THE POTENTIAL OF FIBER MIXTURES TO DELIVER PREDICTABLE AND ROBUST MICROBIOME BENEFITS

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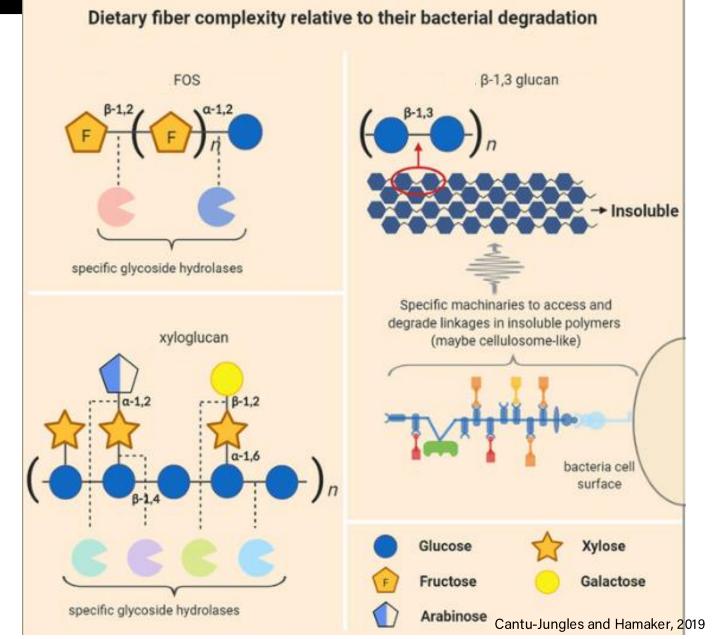
Whistler Center for Carbohydrate Research **Disclosure: RiteCarbs LLC**

Fibers can be selected to promote specific gut bacteria

- Dietary fibers are the main energy source to beneficial gut bacteria
- Bacterial have specific machineries for fiber degradation
- Dietary fibers can stimulate the growth of specific bacteria from the large intestine

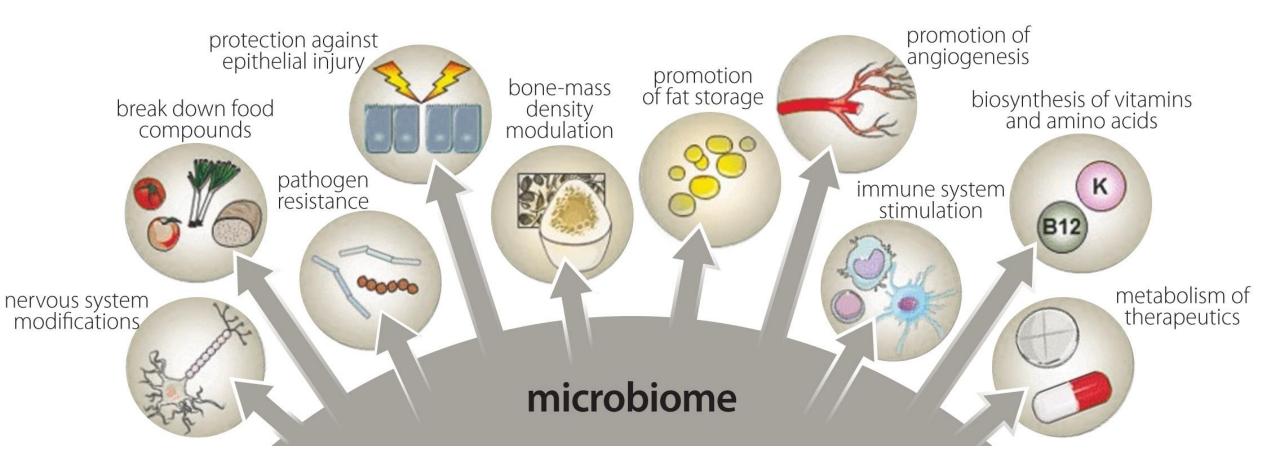


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Which bacteria to promote?

The gut microbiome is a collection of microbes with specific functions

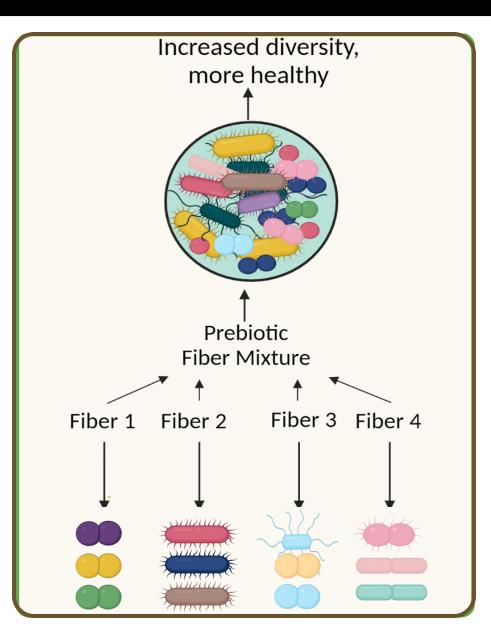




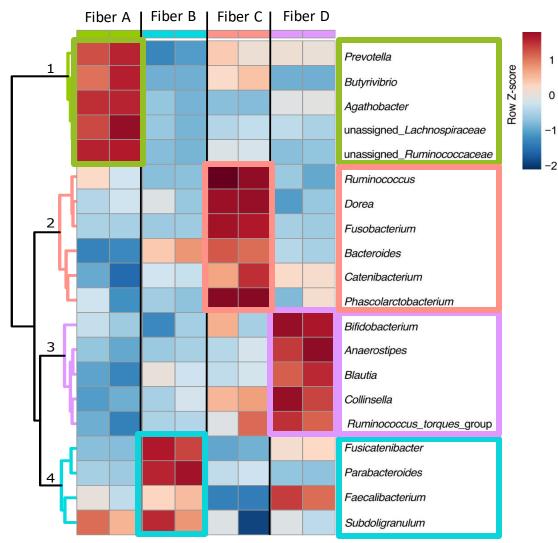
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Design of prebiotics for core beneficial gut bacteria

- Group-matched prebiotics : Fibers that precisely align to groups of taxa
- Blends of group-matched prebiotics : Support of core groups of beneficial bacteria in the gut



Designed prebiotic mixtures for core gut microbiota support – Parkinson Disease



Complementary groups of bacteria and metabolites are promoted with the designed prebiotic mixture*



<u>Nat Commun.</u> 2023; 14: 926. Published online 2023 Feb 18. doi: <u>10.1038/s41467-023-36497-x</u> PMCID: PMC9938693 PMID: <u>36801916</u>

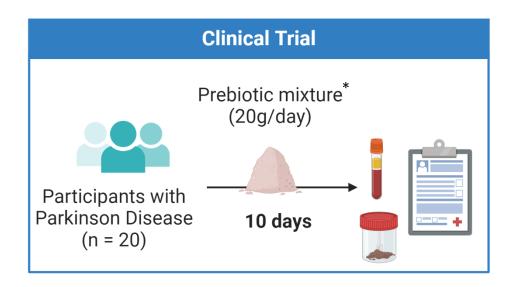
An open label, non-randomized study assessing a prebiotic fiber intervention in a small cohort of Parkinson's disease participants

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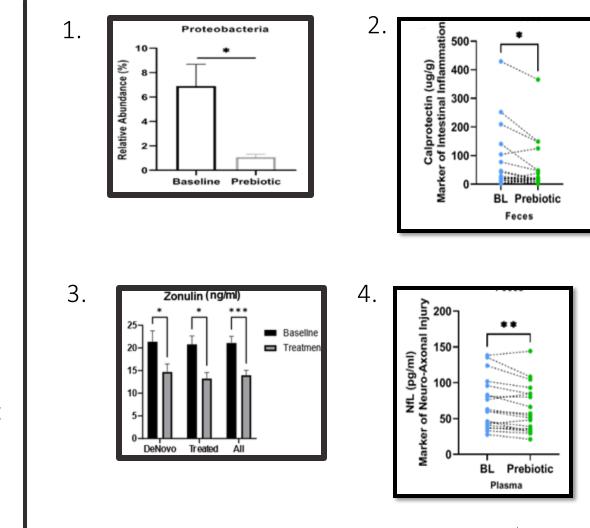
*RiteCarbs (patent pending)

Designed prebiotic mixtures for core gut microbiota support - - Parkinson Disease



✓ Highly tolerable¹

- ✓ Effective¹ in only 10 days to:
 - 1. Promote a healthier balance of bacteria in the gut
 - 2. Reduce intestinal inflammation
 - 3. Improve gut barrier function
 - 4. Reduce brain injury



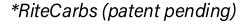
(Hall et al, *Nat. Commun.* 2023)

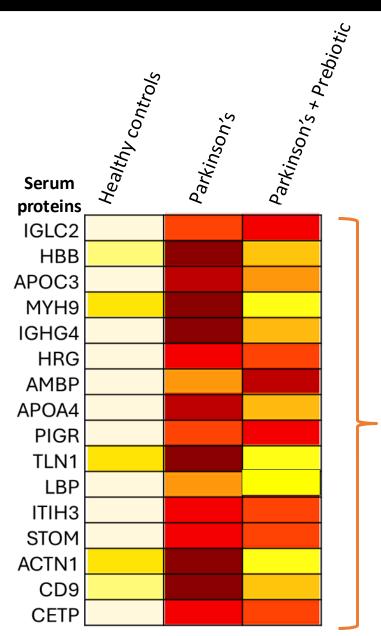
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Designed prebiotic mixtures for core gut microbiota support – Parkinson Disease

Food as medicine

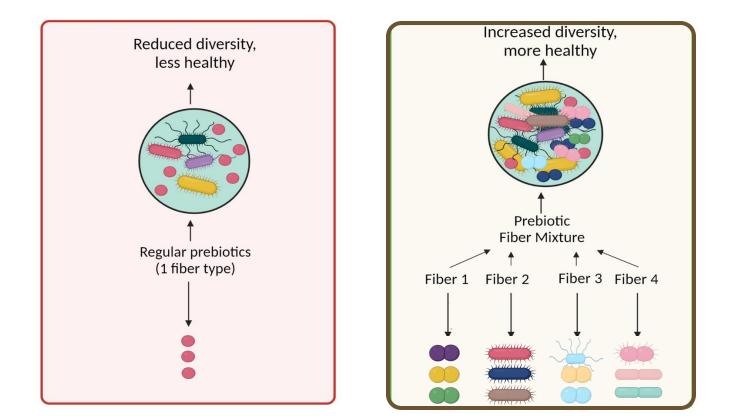
Activation of pharmatargeted biochemical pathways



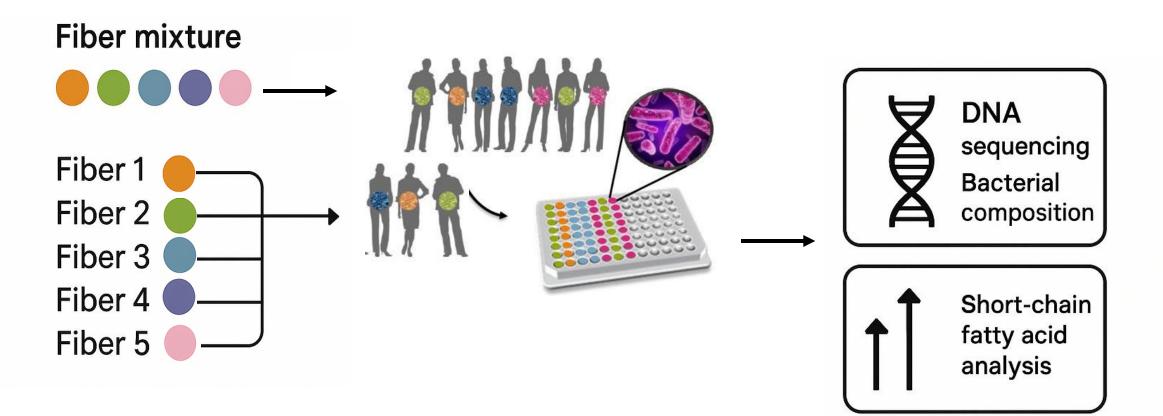


Prebiotic* treatment improved Inflammation Immune response Neural development and repair **Dopaminergic neuron functions Blood-brain barrier** Neurotransmission Lipid metabolism 9

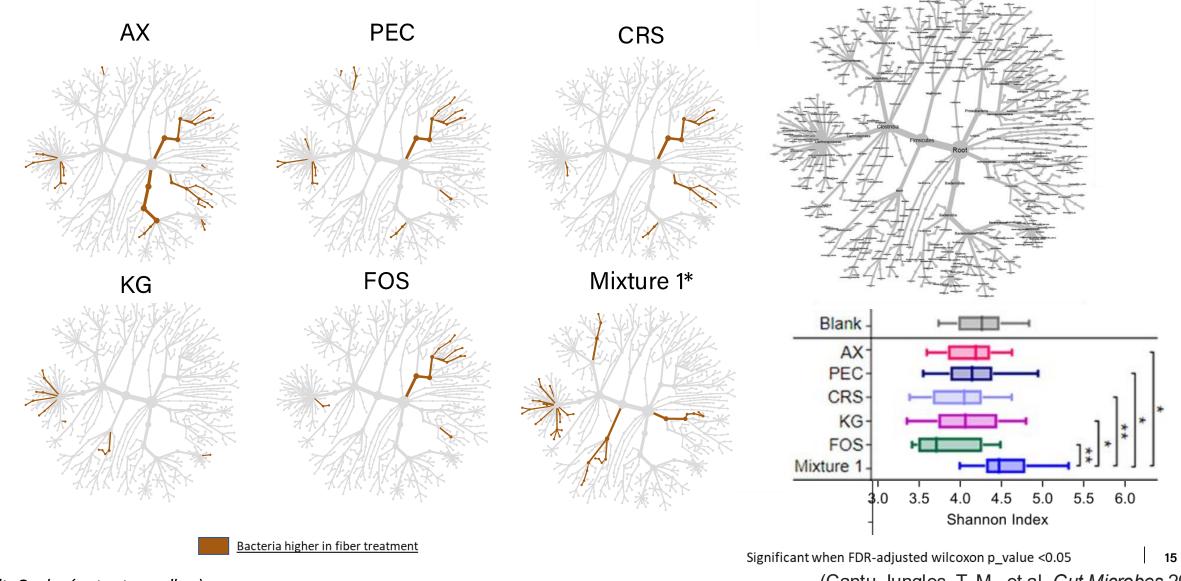
- 1) Provide support to a larger number of beneficial bacteria
- 2) Have more consistent responses in different individuals



Collaboration: Frank Schuren (TNO)



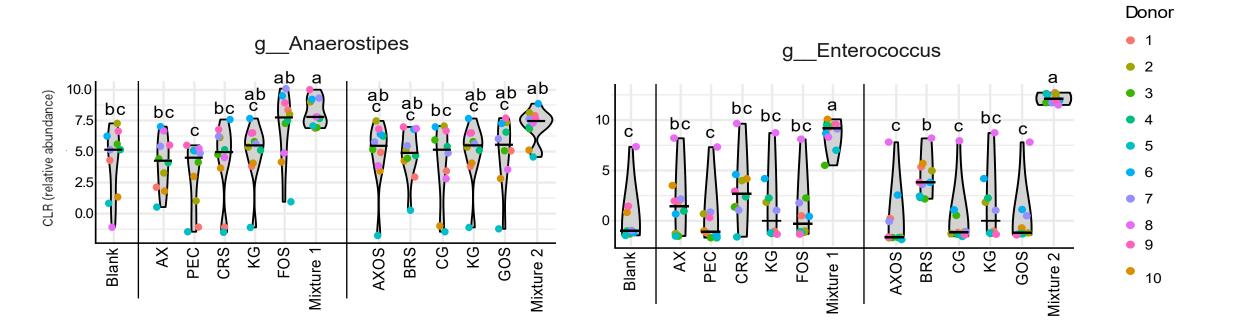




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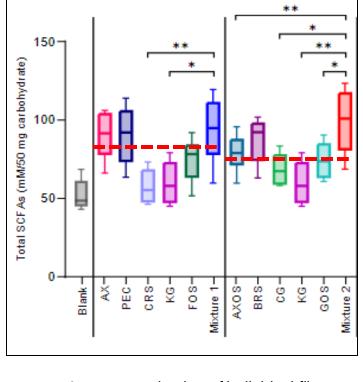
(Cantu-Jungles, T. M., et al. Gut Microbes 2025)

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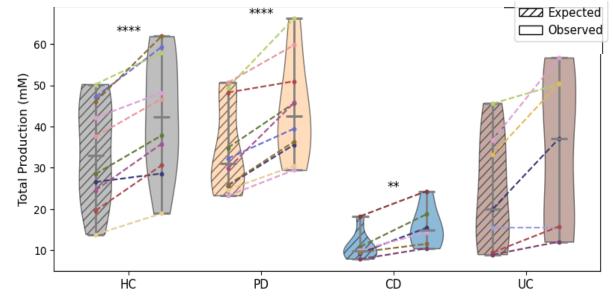
- Provide support to a larger number of beneficial bacteria
 Have more consistent responses in different individuals
 - > Unexpected findings synergistic interactions that:
 - 1) Boost short chain fatty acid production above expected





-- Average production of individual fibers

- <u>Expected SCFAs production</u> with the mixture:
 - Calculated from single *in vitro* fermentation data
 - Assuming additive effects
 - Each of 4 components contributes 25% SCFA produced individually

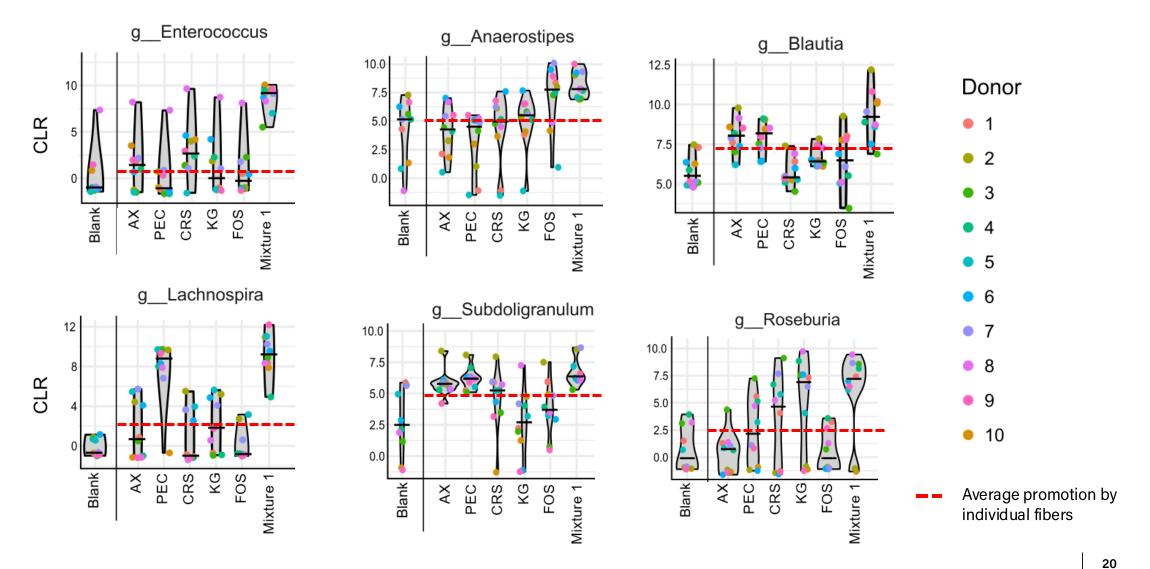


(unpublished data)

HC: healthy controls; PD: Parkinson disease; CD: Chron's disease and UC: ulcerative colitis

- 1) Provide support to a larger number of beneficial bacteria
- 2) Have more consistent responses in different individuals
 - > Unexpected finding synergistic interactions that:
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 - 2) Promote taxa at higher levels than any of its individual components



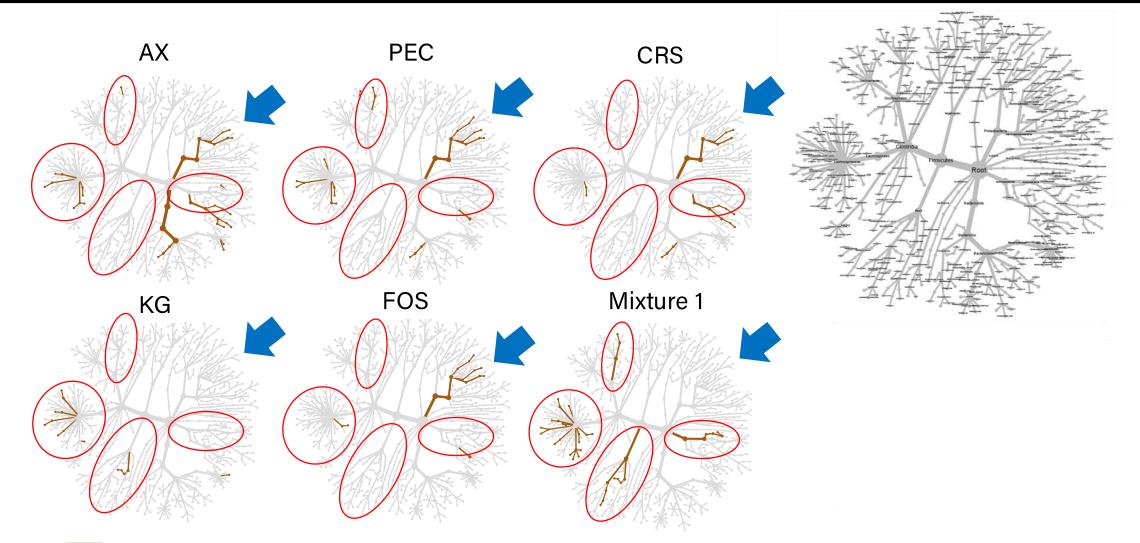


Genus differentially promoted across one or more treatments compared to the blank – identified through ANCOM

(Cantu-Jungles, T. M., et al. Gut Microbes 2025)

- 1) Provide support to a larger number of beneficial bacteria
- 2) Have more consistent responses in different individuals
 - > Unexpected finding synergistic interactions that:
 - 1) Boost short chain fatty acid production above expected
 - 2) Promote taxa at higher levels than any of its individual components
- (3) Taxa that are not supported by any of the mixture components are supported by the mixture



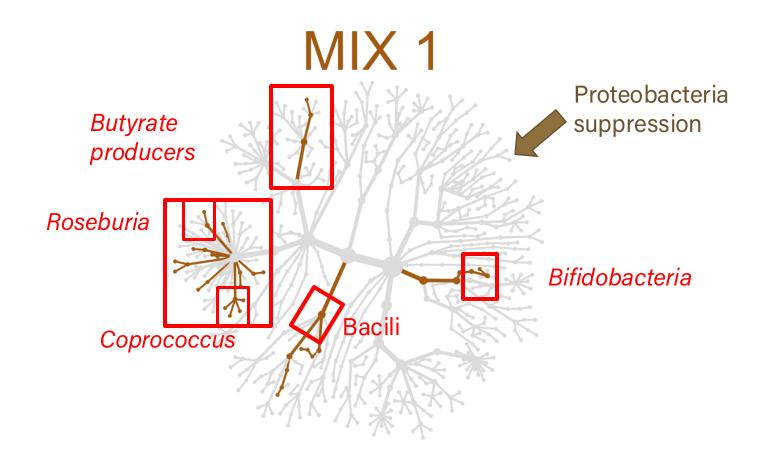


Bacteria higher in fiber treatment

Significant when FDR-adjusted wilcoxon p_value <0.05

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Mixture 1 - Targets



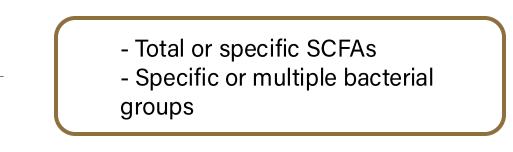
Designed fiber mixtures have potential to lead to:

> Provide support to a larger number of beneficial bacteria

- Promote unique bacterial taxa that might not be promoted with individual fibers
- Prevention of pathogenic bacteria growth
- Consistent responses across people
- Intense responses in the gut microbiota (more effective, act faster?)

Perspectives

- 1) Design fibers mixtures to support specific groups of beneficial bacteria
- 2) Improve synergistic interaction of fibers to promote specific targets



Perspectives

- 1) Design fibers mixtures to support specific groups of beneficial bacteria
- 2) Improve synergistic interaction of fibers to promote specific targets

Total or specific SCFAsSpecific or multiple taxa

Mathematical approaches, machine learning

Knowledge based hypothesis generation

Q&A

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