HMP2Data: Integrative Human Microbiome Data R Bioconductor package

https://github.com/katiasmirn/HMPDataWorkshop

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Integrative Human Microbiome Project (iHMP) https://hmpdacc.org/ihmp/



Pregnancy & Preterm Birth

The Vaginal Microbiome Consortium team at Virginia Commonwealth University is conducting the Multi-Omic Microbiome Study: Pregnancy Initiative (MOMS-PI) in



Onset of Inflammatory Bowel Disease (IBD)

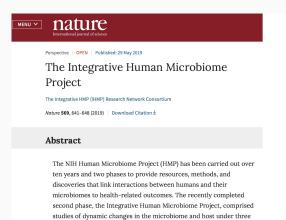
The Inflammatory Bowel Disease (IBD) Multi'omics Data (IBDMDB) research team, led by Ramnik Xavier and Curtis



Onset of Type 2 Diabetes

Type 2 diabetes mellitus (T2D) is a significant health problem facing our nation. In a collaborative effort to systematically understand diabetes and its etiology, the

Novel data: first results just out of press

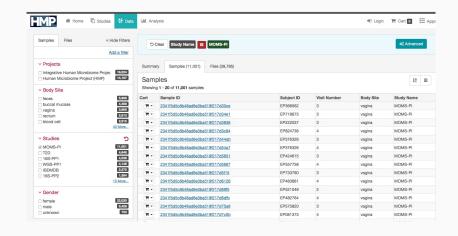


conditions: pregnancy and preterm birth; inflammatory bowel diseases;

and stressors that affect individuals with prediabetes. The associated



Human Microbiome Project Data Portal (https://portal.hmpdacc.org/)



Open data, but...

- Need Aspera client (or similar) to download the data
 - not every lab/researcher has these expertise
- After download still need to:
 - 1. map taxonomy ids from the database
 - 2. merge with meta data not available on the DAC portal
 - 3. construct phylogenetic tree
 - 4. merge -omics modalities

HMP2Data package

- Motivated by HMP16SData (<u>Bioconductor link</u>)
- Currently under review by Bioconductor
- When accepted can be installed using

```
BiocManager::install("HMP2Data")
```

Note: not available yet

Development version

 $In stall\ from\ John\ Stansfield's\ Git Hub$ https://github.com/jstansfield0/HMP2Data

```
if(!require("HMP2Data")){
    BiocStyle::CRANpkg("devtools")
    devtools::install_github(
        "jstansfield0/HMP2Data")
}
library(HMP2Data)
```

Multi-omics microbiome study pregnancy initiavite (MOMS-PI)



(http://vmc.vcu.edu/momspi)

Analysis opportunities

- Early results: term and pre-term birth
- Only a subset of samples was used
- More available through DAC portal
- Novel longitudinal and multi-omics models not explored
- Many opportunities for statistical models development and data exploration!
- 16S and cytokines data

Detailed study design:

https://www.nature.com/articles/s41591-019-0450-2

Data structure

Available as:

- Matrix format for metadata, 16S, taxa table
- Phyloseq
- SummarizedExperiment for cytokines
- MultiAssayExperiment:
 - combine 16S and cytokines

Data objects

16S data as Phyloseq object

```
momspi16S_phyloseq <- momspi16S()
momspi16S_phyloseq

*** phyloseq-class experiment-level object

*** otu_table() OTU Table: [ 7665 taxa and 9107 samples ]

*** sample_data() Sample Data: [ 9107 samples by 13 sample variables ]

*** tax_table() Taxonomy Table: [ 7665 taxa by 7 taxonomic ranks ]
```

Cytokine data as SummarizedExperiment object

```
momspiCyto <- momspiCytokines()
momspiCyto

** class: SummarizedExperiment

** dim: 29 1396

** metadata(0):

** assays(1): ''

** rowmames(29): Eotaxin FGF ... FGF basic IL-17

** rowData names(1): cytokine

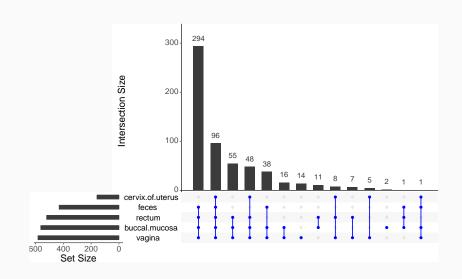
** colnames(1396): EP004835_K10_MVAX EP004835_K20_MVAX ...

** EP996091_K40_MVAX EP996091_K60_MVAX

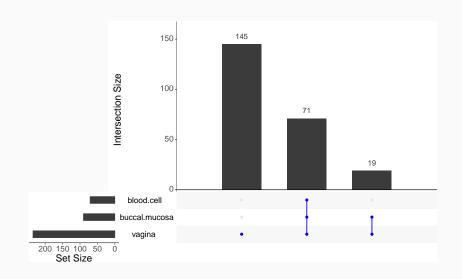
** colData names(13): file_id md5 ... study_full_name project_name
```

	MOMS-PI 16S		MOMS-PI Cytokines		IBD 16S		T2D 16S	
	N	%	N	%	N	%	N	%
Body Site								
buccal mucosa	3313	36.38	311	22.28	0	0	0	0
cervix of uterus	162	1.78	0	0	0	0	0	0
feces	765	8.4	0	0	178	100	1041	47.15
rectum	2679	29.42	0	0	0	0	0	0
unknown	146	1.6	0	0	0	0	0	0
vagina	2042	22.42	979	70.13	0	0	0	0
blood cell	0	0	106	7.59	0	0	0	0
nasal cavity	0	0	0	0	0	0	1167	52.85
Sex								
male	0	0	0	0	84	47.19	1248	56.52
female	9107	100	1396	100	94	52.81	947	42.89
unknown	0	0	0	0	0	0	13	0.59
Race								
african american	0	0	0	0	5	2.81	117	5.3
asian	0	0	0	0	0	0	235	10.64
caucasian	0	0	0	0	164	92.13	1657	75.05
ethnic other	0	0	0	0	8	4.49	73	3.31
hispanic or latino	0	0	0	0	0	0	126	5.71
american indian or alaska native	0	0	0	0	1	0.56	0	0
unknown	9107	100	1396	100	0	0	0	0
total samples	9107	100	1396	100	178	100	2208	100

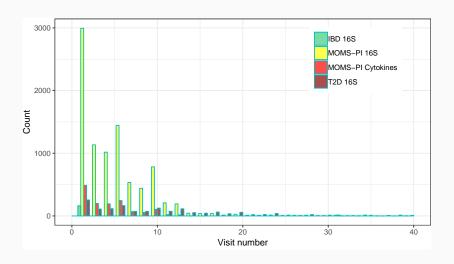
MOMS-PI 16S common samples



MOMS-PI Cytokines common samples



Multiple visits



Metadata (MOMS-PI) "dbGap"



Multi-Omic Microbiome Study-Pregnancy Initiative (MOMS-PI)

dbGaP Study Accession: phs001523.v1.p1

Request Access

Authorized Access

- · Data access provided by: dbGaP Authorized Access
- · Release Date: May 02, 2019
- Embargo Release Date: May 02, 2019
- Data Use Certification Requirements (DUC)
- · Use Restrictions

	Is IRB required?		Number of participants
Disease-Specific (Pregnancy Complications, IRB, PUB, MDS)		NICHD-DAC (<u>HD-DAC@mail.nih.gov</u>)	597

<u>List of components</u> downloadable from <u>Authorized Access</u>

Publicly Available Data (Public ftp)

Connect to the <u>public download site</u>. The site contains release notes and manifests. If available, the site also contains data dictionaries, variable summaries, documents, and truncated analyses.

Note: not available yet

Metadata (IBD): available through the package

Published results:

https://www.nature.com/articles/s41586-019-1237-9

```
Table1Var <- c("subject_gender", "Age.at.diagnosis", "race", "visit_number",
               "site name", "Education, Level", "Antibiotics", "BMI")
demo data = sample data(IBD)[ .c("sample id", "subject id", Table1Var, "diagnosis")] %>%
           data.frame()
#all <- CreateTableOne(vars = Table1Var, data = demo data)
#kableone(all)
stratified = tableone::CreateTableOne(
 vars = Table1Var.
 data = summarytools::unlabel(demo data), strata = "diagnosis", includeNA = TRUE)
stratified <- print(stratified, printToggle = FALSE, showAllLevels = FALSE)
stratified[.!(colnames(stratified) %in% "test")]%>%
 knitr::kable(format = "html", caption = "Characteristics of the Study Cohort",
               col.names = c("Crohn's disease (CD)", "nonIBD",
                             "ulcerative colitis (UC)", "p-value"))%>%
 kable styling("striped", full width = T)
```

	CD	nonIBD	uc	p- value
n	86	46	46	
subject_gender = male (%)	43 (50.0)	24 (52.2)	17 (37.0)	0.264
Age.at.diagnosis (mean (SD))	21.34 (11.31)	NaN (NA)	25.65 (15.21)	0.067
race (%)				0.002
American Indian or Alaska Native	1 (1.2)	0 (0.0)	0 (0.0)	
Black or African American	0 (0.0)	0 (0.0)	5 (10.9)	
More than one race	0 (0.0)	2 (4.3)	2 (4.3)	
Other	4 (4.7)	0 (0.0)	0 (0.0)	
White	81 (94.2)	44 (95.7)	39 (84.8)	
visit_number (mean (SD))	5.33 (10.80)	2.35 (6.39)	1.67 (4.57)	0.034
site_name (%)				<0.001
Cedars-Sinai	21 (24.4)	2 (4.3)	23 (50.0)	
Cincinnati	28 (32.6)	20 (43.5)	11 (23.9)	
MGH	24 (27.9)	18 (39.1)	6 (13.0)	
MGH Pediatrics	13 (15.1)	6 (13.0)	6 (13.0)	

Current and future directions

- Revise the package to include MOMS-PI dbGap data
- Already implemented in HMP16SData
- Package vignettes:
 - 1. HMP2Data package introduction
 - 2. Analysis workflow for MOMS-PI, IBD and T2D studies
 - 3. Multi-omics analysis combining 16S and cytokines data
- June 25: Bioconductor 2019 workshop on HMP16SData and HMP16SData packages