#### GI MICROBIOME & HEALTH: A REVIEW

Nancy C. Kois, MD, FCAP Contemporary Pathology Service



# DISCLOSURES

• There are no disclosures

# OBJECTIVES

- Definitions: GI microbiota, GI microbiome, probiotic, prebiotic
- Current ideas: GI microbiome and health
- Current ideas re: GI microbiome maintenance
- Current microbiome projects and goals

# CURRENT ADVANCES

- NGS and gene editing (e.g., CRISPR)
- Nanotechnology
- Quantum computing
- Artificial Intelligence
- Online consumer
- Smart everything



#### PUBLISHED IN HEALTHLINE JAN 2019

- "Your gut microbiome is made up of trillions of bacteria and other microorganisms-both friendly and unfriendly.
- Maintaining the right balance of friendly and unfriendly bacteria in your gut is touted to improve digestion, reduce inflammation, decrease anxiety, and improve brain function and mood.
- A healthy balance of gut bacteria is said to boost metabolism, eliminate cravings, and help you shed unwanted weight."

A. Petri, MS, RD

## GI MICROBIOME

- · GI microbiota = all the bacteria in our GI tract
- GI microbiome = all the bacterial genes in our GI tract



## GI MICROBIOME-THINK ABOUT THIS

- · Human body has more bacterial cells than human cells
- Human body has more bacterial genes than human genes
- · Bacterial metabolites influence human biochemistry
- Bacteria multiply rapidly, mutate, and exchange genes
- Our body bacterial genome is something we can change



# CURRENT ESTIMATES

HUMANVS BACTERIAL CELLS
30 trillion human cells

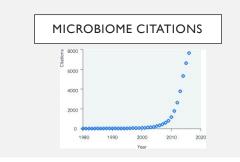
- 20,000 hur
- 39 trillion bacterial cells

43% vs 57% for cells



2-20 million bacterial genes

1% vs 99% for genes





### COLONIC BACTERIA

- Bacteroides fragilis

- Bacteroides fragilis
   Bacteroides relaninogenicus
   Bacteroides crails
   Enterococcus faecalis
   Enterococcus faecalis
   Enterobacter species
   Klebsiella species
   Bildobacterium bildum
   Staphylococcus aureus
- Lactobacillus species Cloatridium perfringes Proteus mirabilus Cloatridium teani Cloatridium septicum Pesudononas aeruginosa Salmonella enterica Fecalitoacterium prusinkti Peptoarreptococcus species Peptococcus species

# SCIENTIFIC PUBLICATIONS



# WHAT THEY ARE SAYING

#### GI bacteria:

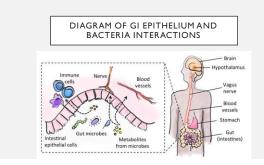
- regulate the health of GI tract epithelial cells
  there are good GI bacteria and not good GI bacteria
  number and type are regulated by our diet
- influence body metabolism, body weight, immune system, behavior diseases, response to medications
- influence food choices
- GI bacterial diversity is favorable
  Diet influences GI bacterial diversity
- · Diet high in fiber promotes GI bacterial diversity · Role for fecal transplant in health care



# MORE DEFINITIONS

- Probiotic=bacteria that are beneficial to host GI
- Prebiotic=foods we eat that feed good bacteria





VIDEO

## GI MICROBIOME BELIEVED TO PLAY A ROLE IN:

#### • obesity

- metabolic syndrome
- cancer
- IBD
- IBS
- arthritis • asthma
- diabetes
- I USED TO THINK CORRELATION IMPLIED CAUSATION. I THEN I TOOK A STATISTICS CAUSE. NOW I DONT. WELL, MARKE 8 8 R 9

CNS disease

heart disease

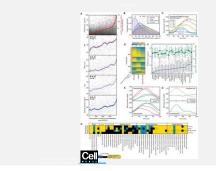
liver disease

GI ulcers

#### ZEEVI ET AL 2015

# "PERSONALIZED NUTRITION BY PREDICTION OF GLYCEMIC RESPONSES"

- 800 people, continuous glucose monitor, measured their microbiome, measured their response to the same meals
- People eating identical meals present high variability in post-meal blood glucose response
   The response was related to microbiome
- $^\circ~$  Were able to design personalized diet to lower glycemic response
- · Response to diet is predicted by microbiome
- \* Change in diet can change microbiome and glycemic response



#### MICROBIOME AND WEIGHT

- Gl microbiome analysis can predict with 90% certainty whether or not a person is overweight
   Human genome analysis can predict with 57% certainty whether or not a person is overweight.

- Twin studies
- · overweight twin had decreased bacterial diversity.
- Non overweight twin had more: Christensenella and Akkermansia
- Germ free mice experiments



#### MICROBIOME AND IMMUNE SYSTEM

- GI microbiome believed to influence the development of the innate and adaptive host immune systems
- Scher, J, et al "Expansion of intestinal Prevotella copri correlates with enhanced susceptibility to arthritis"
- Believed to be a role for intestinal bacteria in supporting the systemic immune response required for joint inflammation
- \* Also believed association with increased prevalence of asthma



#### MICROBIOME AND INTESTINAL DISEASE

- 2019 Nature Communications
- Haberman,Y et al:"Ulcerative colitis mucosal transcriptomes reveal mitochondriopathy and personalized mechanisms underlying disease severity and treatment response"
- Identified positive associations between genes and pathways associated with UC severity and response to treatment and disease-linked microbial taxa.



#### MICROBIOME AND RESPONSE TO TUMOR TREATMENT

- The ability of gut microbiota to alter response to chemotherapy has been shown in animal models
- "Antibiotic treatment prior to immune checkpoint inhibitor therapy as a tumor-agnostic predictive correlate of response in routine clinical practice."
- · Pinato,J et al, presented March 1, 2019
- · Medications, esp antibiotics affect gut microbiome
- · ?chemicals in food, ?artificial sweetners

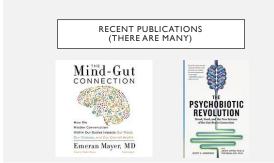
### MICROBIOME AND CNS DISEASE

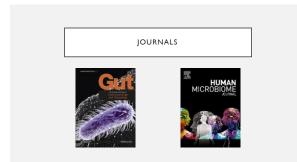
Nutritional psychiatry:

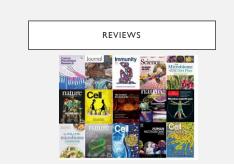
- Jostin Honar psychiat 37
   Josta, Fe al. YA randomized controlled trial of dietary improvement for adults
   with major depression" 2017
   Parletta, N et al. 'A Mediterranean-style dietary intervention
   supplemented with fish oil improves diet quality and mental health
   in people with depression" 2017

- \* Role in neurodegenerative diseases: MS, Parkinson's









### PROJECTS AND ORGANIZATIONS

- Human Microbiome Project
- American Gut Project
- Center for Microbiome Innovation-Al for Healthy Living
- American-British Gut Project
- Infectious Disease and Microbiome Project
   Global FoodOmics Project
- Mayo Clinic Microbiome Project



# YOUR AMERICAN GUT SAMPLE

#### MICHAEL POLLAN What's in your American Gut sample?

~		Fittidadel Baranoldoni Possidadoria Admitisatoria Venucomorbia Terrehoda	Your most abundant microbes:		Your most enriched microbes:			
. *	- Page		Таколопу	Sample	Taxonorty	Sample	Papulation	Fold
			Genus Phavotella	24.9%	Genus Closhidury	2.5%	0.3%	78
			Family Auminococcaceae	13.4%	Genus Finegoldia	0.7%	0.0%	17x
· _		chacteria barteria	Family Lachnospiraceoe Genus Besternoles	10.1%	Genus Arevotella Genus Collosella	24.9%	2.9%	9×
711		- Citer	This sample included the follow sam taxa: Genus Vanbaculum, Genus Aelaserie, Genus Campylobacter: Order ML615J-28					
How do	your gut microl	bes co	impare to other	rs?	• Yes + Ohers	• Numing		. *
	and the second	head						9
	Different body sites		Different ages and pop			The Ameri	an Gat popy	witten

### PROJECT GOALS AND ACHIEVEMENTS

- · Collect and categorize microbiome data

- Data is open and free for research
  ApplyAl to interpretation of microbiome data
  Review microbiome taxa to predict disease

#### ROLE OF FECAL TRANSPLANT

- · History of use with C. difficile
- Current modes of transplant



#### RESEARCH EFFORTS

- Dr. Tim Spector
- King's College London
- Author:
- Identically Different:Why You Can Change Your Genes 2012
- The Diet Myth:The Real Science Behind What We Eat 2015
- Dr. Rob Knight • UCSD, La Jolla, CA
- Cofounder:American Gut Project
- Cofounder: Earth Microbiome Project

# CONCLUSIONS FROM DR. SPECTOR

#### What we Know:

-Probiotics have beneficial effects on human health -Gut microbes influence human energy metabolism -Diet and medication have a strong influence on gut Diet and medication have a strong influence on gut microbiota
 Microbiota composition influences response to chemo and immunotherapy
 Microbiome composition defines glucose response to foods and can be used to personalize diet
 Dietary fiber intake influences gut microbiota and is related to better health

# What we don't know:

-Are natural probiotics in food better than probiotic supplements? Should we take them preventively -Can microbes influence food choices and

-Can microbes influence food choices and appetite? -Do low dose antibiotics in food affect human health? -The effect of pesticides in food on the gut microbiome? Is organic food better? -Should all new drugs and food chemicals be tested on the gut microbiota?

#### DIETARY RECOMMENDATIONS FOR HEALTHY GI MICROBIOTA

- \* Plant-based complex carbohydrates (variety important to promote diversity)
- Plant-derived fats
- Grains
- Fish and chicken better
- Polyphenols (olive oil, red wine)
- Plant products regarded as anti-inflammatory (turmeric, curcumin, ginger)
- Naturally fermented foods (contain microorganisms) Avoid red meat and animal fat
- Avoid refined sugar and processed food



# WHY ARE THESE FOODS REGARDED AS GOOD?

These foods are believed to:

- $^{\circ}\,$  increase bacterial diversity (bacterial food source)
- increase number of health-promoting bacteria
- Health-promoting bacteria:
- are believed to reduce intestinal permeability ("leaky gut")
- are believed to be good for brain

### IN SUMMARY

 "Research suggests that the relationship between gut flora and humans is not merely commensal (a non-harmful coexistence), but rather is a mutualistic, symbiotic relationship."



# GI MICROBIOME!

- GI microbiome is important in health and disease
- Thank you for your attention
- nck@usa.net

