

Gut-Brain Connection in Parkinson's Disease

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Disclosures

None

An anatomical illustration showing the human digestive system on the left and a human head in profile on the right. The digestive system, including the stomach and intestines, is highlighted in red. The brain is highlighted in pink. A large, curved yellow arrow points from the digestive system towards the brain, indicating a connection or flow of information. The background is dark, making the anatomical structures stand out.

Digestive Disease...

...Neurodegenerative disease

Goal: To heal the aging digestive system and harness its power to combat neurodegeneration

Digestive disorders in Parkinson's disease

Almost 75% suffer

Underrecognized; many are undiagnosed

Common digestive issues in Parkinson's disease

Dysphagia (difficulty swallowing)

Gastroparesis (slow stomach emptying)

Malabsorption

Small intestinal bacterial overgrowth

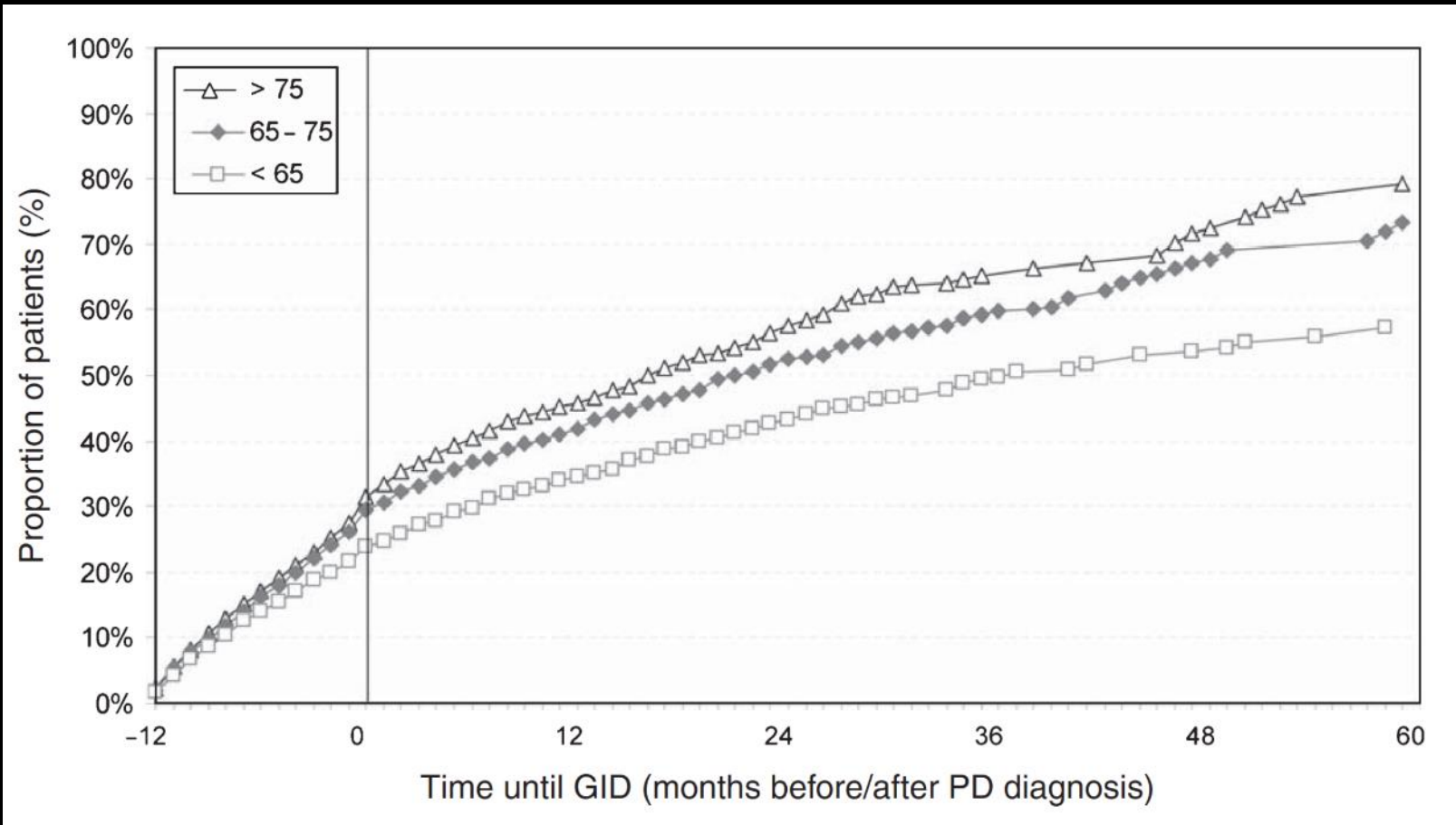
Constipation

Rectal evacuation disorders

Abdominal pain

Weight loss

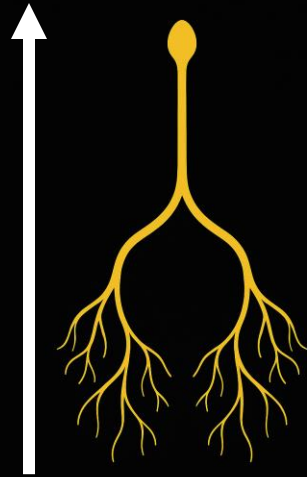
Gastrointestinal dysfunction (GID) in PD



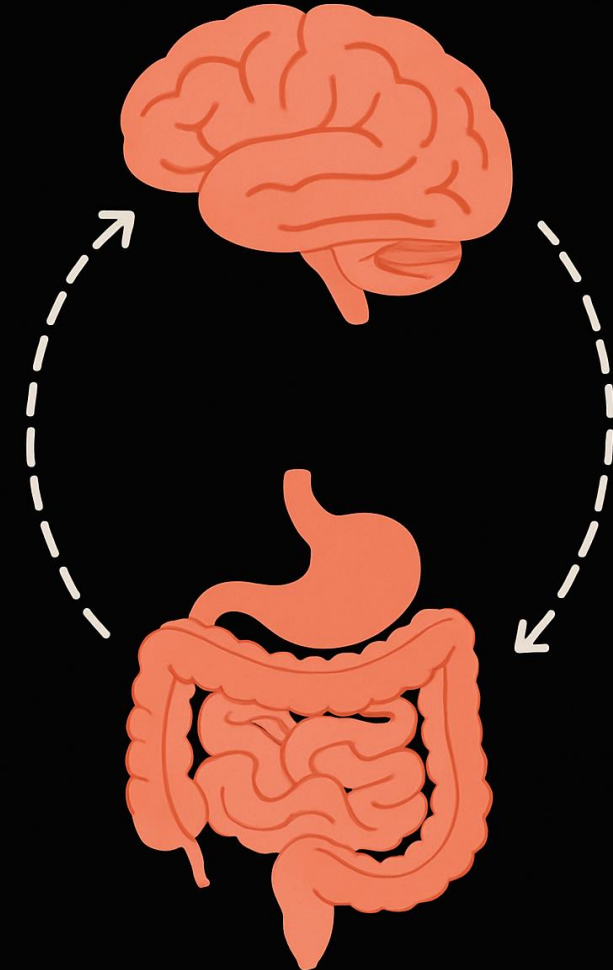
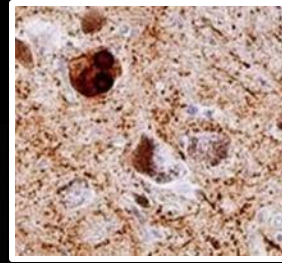
Makaroff, et al. (2011)

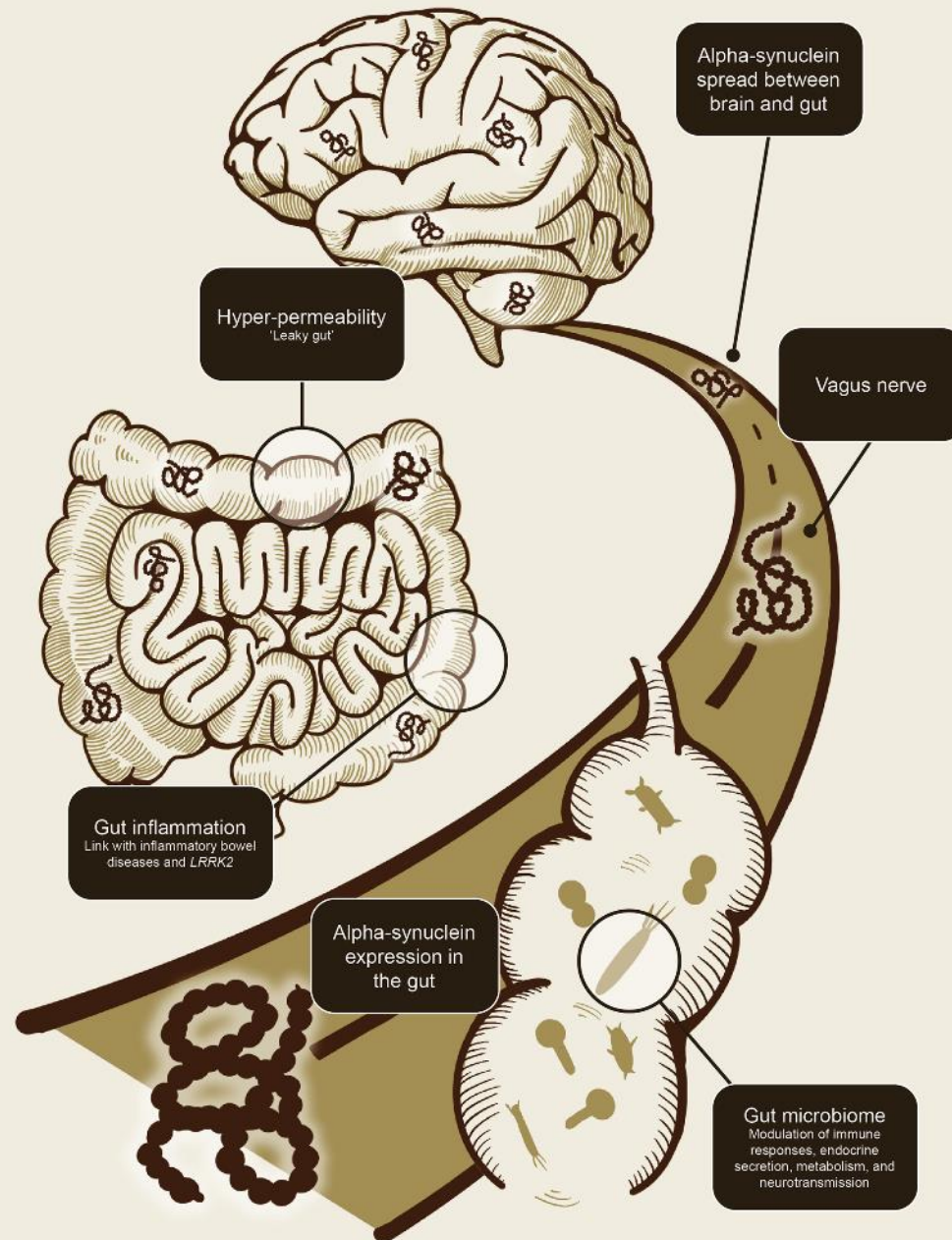
α -synuclein in the brain

Vagus nerve



α -synuclein in the colon





Healthy

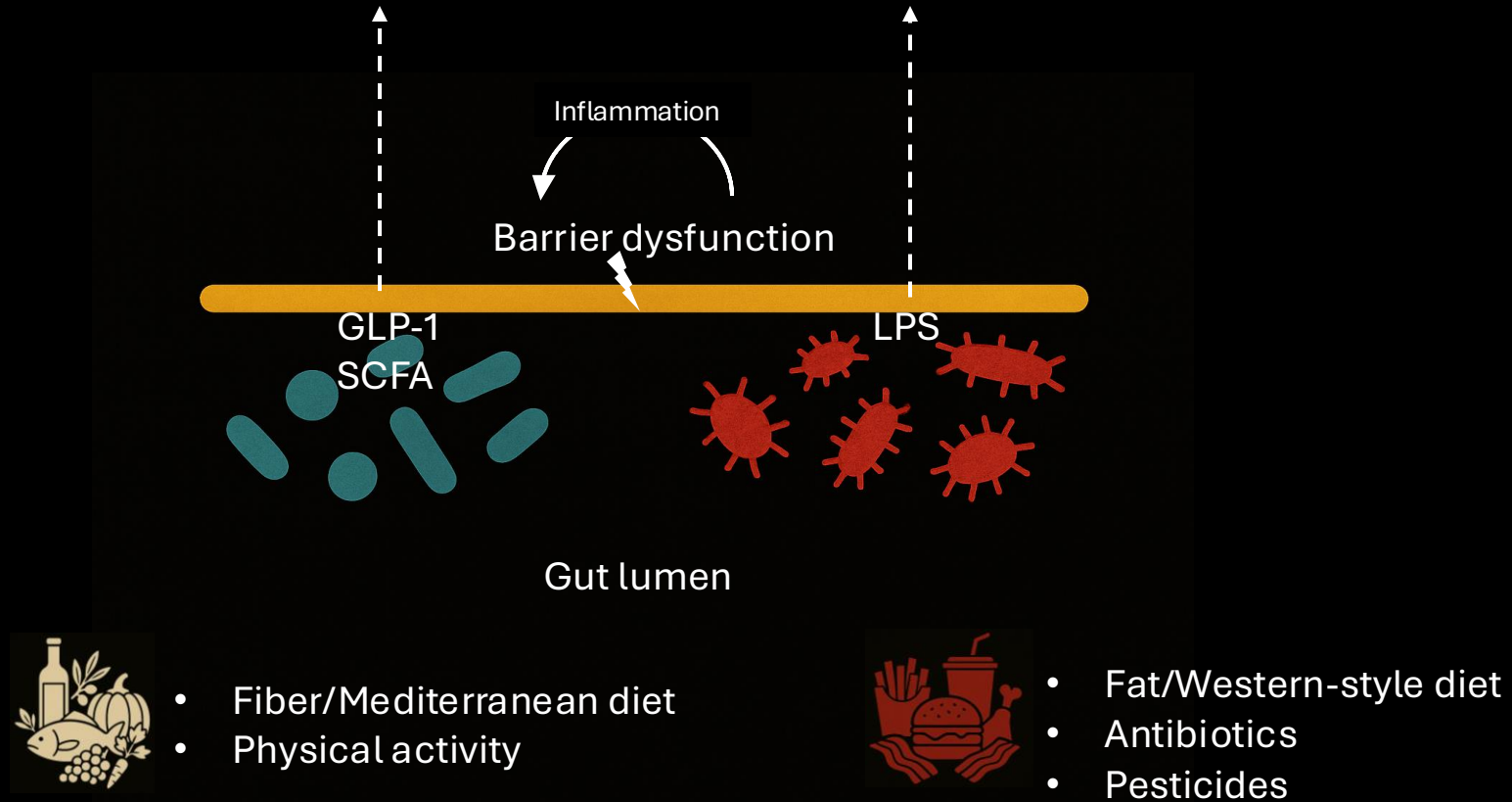


- Neuroprotection
- Low systemic inflammation

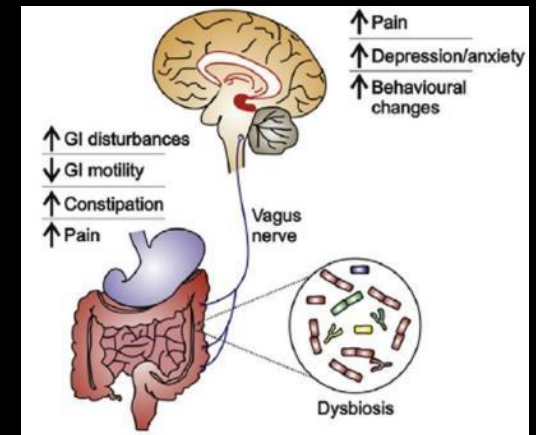
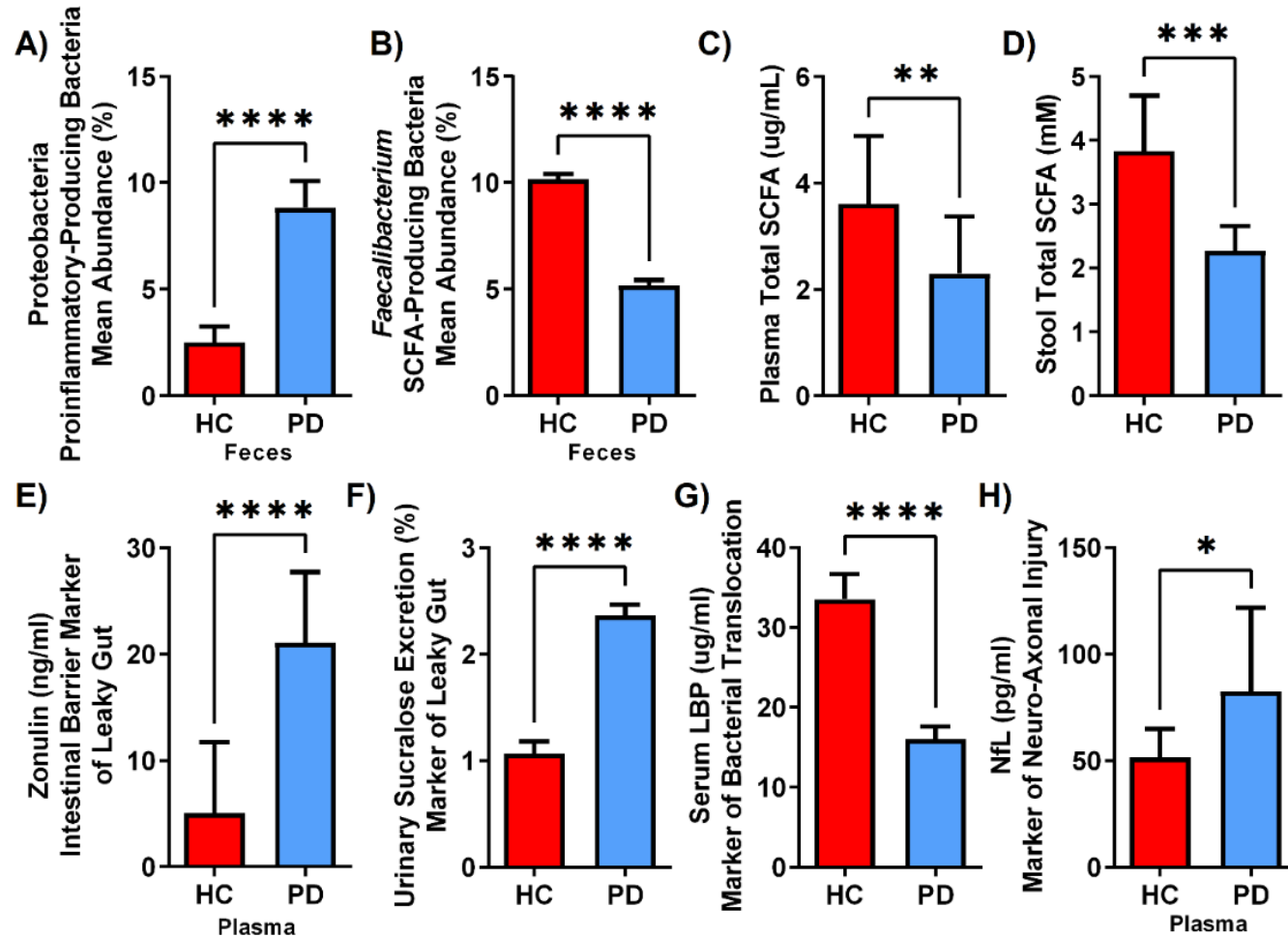
Unhealthy



- Neurodegeneration (microglial activation)
- α -synuclein aggregation
- High systemic inflammation



Gut microbiota changes in PD





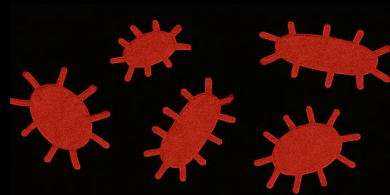
Diet



Environment



Genetics



Altered gut microbiota

Inflammation



Diminished gut
barrier function



Reduced neural protection
by healthy bacteria



Neurodegeneration





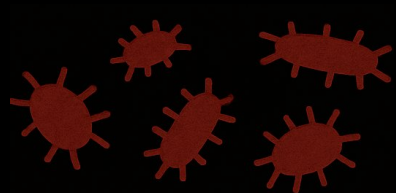
Diet



Environment



Genetics



Altered gut microbiota

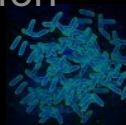
Inflammation



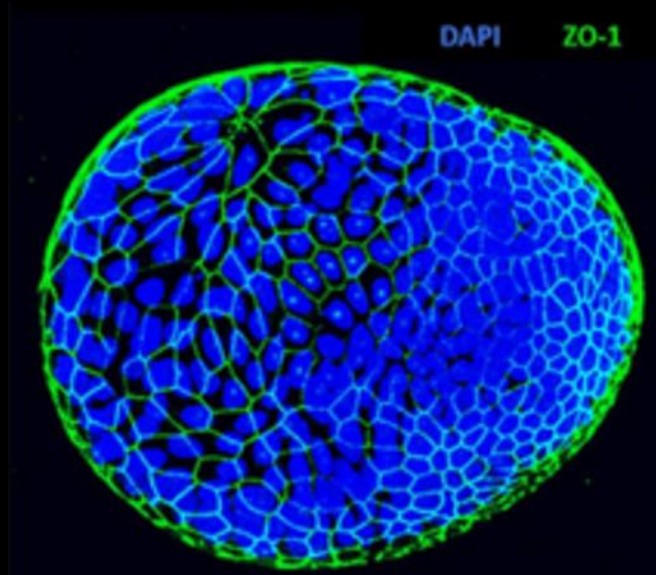
Diminished gut
barrier function



Reduced neural protection
by healthy bacteria



Neurodegeneration





Diet

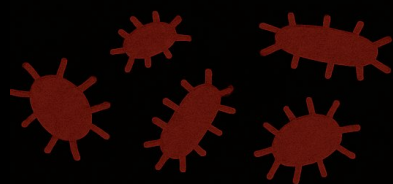


Environment



Genetics

Altered gut microbiota



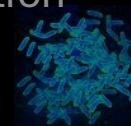
Inflammation



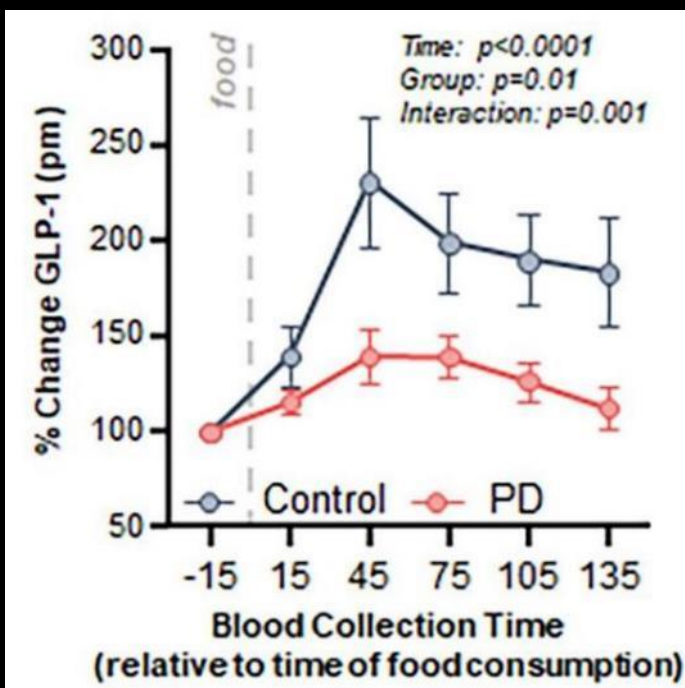
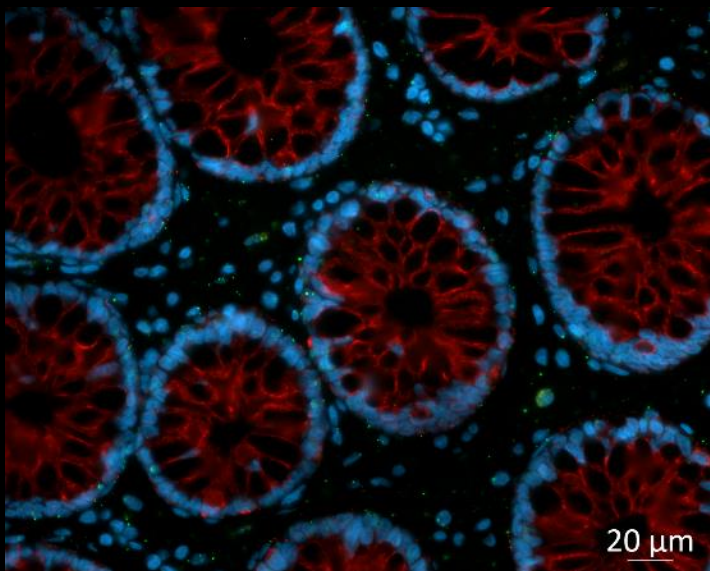
Diminished gut barrier function



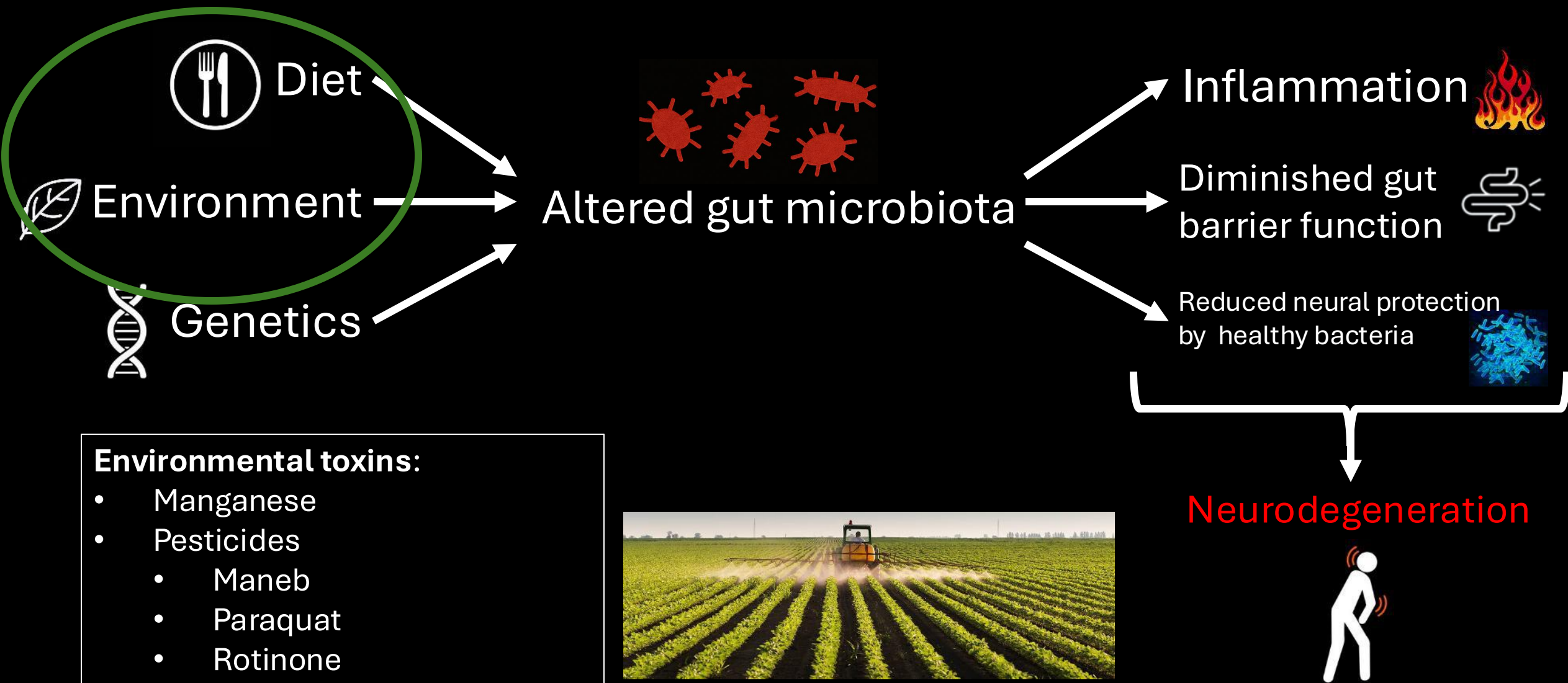
Reduced neural protection by healthy bacteria



Neurodegeneration



Manfreedy, et al. (2021)



Diet

Goal: To fuel the body's 'healthy' microbiota while preventing growth of pro-inflammatory bacteria.

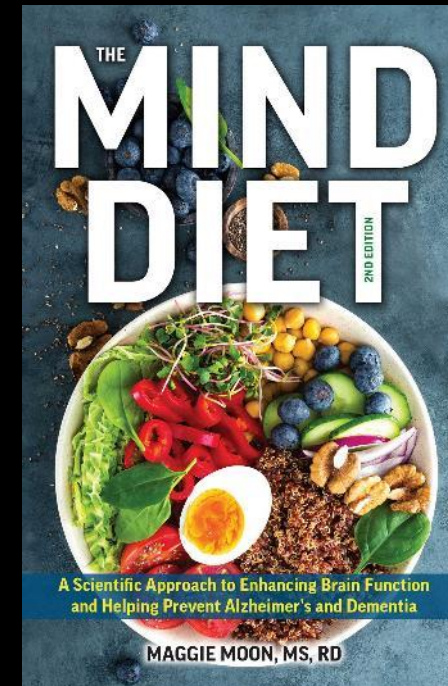
- Green leafy vegetables
- Fruits
- Whole grains
- Legumes
- Nuts
- Lean meats (chicken, fish)
- Olive oil



Mediterranean Diet

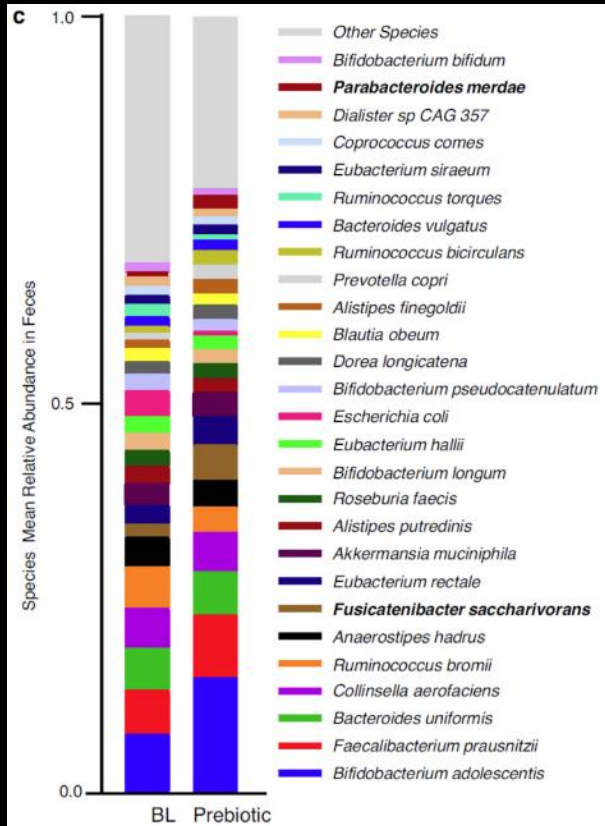
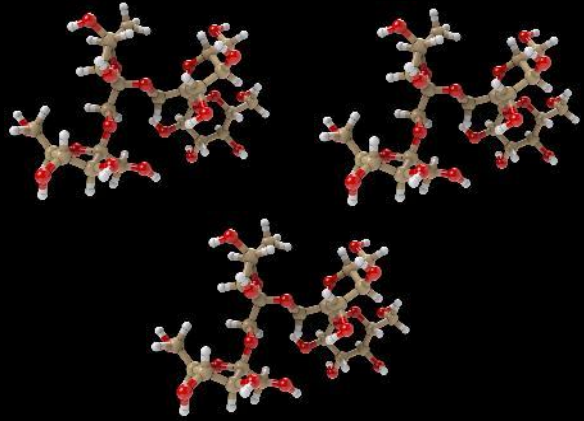


MIND Diet

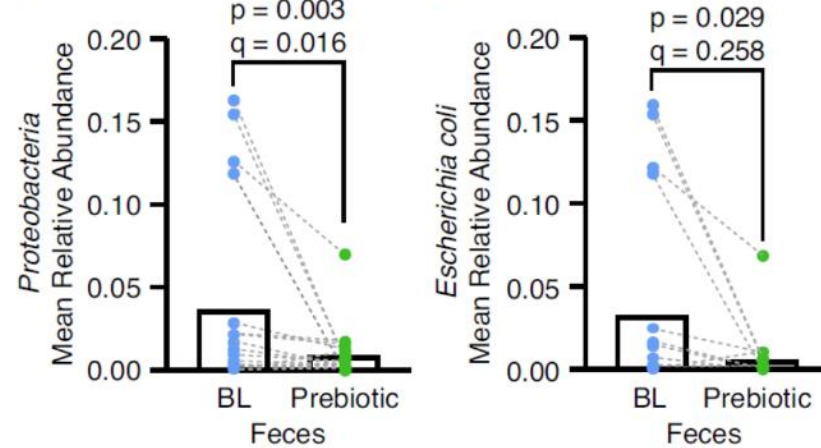


A targeted approach for Parkinson's patients

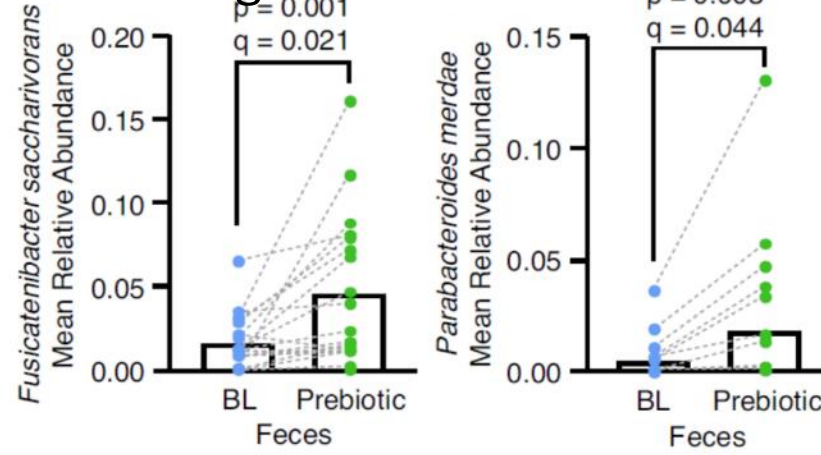
Custom-designed prebiotic fibers



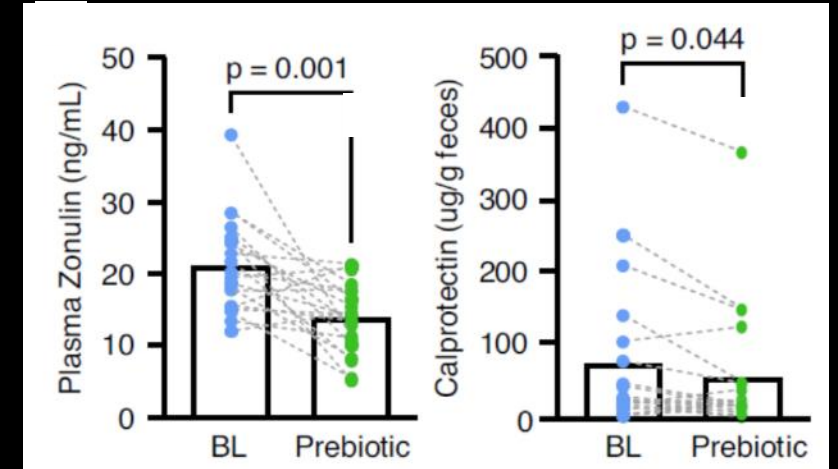
Decrease “bad” bacteria



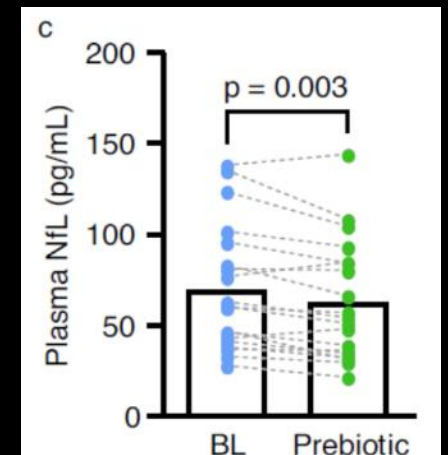
Increase “good” bacteria



Decrease gut inflammation

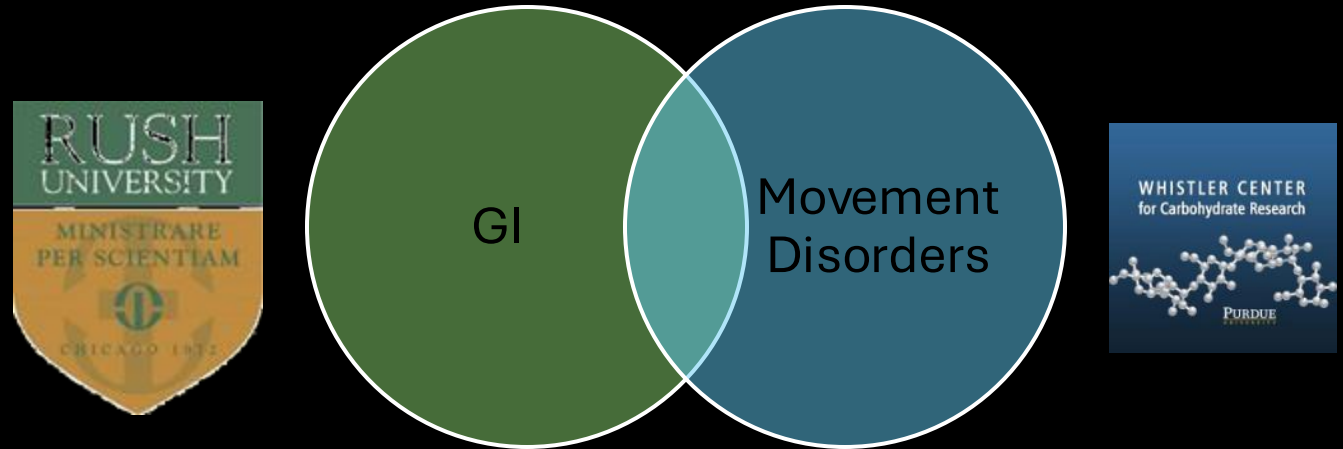


Protect neurons



Dietary strategies tend to **work better for early-stage Parkinson's**.
It is more difficult to restore gut-brain function once there is substantial brain involvement in advanced disease.

Experience from PD-GI Clinic



Constipation and bloating are major hurdles to dietary tolerance in advanced disease.

PD Constipation



α -synuclein accumulation

Inflammation

Gut barrier dysfunction

Impaired intestinal motility

Pelvic floor dyssynergia

PD Constipation



α-synuclein accumulation

Inflammation

Gut barrier dysfunction

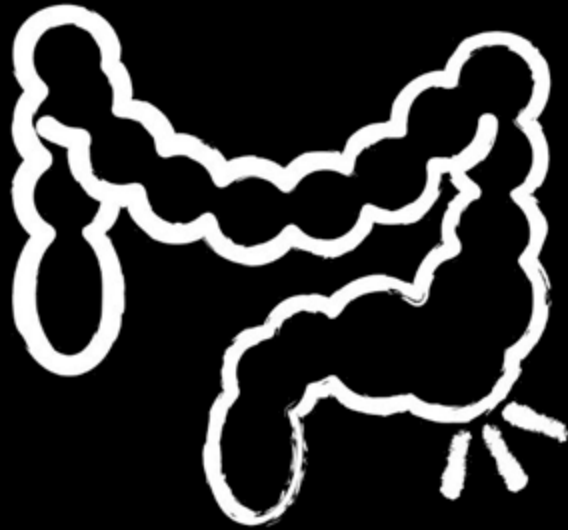
Impaired intestinal motility

Pelvic floor dyssynergia

Distinct entity from IBS

5 or fewer spontaneous bowel movements (not aided by medications) a week with **lack of complete evacuation**.

PD Constipation



α -synuclein accumulation

Inflammation

Gut barrier dysfunction

Impaired intestinal motility

Pelvic floor dyssynergia

- Diet
- Exercise
- Special laxatives to reduce gut-brain inflammation

CLINICAL TRIALS ONGOING

PD Constipation



α -synuclein accumulation

Inflammation

Gut barrier dysfunction

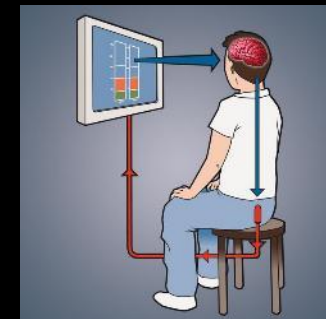
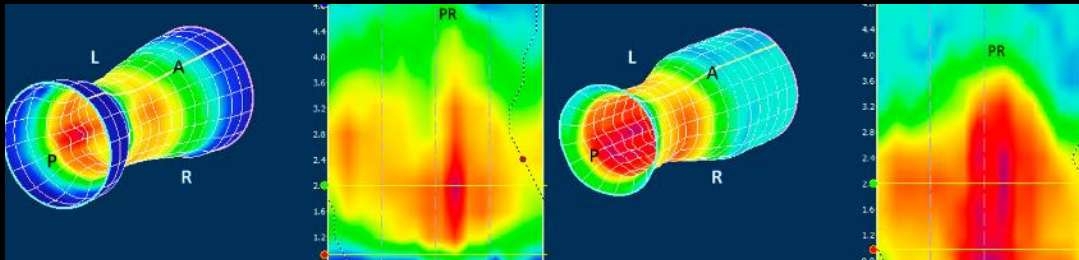
Impaired intestinal motility

Pelvic floor dyssynergia

• Anorectal manometry



• Biofeedback therapy





Rx

PATIENTE NAME _____

ADDRESS _____

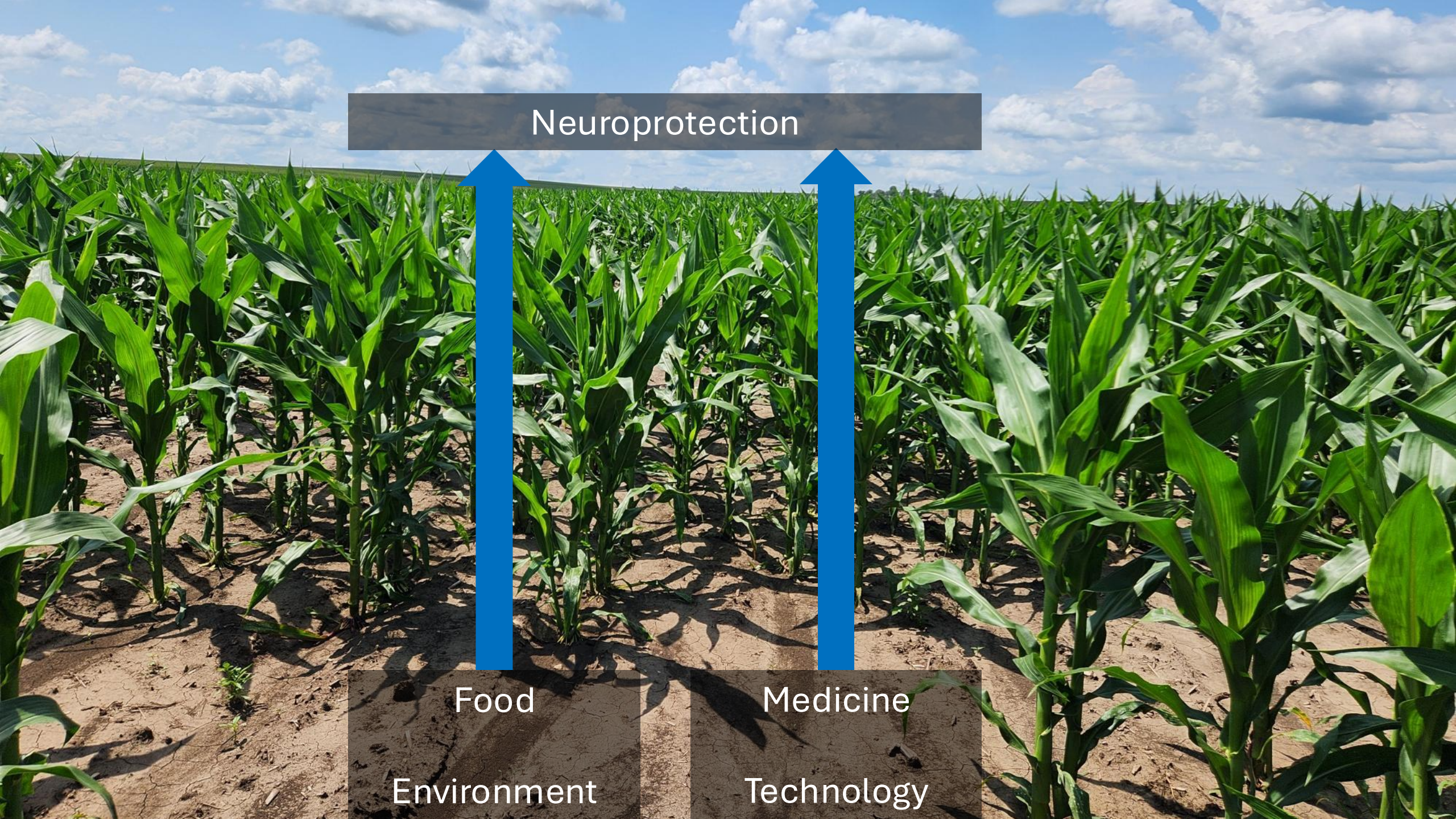
Description: _____

fruit _____

Vegetable _____

SIGNATURE _____

DATE _____



Neuroprotection

Food

Environment

Medicine

Technology

Thank you!

DHC Group:

Harichandana Punukula, PharmD, MS
Jane Baker, ARNP
Ravi Mallavarapu, MD

Motility Center: Stephanie Nerison, RN, BSN, CGRN;
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Lijuan Zhang, MD
Zlata Bogin, MS

Neurology: Christopher Goetz, MD;
Deborah Hall, MD, PhD



Purdue Lab:

Bruce Hamaker, PhD
Thaisa Cantu-Jungles, PhD

To learn more about our Parkinson's trials or participate:

Email: hpunukula@cvmssp.com

To schedule a Parkinson's Digestive Health consultation:

Call: (319) 234-5990 (Dr. Manfready clinic)

Waterloo, IA location

Questions?