Beebiome, these databases also include manual curation. Manual curation poses
- problems of sustainability for a small community such as bee microbiome researchers, while our
- automated filtering according to criteria defined by the community allows to keep Beebiome
- 147 updated continuously. Importantly, these criteria can easily be adapted or updated according to
- 148 community needs and feedback. For example, following the discovery by users that our criteria
- 149 could include bee associated beetle pests, we updated these criteria to exclude all of Metazoa
- and Viridiplantae, thus restricting to bacterial and eukaryotic microbes. To avoid manual
- 151 curation, ensure metadata standardization, and enable the Beeebiome portal to correctly detect

- as much data as possible, an important future goal of the community should be to establish
- 153 guidelines for how to deposit beebiome datasets into public repositories (e.g. ENA EBI
- 154 checklist). This would also allow more fine-grained filtering of the datasets. For example,
- 155 datasets coming from different life stages or body sites of the same host species should ideally
- be distinguishable. Also, while the possibility to filter datasets by library source or library
- 157 strategy allows to download datasets of different types, amplicon sequence data e.g. will include
- datasets of amplicons coming from different genes (.e.g rpoB or 16S rRNA gene ) or different
- 159 regions of a given gene.
- 160

161 Another possible area to explore in the future is to provide access to processed datasets, and 162 make analysis tools or pipelines available via the portal. This will facilitate data usage and help

- make analysis tools or pipelines available via the portal. This will facilitate data usage and help in standardizing analysis pipelines as much as possible. We hope that the community will find
- 164 the current tool already helpful and help us to develop the portal further into the directions
- 165 discussed above.

# 166 Data and methods

## 167 Data

168 Beebiome stores metadata on bee microbiomes which come from NCBI Biosample, Bioproject,

and SRA entries (7). Data is retrieved using a Perl script generated by Ebot (11), modified to

170 retrieve relevant metadata. We do not restrict metadata to those which follow a specific

171 standard, such as GSC MIxS (12, 13) or FAANG metadata standards (14). However, NCBI

172 recommends submitting data according to GSC MIxS packages. These packages include

173 attributes defined by the GSC to formally describe and standardize sample metadata. NCBI

submission asks the use either of the GSC MIxS packages or of NCBI packages, forcing the

submitter to give a minimum of information. Entries in Beebiome are represented by the fields

176 detailed in **Table 2**.

BeeBiome entry fields	Standards					
BioProject acc	PRJD# or PRJEB or PRJNA# (NCBI BioProject accession)					
BioSample acc	SAMN# (NCBI BioSample accession)					
SRA experiment entries	Integer					
NCBI Nucleotide entries	Integer					
Assay types	NCBI Strategy enum					
Center name	Free text					

177 **Table 2**: Beebiome entries and corresponding standards.

Library layouts	SINGLE or PAIRED
Library sources	NCBI list, subset to: GENOMIC, TRANSCRIPTOMIC, METAGENOMIC, VIRAL RNA, or OTHER
Organism	NCBI Taxonomy scientific name
Host	NCBI Taxonomy scientific name
Intrument	NCBI Instrument enum
Geo. loc. name	Free text
Collection date	YYYY-MM-DD or YYYY-MM or YYYY

178

Figure 3 presents a broad overview of Beebiome generation. The details of Step 1, to retrievemetadata, are as follows:

- Request NCBI Taxonomy: retrieve species under the taxonomic level 'Apoidea'. We use
   scientific names, common names, genbank common names and synonyms for the next
   point (called *all names*).
- Request to NCBI BioSample using request 1 result: retrieve samples having one of the all\_names (any fields), having an attribute named host and having an organism that is not a Metazoa or Viridiplantae.
- 187host[Attribute Name] AND (Apis mellifera OR honey bee OR188European honey bee OR Western honey bee OR bee OR honeybee OR ...)189NOT Metazoa[Organism] NOT Viridiplantae[Organism]

190 A query is built on the template 'host[Attribute Name] AND (<names>) NOT

191 Metazoa[Organism] NOT Viridiplantae[Organism]', where <names> is names 192 contained in all\_names separated by `OR'.

- 193To avoid network and technical problems due to large files, we do several requests by194generating <names\_separated\_by\_OR> with a maximum of 300 names. Thus the same195data can be recovered in different files (for instance, a BioSample can be retrieved196several times).
- Request to NCBI BioProject, NCBI SRA and NCBI Nucleotide using request 2 result:
   retrieve more metadata and/or links between Biosample and these databases.
- 199 Beebiome is automatically updated every first Saturday of the month, according to these steps.
- 200

# Pipeline



201

- 202 **Figure 3:** Overview of the Beebiome database generation.
- 203 Step 1: Perl script; Step 2: Java API; Step 3: Java API and React webapp.
- 204

### 205 Database and views

206 BeeBiome data is stored in a PostgreSQL database. The API is in Java and Spring boot, and 207 the webapp in React. The API is used to import data from NCBI NCBI XML files:

- 208 1. Read files to put them into NCBI Java objects (built from NCBI XSD files)
- 209
  2. Filter out BioSamples where the host is not one of the *all\_names* (value of the attribute host (which is a free text in NCBI submit format) should be an exact match with one of the *all\_names*)
- 212 3. Convert NCBI Java objects to BeeBiome Java objects (each BeeBiome Java object is
  213 equals to a table into the database)
- 214 4. Save data into database
- An SQL view is generated to save time when there is a request. To generate this view,
   the query filters out biosamples with any SRA experiment.
- 217
- The same API which is used to generated these views also allows to retrieve metadata in JSON format with the following URLs:
- 220 https://beebiome.org/beebiome/sample/all for 'Browse' page, the 'basic
- 221 search' restriction is done by the webapp.
- 222 https://beebiome.org/beebiome/sample/{query} for 'Advanced search' page to
- 223 retrieve entries with a BioSample accession containing {query}

#### Code and data availability 224

- 225 All code is available at https://github.com/BeeBiome-consortium/beebiome-data-portal under
- 226 GPL 3.0 license. All data in Beebiome is distributed under CC0. Other information follows the
- 227 original licenses, e.g. supplemental data from Engel et al (1) is under CC-By-NC-SA 3.0.

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