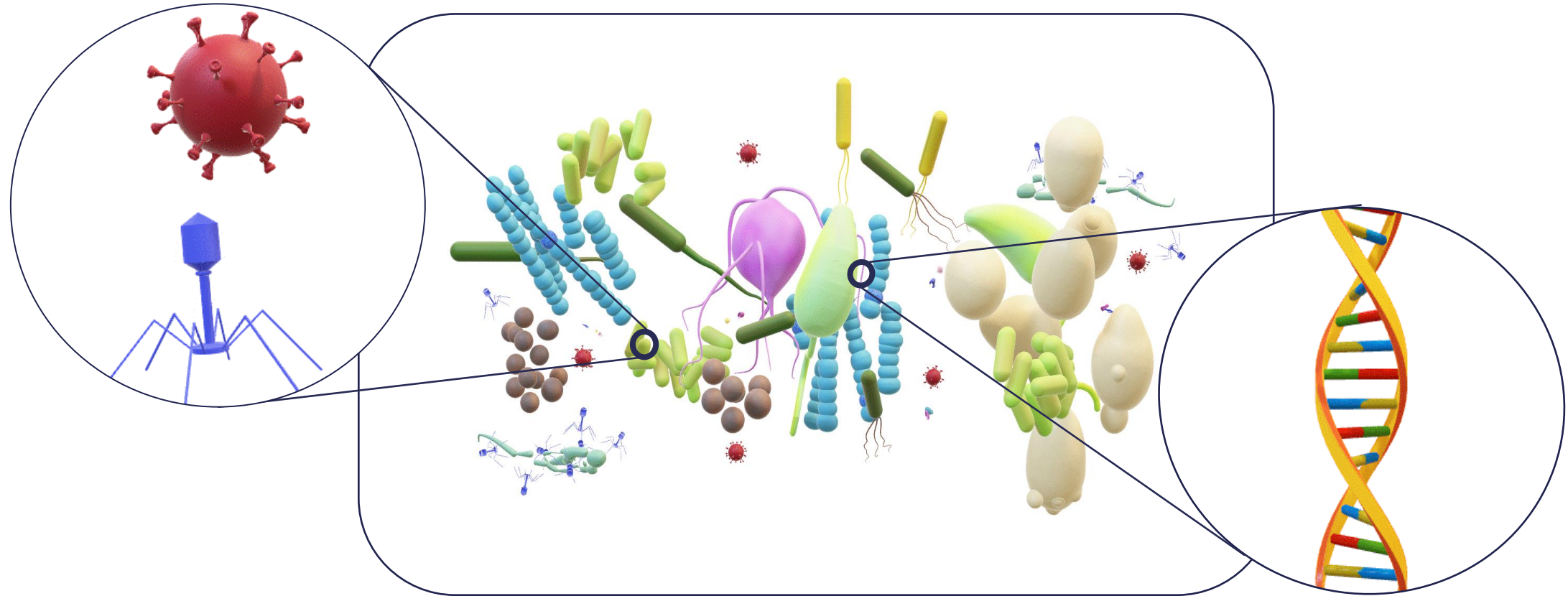


Microbiome research and the meaning of Open Science

Birgit Wassermann and Gabriele Berg
Institute of Environmental Biotechnology
TU Graz

The microbiome

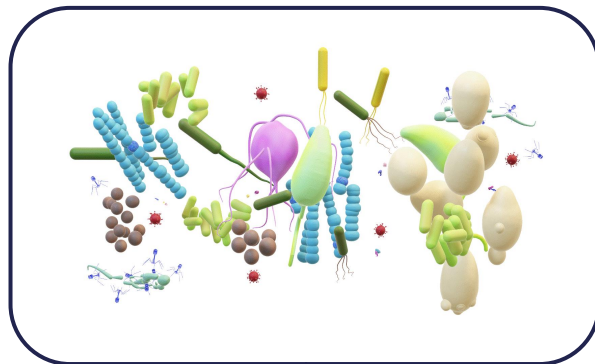


The microbiome encompasses all **microbial organisms** (Bacteria, Archaea, Fungi, Algae, Protists), relic DNA, **Phages, Plasmids, Viruses**, and the **entirety of their functions**.

The microbiome

and the necessity to research

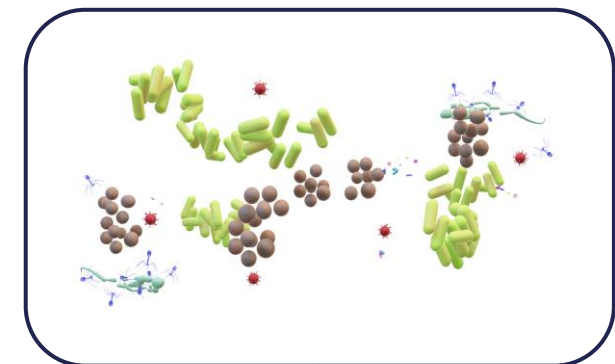
The microbiome is in control of all biogeochemical cycles and interconnects all life on earth.



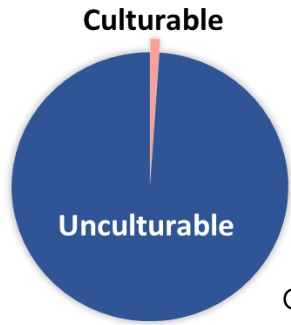
Anthropogenic impact on the microbiome causes:

- Loss of ecosystem stability
- Loss of resilience to abiotic stresses across all life forms
- Spread of plant pathogens
- Increase of chronic diseases
- Increased incidence of human pathogens
- Spread of antibiotic resistance

Planetary health problems result from a massive disconnection between us and nature.



Data explosion



15 years since we can capture the *Unculturable*



Frederick Sanger

1977



Woese

16S rDNA

1980



Kary Mullis (PCR)

Use of 16S rDNA for bacterial taxonomy

1990

illumina

2006

2005

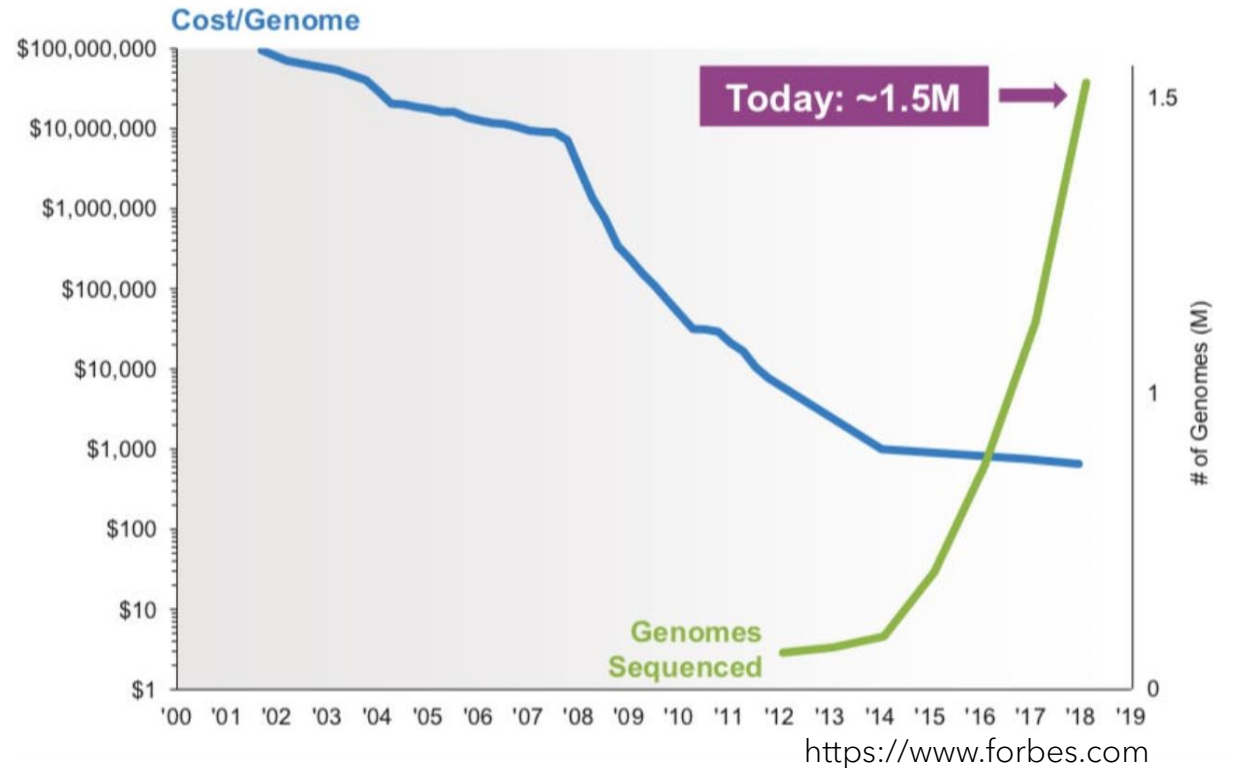
Roche

454

SEQUENCING

High Throughput Sequencing

2020



The omics technologies:

Metagenomics: Who is there and what can they do? → DNA

Metatranscriptomics: What are they doing at the moment? → RNA

Metaproteomics: What is their metabolic potential? → Proteins

Metabolomics: What are they producing? → Metabolites

Microbiome research generates a tremendous amount of data.

All data are publicly available.

Microbiome research



Google Scholar
1,110,000 articles

THE LANCET
173 articles

Science
1,472 articles

Cell
5,393 articles

nature
12,556 articles

<https://mgm.duke.edu/centers-and-cores/microbiome-core-facility>

Publication of microbiome research requires

- ✓ Deposition of the whole dataset in a public repository that can be accessed by everyone world-wide
 - ✓ Provision of metadata
 - ✓ The FAIR concept (Findable, Accessible, Interoperable, and Reusable) is implemented
 - ✓ Upload and download is free of charge
 - ✓ Researchers can choose between three repositories that are interlinked



Operated by the European Molecular Biology Laboratory (EMBL) and supported by European Commission, the British Biotechnology and Biological Sciences Research Council (BBSRC) and the Wellcome Trust (WT)



Operated by the United States National Library of Medicine (NLM), a branch of the United States National Institutes of Health (NIH) and funded by the government of the United States.



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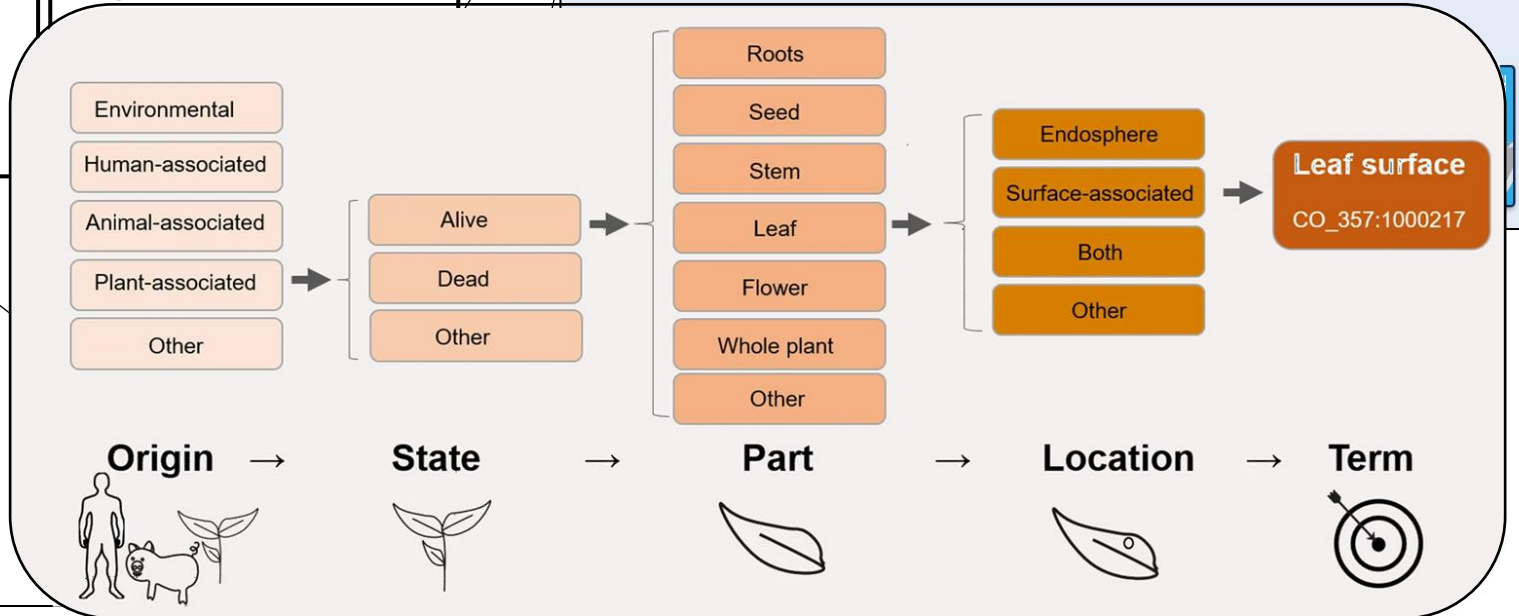
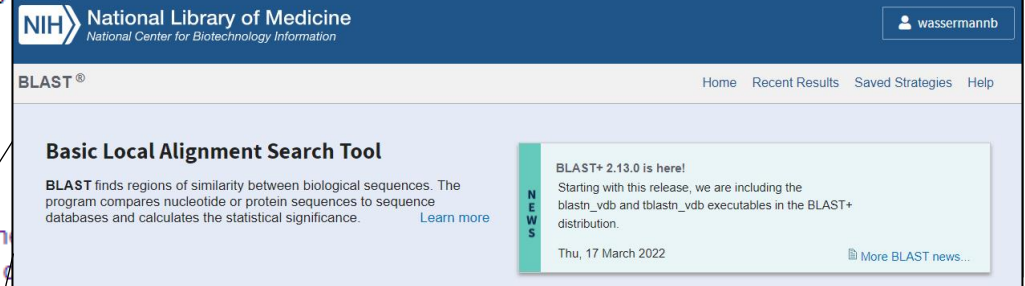
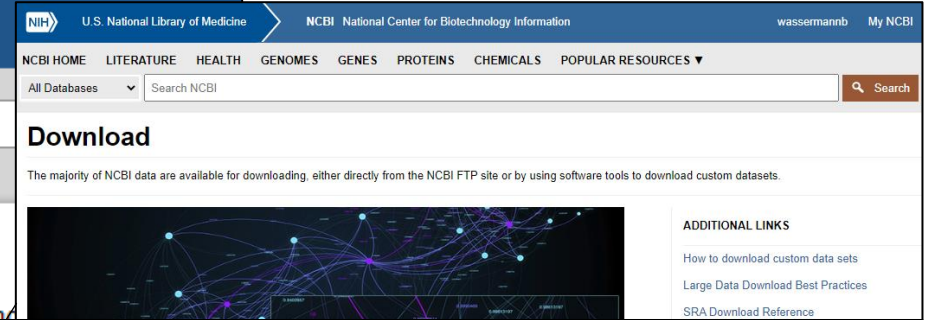
Download

Transfer NCBI data to your computer

Find h class

Develop

Use NCBI APIs and code libraries to build applications

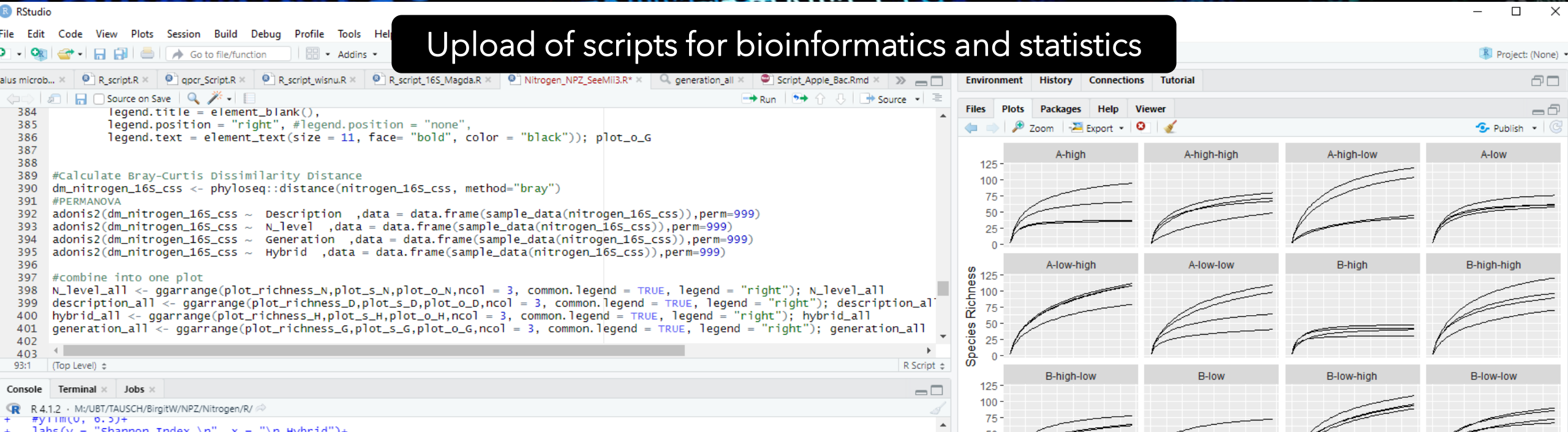


Upload of DNA and RNA sequences



<https://www.statnews.com/>

Upload of scripts for bioinformatics and statistics



The screenshot displays the RStudio environment. The script editor contains R code for calculating Bray-Curtis Dissimilarity Distance and performing PERMANOVA. The console shows the execution of the script. The Environment pane is empty. The Plots pane displays a grid of 12 plots showing Species Richness versus an unlabeled x-axis variable. The plots are arranged in a 3x4 grid, with rows labeled A, B and columns labeled high, low, high-low, low-high. Each plot shows multiple curves representing different samples or conditions.

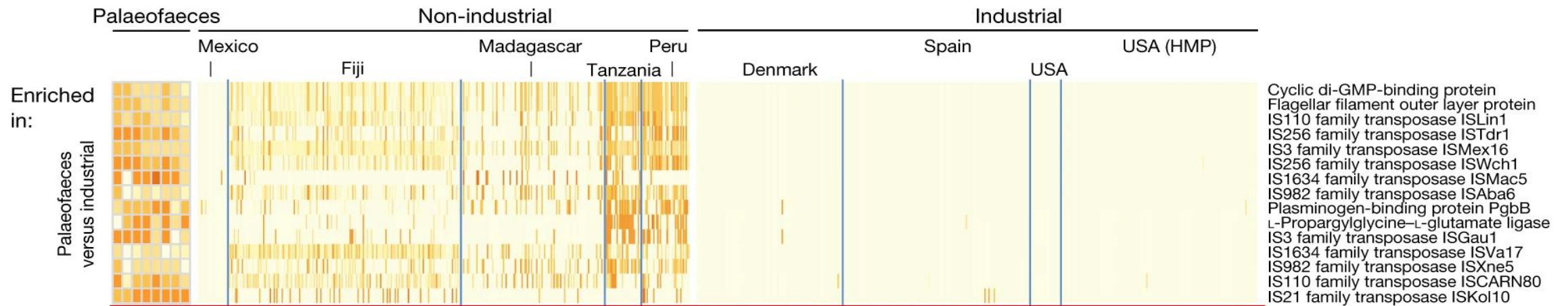
```
384 legend.title = element_blank(),  
385 legend.position = "right", #legend.position = "none",  
386 legend.text = element_text(size = 11, face = "bold", color = "black"); plot_o_g  
387  
388  
389 #calculate Bray-Curtis Dissimilarity Distance  
390 dm_nitrogen_16S_css <- phyloseq::distance(nitrogen_16S_css, method="bray")  
391 #PERMANOVA  
392 adonis2(dm_nitrogen_16S_css ~ Description ,data = data.frame(sample_data(nitrogen_16S_css)), perm=999)  
393 adonis2(dm_nitrogen_16S_css ~ N_level ,data = data.frame(sample_data(nitrogen_16S_css)), perm=999)  
394 adonis2(dm_nitrogen_16S_css ~ Generation ,data = data.frame(sample_data(nitrogen_16S_css)), perm=999)  
395 adonis2(dm_nitrogen_16S_css ~ Hybrid ,data = data.frame(sample_data(nitrogen_16S_css)), perm=999)  
396  
397 #combine into one plot  
398 N_level_all <- ggarrange(plot_richness_N, plot_s_N, plot_o_N, ncol = 3, common.legend = TRUE, legend = "right"); N_level_all  
399 description_all <- ggarrange(plot_richness_D, plot_s_D, plot_o_D, ncol = 3, common.legend = TRUE, legend = "right"); description_all  
400 hybrid_all <- ggarrange(plot_richness_H, plot_s_H, plot_o_H, ncol = 3, common.legend = TRUE, legend = "right"); hybrid_all  
401 generation_all <- ggarrange(plot_richness_G, plot_s_G, plot_o_G, ncol = 3, common.legend = TRUE, legend = "right"); generation_all  
402  
403
```

Re-use of mutiple datasets increases the value of the study

The added value of sharing data

Loss of gut microbial diversity is significantly associated with chronic diseases in industrial populations.

498 genomes from human paleofaeces compared to 789 publicly available present-day gut samples



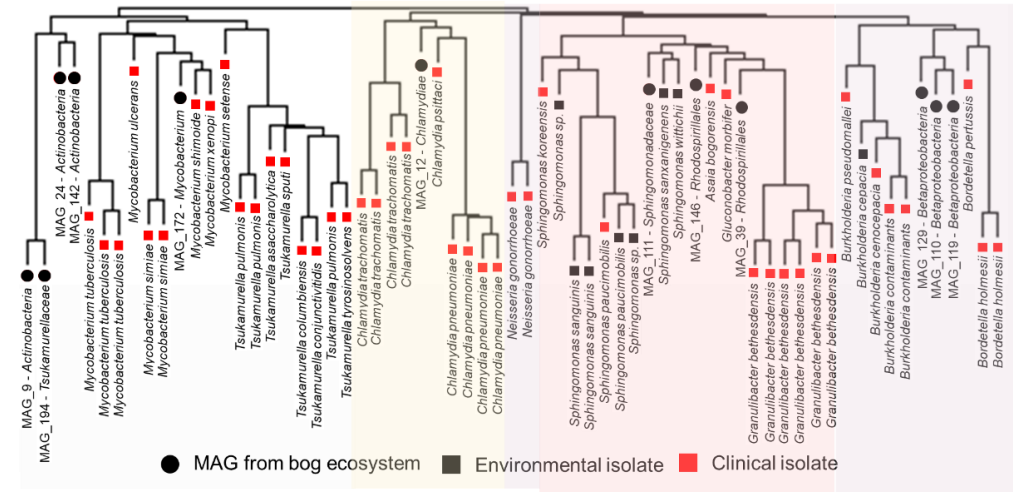
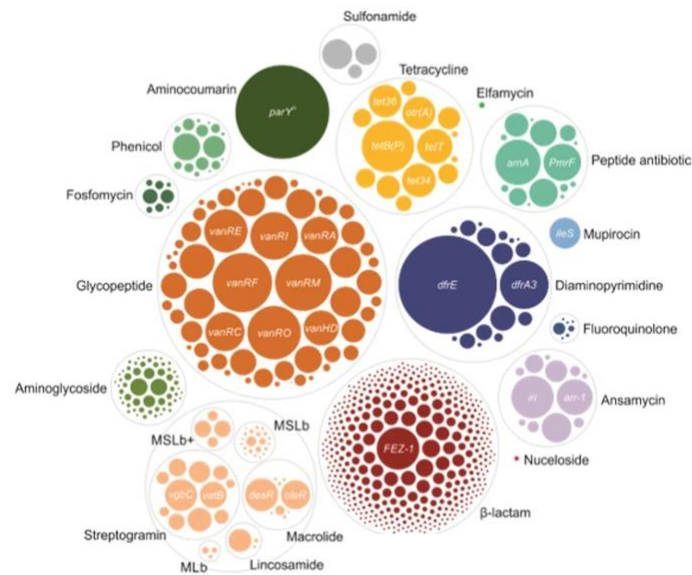
Ancient human stool samples (1000–2000 years old)

- ✓ are more similar to present-day non-industrialized human gut samples and distinct from industrialized samples.
 - ✓ contain important health-beneficial microorganisms
 - ✓ contain less antibiotic-resistance genes

The added value of sharing data

Bog ecosystems store 30% of global carbon. Ongoing global warming will make them one of the largest CO₂ sources.

Austrian bog microbiomes and publicly available datasets from Norway, Germany, and Sweden

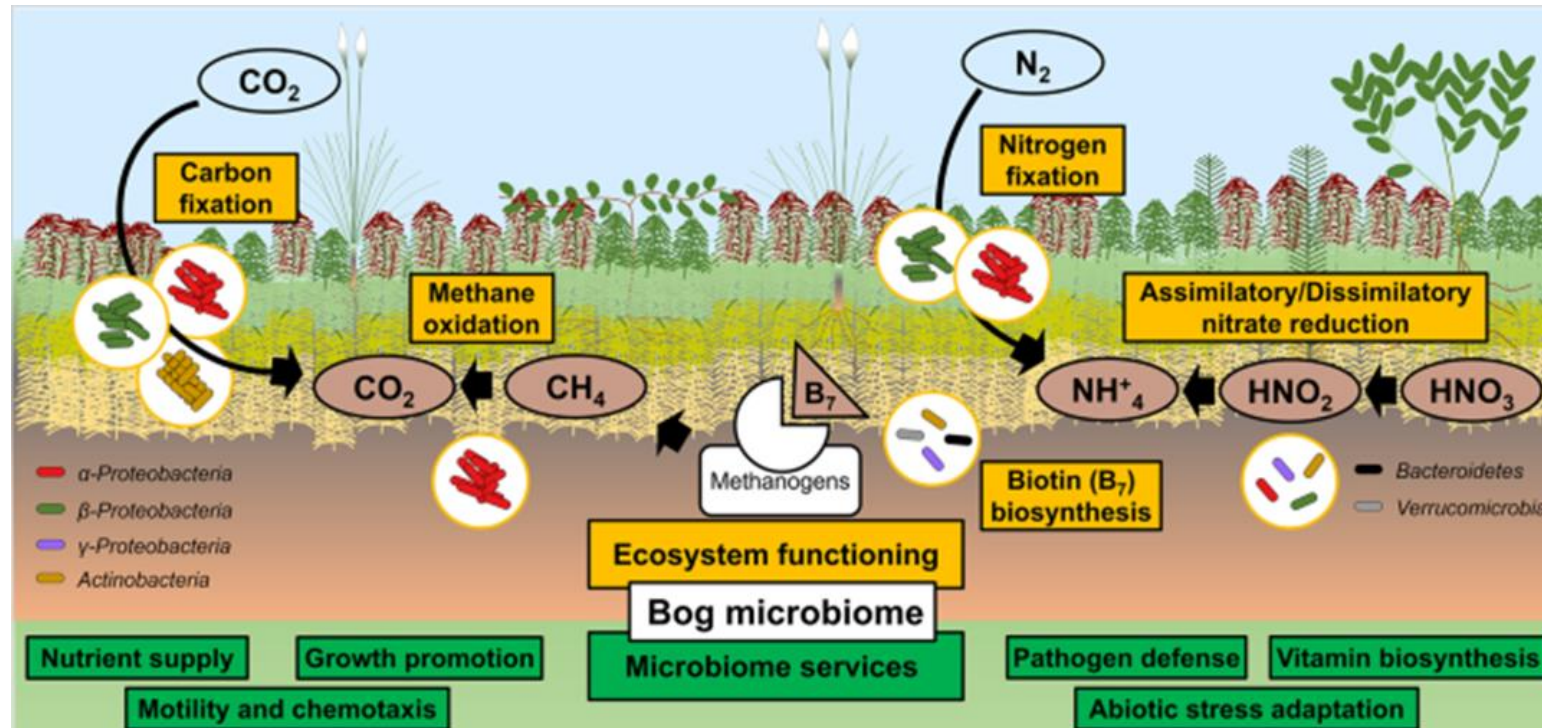


Evolutionary old bog ecosystems harbor

- ✓ a high diversity of antibiotic resistance genes
- ✓ microorganisms with sequence similarity to human pathogens.

The added value of sharing data

Bog ecosystems store 30% of global carbon. Ongoing global warming will make them one of the largest CO₂ sources.



Evolutionary old bog ecosystems harbor

- ✓ microbiomes that are essential for carbon capture
- ✓ plant-beneficial microorganisms that can be used for sustainable agriculture.

The issue with 'data about data'

- ✓ Compared to the massive amount of data, only few products from microbiome research were patented (8,000 patents worldwide).
 - ✓ Generalization of findings by performing integrative analyses is difficult.
- ✓ Metadata should provide all information to repeat a study and to reuse data in a broader context.
 - ✓ A common solution is necessary for this fast-developing research field.

Number of citations, reanalyses, downloads, views, and connections (April 2019)

omics type	Number of citations	Number of cited datasets	Number of reanalyses	Number of reanalysed datasets	Number of downloads	Number of downloaded datasets	Number of views	Number of viewed datasets	Number of connections	Number of datasets with connections
Genomics	8152	3389	1103	872	1,210,799	54,336	1,233,388	13,441	1,041,407,105	313,549
Metabolomics	827	117	–	–	49,907	321	253,428	2726	340,483	1340
Models	3	3	7190	239	–	–	435,859	7262	12,880,012	7200
Multiomics	9111	2053	5013	2422	179,669	2694	860,092	7848	16,453,633	7849
Proteomics	4624	1793	3344	567	153,548	5392	1,417,107	13,015	51,857,985	20,577
Transcriptomics	665,022	50,699	10,527	8062	208,383	3675	14,793,937	119,139	27,696,366	118,500

The availability of datasets led to the most important discoveries in microbiome research for medicine, society and planetary health.

"One of the biggest contributions scientists leave is the potential of their research being reused in the future!"

Tweet by Ulisses Nunes da Rocha (UFZ, Leipzig)

Thank you for your attention!

