

#### **Nutrition and Parkinson's Disease**

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#### **DISCLOSURES**

- Owner of Acacia Health Ltd, Victoria BC
- No shares in fruits and vegetables to disclose





#### **LEARNING GOALS**

- Deeper understanding of nutrition and its relation to PD
- Understanding of Neurological Reserve and how nutrition affects it
- Working knowledge of the microbiome and how to feed it well
- How to determine if protein is causing problems with Levodopa, and what to do about it
- Changed nutrient needs in PwP
- How the Positive Deviants behave (food/exercise)





## WHAT ARE WE TALKING ABOUT?

- What are nutrients?
  - Definition of nutrient:
    - A substance that provides nourishment essential for growth and the maintenance of life. Oxford English Dictionary
      - "fish is a source of many important nutrients, including protein, vitamins, and minerals"
- What nutrients do not enter the body through the gut?
- What nourishes us that is something we DO vs something we RECEIVE?





- Nutrients form the building blocks of your entire body, brain included
- Nutrients/diet affects the 100 trillion organisms that make up the gut microbiome – which has far reaching effects on the entire body, including the brain.
  - Even non-dietary 'nutrients' effect gut microbiome population and diversity
    - Sun exposure
    - Exercise
    - Pet dander (is this a nutrient ②)





- What/when/how you eat affects medication kinetics
- Gut health is an important aspect of PD PwP often report gut symptoms as problematic, diminishing quality of life – constipation, GERD, SIBO, Gastroparesis
  - What you eat affects gut health





- Evidence that nutrition plays a role in development of PD
  - Animal studies indicate potential neurotoxic effect of dairy products (increased oxidative stress, contaminants like rotenone, paraquat toxic to SN neurons)
  - 2019 review by Boulos, C. et al looked at 7 cohort studies showing dairy intake increasing risk of developing PD





- Findings are confusing
  - Low fat dairy appears to confer higher risk
  - Dairy intake seems worse for men (although one cohort did show a correlation with dairy intake and women with PD)
  - In Greece milk was a problem but yogurt and cheese were not associated with increased risk.





- Recent meta analysis PD risk increases by 17% for every 200g/day increase in milk intake, 13% for every 10g/day increase in cheese (Jiang W et al 2014).
- Correlation between faster progression and higher intake of milk, yogurt, and ice cream in the CAM Care PD dataset (Mischley LK 2019 – unpublished)





- Also associations with prevention or delayed onset of symptoms
  - Coffee (28% risk reduction developing PD) 3 cups per day maximum benefit (Qi, H. 2014)
  - Tea (3+ cups per day associated with 7.7 year delay in motor symptoms) Green or Black (Kandinov, B. 2009)
  - Exercise
  - PUFAs
  - Smoking (not recommended)
  - Vitamin D levels
  - Increased dietary antioxidants
  - Mediterranean diet





#### **NEUROLOGICAL RESERVE**

- The sum total of Brain reserve (BR) and Cognitive reserve (CR)
- BR = The total volume of our brains
  - Dictated by many factors (genes, nutrition, injury, smoking, toxin, drug abuse, sleep, inflammation, exercise, circulation)
- CR = The number of and complexity of neuronal networks based upon how we use our brain.
  - Higher in multilingual people, post-secondary education, mentally demanding careers, meditators





### **NEUROLOGICAL RESERVE AND YOU**

- Folks with higher neurological reserve are more resilient to all neurdegenerative diseases (PD, AD) and have lower symptomatology with MS
- My goal is for you to see your Neurological Reserve like a savings account – the more you have in it, the more you can afford to lose and still live well.
- Much of what we (NDs) and other health professionals do to help folks with PD comes down to techniques that retain or build neurological reserve.





#### **NEUROLOGICAL RESERVE AND DIET**

- Nutrition plays a key role in neurological reserve
- Provides fuel for brain to build and re-build and repair
- Provides antioxidants and anti inflammatory factors to help the brain protect itself from damage through inflammation, toxins etc.
- Provides precursors for NT and co-factors for enzymatic conversions.
- Encourages (or discourages) the production of promyelination (nerve insulating) factors by augmenting microbiome activity.
- Provides substances that improve neuronal connections (for example: Vitamin D)





### NEUROLOGICAL RESERVE AND NUTRIENTS

- What nutrients support neurological reserve (some examples)?
  - Omega 3 Fatty acids (Wild Fish, Seeds, Nuts)
  - Antioxidants
    - Blueberries
    - Leafy greens
    - Colourful fruits and veggies
    - Spices and herbs (turmeric, ginger, parsley...)
  - Choline (good sources)
    - Fish, egg yolks, Collard greens, Swiss chard, Brussels sprouts, cauliflower, Asparagus, Broccoli





#### **NUTRITION AND GUT MICROBIOME**

- Gut Microbiome
  - We have over 100 Trillion passengers in/on our body
  - Bacteria, fungi, protozoa, viruses
  - The bacteria in our large intestine form the majority of our microbiome.
  - They help us digest our food, regulate our immune system, protect us against pathogenic bacteria/fungi, produce vitamins including B12, thiamine, riboflavin, Vitamin K.





## NUTRITION AND THE GUT MICROBIOME

- Some interesting facts about the gut microbiome
  - More cells in our guts than our entire body
  - Diversity seems to matter less diverse populations associated with obesity, Type 1 DM, other chronic illnesses
  - Dust from the homes of dogs appear to reduce immune response to allergens in general by adjusting the microbiome.
  - Eating junk food regularly for as little as two weeks creates massive changes in the microbiome towards organisms that encourage inflammation.





### PWP HAVE ALTERED GUT MICROBIOMES

- Studies so far have poor reproducibility only a few signals that seem relatively consistent so far.
- Most studies looked at medicated patients, and some medications appear to affect the microbiome themselves (COMT inhibitors, Anticholinergics) – the consequences of these alterations are currently unknown and could be positive or negative.
- PD patients appear to have (vs. Controls)
  - Higher Lactobacilli, Bifidobacteria, Akkermansia
  - Lower Prevotella, Lachnosperaceae



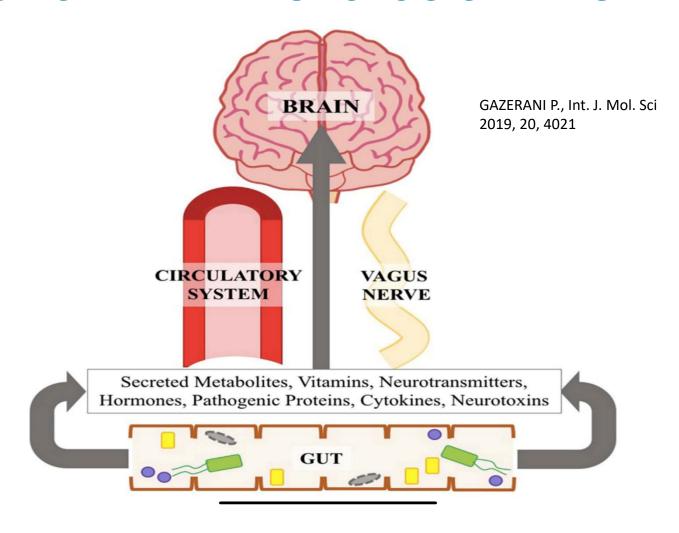


## **GUT MICROBIOME CHANGES IN PWP**

- Lachnospiraceae is a major producer of short chain fatty acids (SCFA) in the colon
- SCFA depletion has been implicated in the pathogenesis of PD in several studies.
- SCFA are produced by healthy gut bacteria breaking down fiber (think veggies, fruits and whole grains) in your colon.
- SCFA are the main source of energy for the cells lining the colon.
- SCFA local and distant effects on gut integrity, inflammation (body and brain), immune system homeostasis











- Microbiota in the gut produce large amounts of neurotransmitters, vitamins, biologically active metabolites (SCFA), hormones, cytokines, pathogenic proteins, and neurotoxins
- These have far reaching and impressive effects on the immune system, levels of neurotransmitters in the brain (not necessarily directly), neuroinflammation, blood brain barrier integrity, and more.





- Example mouse model for epilepsy ketogenic diet found to only be effective for reducing seizures if gut microbiota present.
  - Two strains that are high in mice fed ketogenic diet are known to increase hippocampal GABA and glutamate by decreasing gut Gamma glutamylation.





 In MS research – bacterial metabolites have been found to gain access to the brain and control microglial cell activation (cells that play a large role in inflammation in the brain) – and this can be augmented by diet.





## HOW TO IMPROVE YOUR MICROBIOME

- We can cultivate a healthier microbiome by what we eat and by exercising, getting sun exposure, and ensuring adequate sleep.
- Limiting sugar, fried foods, highly processed foods, and artificial sweeteners, reduces the harmful effects these substances have on our microbiome.
- Increasing 'prebiotic fibers' provides fuel for healthy gut microbes – think asparagus, garlic, onions, whole grains





## HOW TO IMPROVE YOUR MICROBIOME

- Eat fermented foods especially from vegetables and organic soy such as:
  - Saurkraut
  - Lactofermented pickles (in fridge at grocery store)
  - Any lactofermented vegetable
  - Kim Chi
  - Tempeh
  - Miso soup





## PROGRESSION – WHO DOES IT SLOWLY?

#### Positive Deviance

- In any community, there are certain individuals or groups of individuals who have adopted behaviours and strategies that allow them to thrive despite the collective challenges of that community.
- Knowing what the Positive Deviants in the PD community do, do not do, do more of, or less of, can help guide the greater PD community to adopt their strategies.
- Although correlation is not causation, when a trend towards certain low risk behaviours is associated with better outcomes, what is the harm in adopting those behaviours?





#### PRO PD SCALE

- Developed by Dr Laurie Mischley ND PhD
  - Track folks over time motor and non-motor symptoms
  - Monitor progression
  - What activities, behaviours, dietary differences etc...
     are common among the SLOW PROGRESSORS
  - Presented 2019 data set at PD School 2019





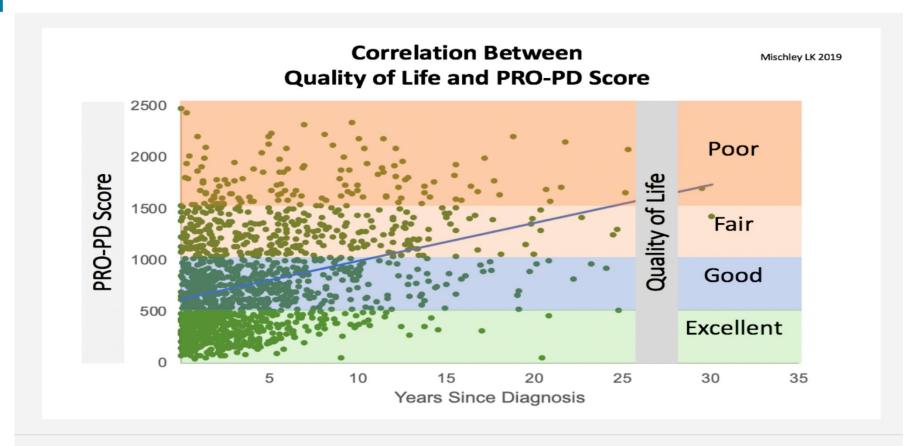
### PRO PD SCALE – WWW.PROPD.ORG

Patient-Reported Outcomes in PD (PRO-PD)		
Please rate the severity of your symptoms over the past 7 days, on average. The more severe and debilitating the symptom, slide right. If you're not having that symptom, slide to the left.		
Slowness * must provide value	Move with ease Severe slowness  Tap the slider above to set a response	reset
Constipation (incomplete bowel emptying) * must provide value	Healthy, daily Require Severe bowel movements medication constipation  Tap the slider above to set a response	reset
Walking * must provide value	I move freely, unable to move with ease  Tap the slider above to set a response	reset
Freezing * must provide value	None Severe, debilitating  Tap the slider above to set a response	reset
Falling * must provide value	Never Occasionally Daily  Tap the slider above to set a response	reset





### PRO PD SCALE



Please contact lmischley@bastyr.edu for questions about using this scale.

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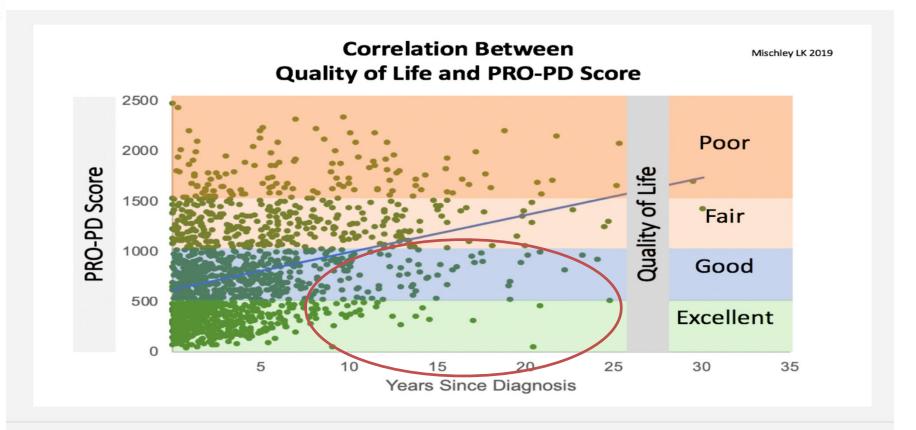
# PRO PD – IMPORTANT KNOWLEDGE

- Average progression is 38 points per year
- Each of 33 motor, mood, and non-motor symptoms are rated on a scale of 0-100
- 0-500 = excellent condition (PD no to very minor impact)
- 500-1000 = good condition (PD creating minimal impacts)
- 1000-1500 = fair condition (PD is impacting life)
- Over 1500 = poor condition (PD with severe impacts)





## PRO PD SCALE – POSITIVE DEVIANTS



Please contact Imischley@bastyr.edu for questions about using this scale.

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#### DIET AND PRO PD

- Intake of specific foods is measured on a 10-point scale
- <1/month, 1/month, 2-3×/month, 1/week, 2–4×/week, 5-6×/week, 1/day, 2–4×/day, 5-6×/day.</li>
- Dose dependent response found for both foods associated with slower progression and those associated with faster progression.





### POSITIVE DEVIANTS EAT (2019 DATA)

- Fresh vegetables
  - 1/wk correlates with 49 point reduction PROPD score
  - 5-6x/d 490 point reduction
- Fresh fruit and veggies
  - 46 400 pt reductions
- Nuts and seeds
  - 40 400 pt reductions
- Non-fried fish
  - 34 350 pt reductions





## POSITIVE DEVIANTS EAT (2019 DATA)

- Wine
- Olive oil
- Fresh herbs
- Coconut oil
- Green tea





### POSITIVE DEVIANTS LIMIT (2019 DATA)

- Dose dependent responses here too
  - Canned fruit
  - Fried food
  - Diet soda (diet soft drinks)
  - Canned vegetables
  - Soda (soft drinks)
  - Beef
  - Ice cream
  - Frozen vegetables
  - Yogurt and Milk





### POSITIVE DEVIANTS AND EXERCISE (2019 DATA)

- Frequency matters
- No benefit seen in folks who exercise 2 or less times per week (30+ min sessions)
- Dose dependent response as frequency goes up
  - 7 days a week 351 pt reduction
  - 6 days a week 343 pt reduction
  - 5 days a week 294 pt reduction
  - 4 days a week 270 pt reduction
  - 3 days a week 201 pt reduction





### POSITIVE DEVIANTS AND EXERCISE (2019 DATA)

- Type of exercise makes a difference:
  - Of the surveyed exercise types
    - Running and Park cycling program showed greatest reductions in the PROPD scale (237 and 228 respectively)
    - Dance and Walking had reductions but less so 64 and 74 respectively
- What do we make of this?
  - Intensity matters?
  - What else?





## POSITIVE DEVIANTS AND BEHAVIOURS (2017)

 When asked about behaviours over the past 6 months(Mischley L, 2017)

PwP who answered "True"	Average change in PROPD score
I routinely prepare meals for others.	- 112.8
I cook most of my meals.	- 135.4
I buy food from local farmers (co-op, farmer's markets).	- 97.2
I try to eat organically grown foods when possible.	- 74.9
I drink from a plastic bottles	+ 11.9





## SOCIAL EXPOSURE – BRAIN NUTRIENT?

- People who answered yes to "I am lonely" had a 354 increase in PRO PD scale.
- Social exposure is recognized as an important factor in preventing/slowing dementia.
- Social exposure causes an increase of BDNF (Brain Derived Neurotrophic Factor) in the brain – "Miracle Grow for the Brain" – required for neuronal development, survival and plasticity.
- PwP who feel they "Have lots of friends" had an average 177 decrease in PROPD





### WHAT ABOUT DIETARY EFFECTS ON MEDS?

- Gastric acid secretion in the stomach provides an acidic environment for foods to be broken down.
- Levodopa has high solubility in acid environments but low in water/neutral pH.
- Approximately 40% of PwP have low stomach acid production – this impairs Levodopa solubilization and subsequent absorption.





### **INCREASING STOMACH ACID**

- Diet/Supplementation can help:
  - Take med with
    - lemon juice (30 ml R. Shinkeigaku study 1994)
    - Vitamin C (as Ascorbic Acid) 500 mg capsule.
    - Apple cider vinegar
    - Betaine HcL capsule
  - Dietary practices that increase HCL secretion/Stomach acidity
    - Eat mindfully, minimize stress during meals





### PROTEIN AND MEDICATION

- Levodopa competes with amino acids from dietary protein for access to the blood stream (from the gut) AND access to the brain (from the bloodstream)
- Signs that this could be a problem for you:
  - Long time for med to kick in
  - More time in off state than typical especially related to high protein meal
  - "Dose Failures" where the med does not seem to work at all – especially related to protein in diet





## PROTEIN AND MEDICATION – WHAT TO DO

- Determine if this is the problem
  - Keep a diet journal with date and time of what you ate or drank, and also track medication times AND results.
    - Bring this to your dietician or doctor
  - OR do a 4-day low protein diet and see if you get better symptom control





#### HIGH PROTEIN FOODS

- Flesh foods (meat, chicken, turkey, pork, fish)
- Eggs
- Most dairy
- Nuts and nut butters, Seeds and seed butters
- Soy beans and soy products (tofu, edamame, soy milk)
- Protein powders
- Ensure or Boost
- Beans and peas, lentils
- Desserts made with eggs or cheese/dairy products





## PROTEIN AND MEDICATION – WHAT TO DO

- If you determine it is an issue (pick one and try it):
  - Take levocarb on an empty stomach 30 min before or 1 hour after meals. Even better – take it with an acid as well.
  - Limit protein intake during the day to prevent the interaction during your most active times.
    - You must still get essential protein with this method
    - Eat a high protein dinner meal
  - Plant protein may interfere less than animal flesh protein and eggs – reserve the latter fo nighttime intake





# CHANGED NUTRIENT NEEDS IN PWP

- Your body has altered nutritional requirements due to PD
- AND recommendation of high frequency exercise for treatment and progression slowing ADDS to the nutrient demands for your system





#### **HYDRATION**

- Increased nighttime urination, low blood pressure, medication effects, altered thirst signaling – all contribute to a greater need for adequate hydration.
- I encourage folks with PD to aim to get 2 liters spaced out through the day (instead of large servings) to help maintain blood pressure, reduce cramping from dehydration.
- Additional water needed before/during/after exercise
- Electrolytes help reduce muscle cramping and improved blood pressure control – have with water around exercise, and during the day if OH or muscle cramping a concern.





#### VITAMIN D

- Folks with PD tend to have lower Vitamin D levels than age matched controls (Sleeman, I. et al, 2017)
- Lower Vitamin D at diagnosis is associated with greater severity in motor symptoms 36 months later (Sleeman, I. et al 2017)
- Supplementation recommended (liquid form or gel caps)
  - 2000-4000 IU D3/day to maintain levels
  - 6000 IU D3/day to attain healthy levels
- I like to test, treat, and re-test because I find it can be challenging, in PwP, to get to healthy levels.





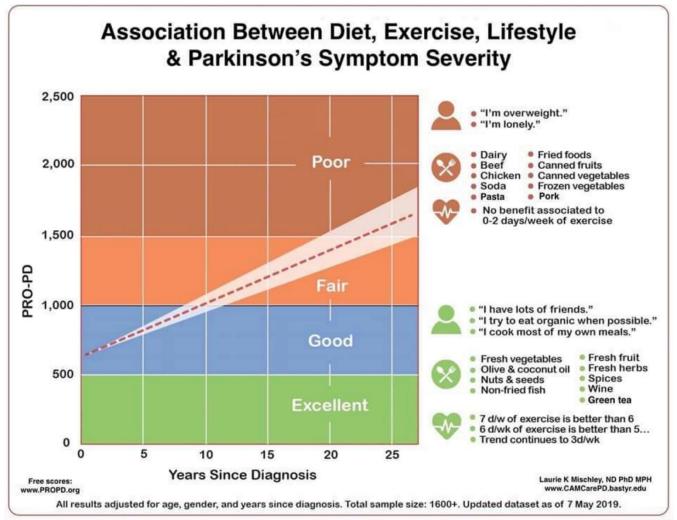
#### **FIBER**

- Fiber is highly therapeutic for PwP, and arguably the needs for fiber increase in folks with the condition.
- Fiber improves levodopa bioavailability
- Fiber provides food for healthy microbiota cultivation, and subsequent gut wall health and integrity
- How to get more fiber
  - Eat 4-6 cups of vegetables per day (with half being leafy greens), 1-2 cups berries and fresh fruit per day
  - Eat whole grains whenever having any grain.
  - Supplement with psyllium husk powder (this can improved levodopa absorption and reduce constipation)





#### TAKE HOME MESSAGES







### THANK YOU

- Contact information:
  - pam@acaciahealth.ca



