



Nutrition for Gut Health & More

Kerry K. McMillen, MS, RD, CSO, FAND
Director, Oncology Supportive Care and Screening

Describe the gut microbiome and factors in development.

Discuss the role of the gut microbiome and impact on oncologic therapies.

Review evidence-based nutrition recommendations for maintaining a healthy gut microbiome.

Recipes for Success

Q + A

Gut Microbiome + Diet

“At a minimum, the diversity of the gut microbiome may be a future biomarker of long-term consumption of a “healthy” versus “unhealthy” diet, which may be linked to potential for disease development.” *Cresci*

Gut microbiome: What is it?

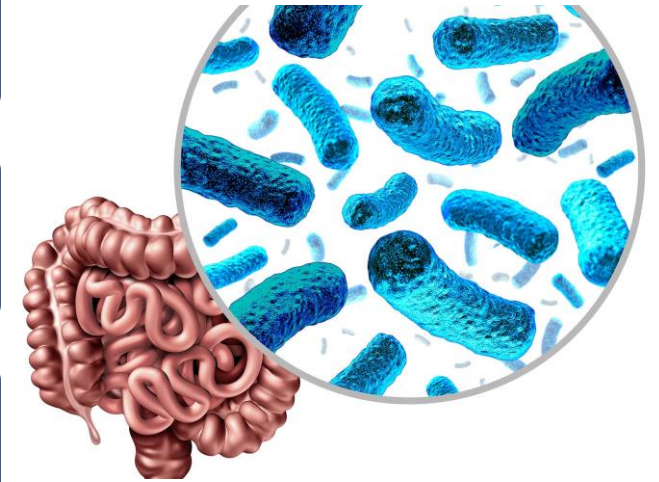
Gut microbiome is comprised of microorganisms such as bacteria, archaea, fungi, and viruses that are distributed throughout the entire GI tract.

Humans are made up of more bacterial cells (40 trillion) than human cells (30 trillion).

The GI tract contains over 3 million genes, which is 150 times more genes than in the genome of a human body.

Gut microbiome contains up to 1,000 species + more than 7000 different strains of bacteria.

Can weigh as much as 1-2 kg (similar to the weight of your brain!).



Gut Microbiome Function

Three main functions:

- Structural
- Protective
- Metabolic

Plays an essential role in:

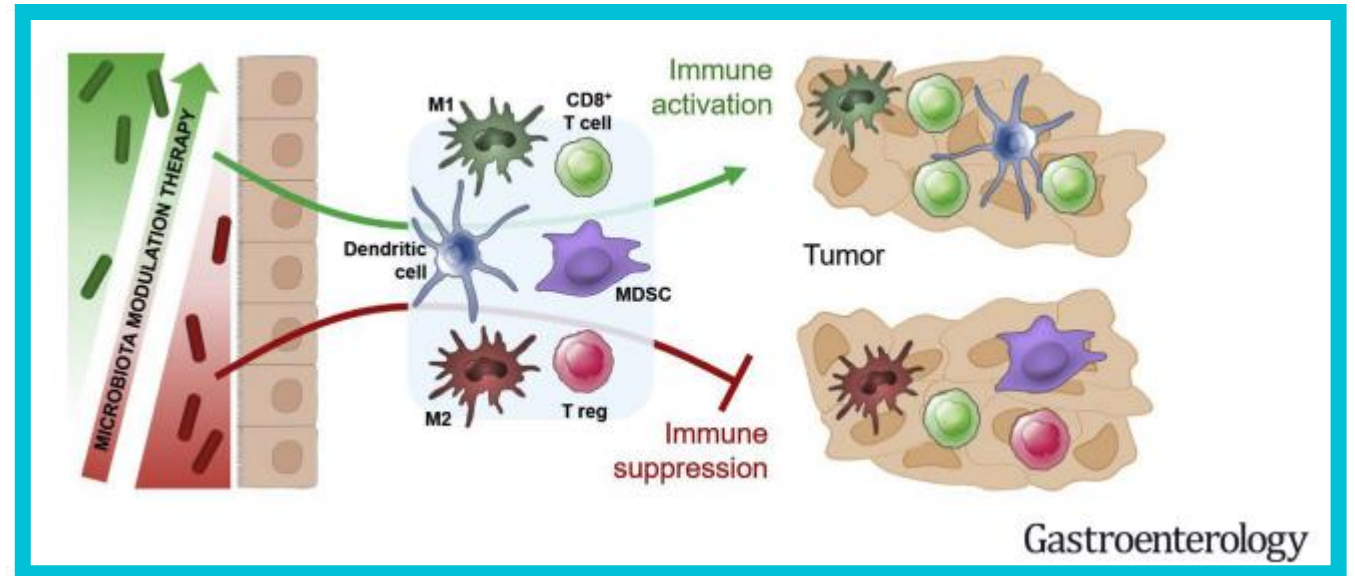
- Nutrient + mineral absorption
- Synthesis of enzymes, vitamins and amino acids
- Production of short-chain fatty acids (SCFAs)

Gut Microbiome + Human Health

Studies show the maintenance of a healthy gut microbiome is inseparable from host health.

Bidirectional relationship between the GI microbiome and innate immune system.

Ongoing investigations are revealing the importance of the gut microorganisms in exerting beneficial effects on human health.



Gut Microbiome Function + Host Health

Fermentation byproducts of gut microbiota (acetate, propionate + butyrate) are important for gut health:

- Provide energy for epithelial cells
- Enhance epithelial barrier integrity
- Provide immunomodulation
- Protect against pathogens

Coordinate action/communication between the gut microbiome and host immune system

- Robust gut microbiota enables the immune system to recognize and attack opportunistic bacteria

Gut Microbiome

Primary Influential Factors



Eubiosis vs Dysbiosis

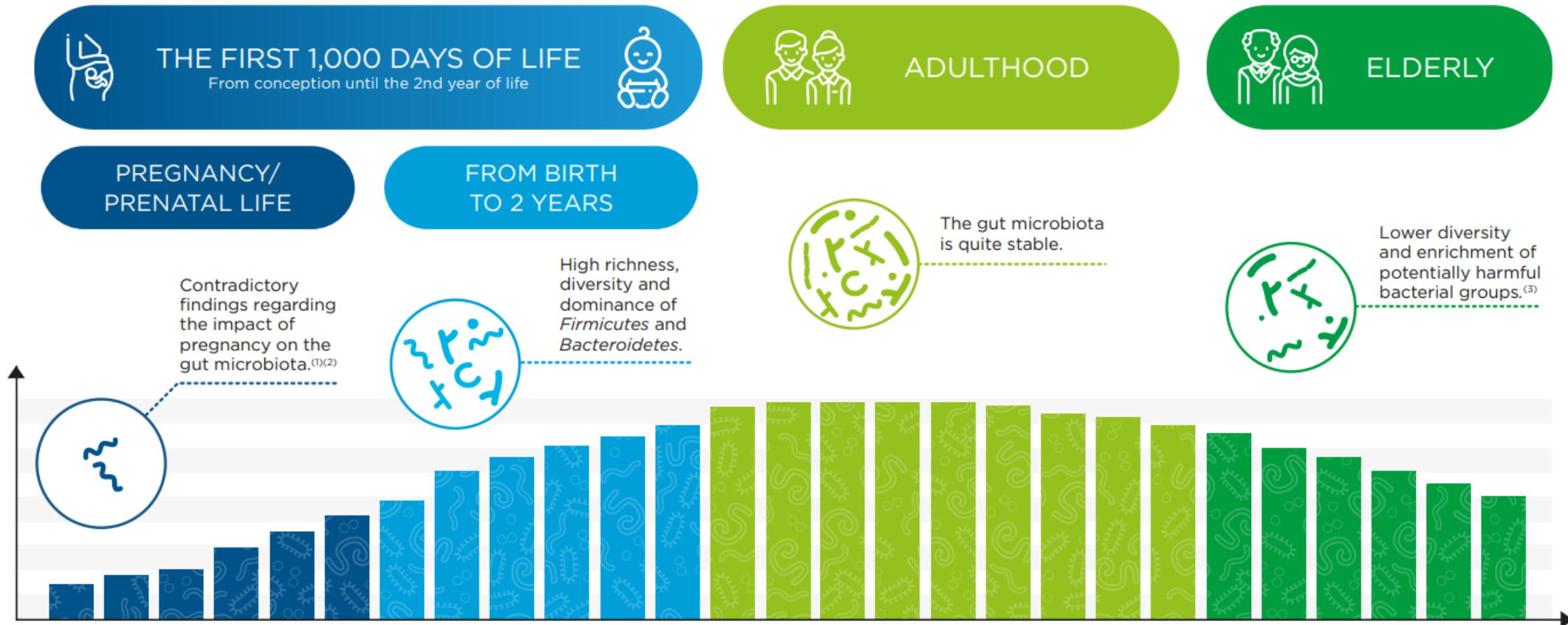
EUBIOSIS = STATE OF HEALTH:

- ✓ Diversity of bacteria in the gut microbiota
- ✓ Balance between pro-inflammatory + anti-inflammatory cytokines
- ✓ An intact/healthy mucosal barrier and mucus layer.

DYSBIOSIS = ALTERATION OF THE MICROBIAL COMMUNITY:

- Causes altered metabolism in the intestine → disturbs functions of the microbiota
- Can result in ↓ diversity + helpful bacteria
- Connected to a wide array of chronic diseases = cardiovascular, metabolic, neurological, autoimmune, GI diseases, obesity and cancer

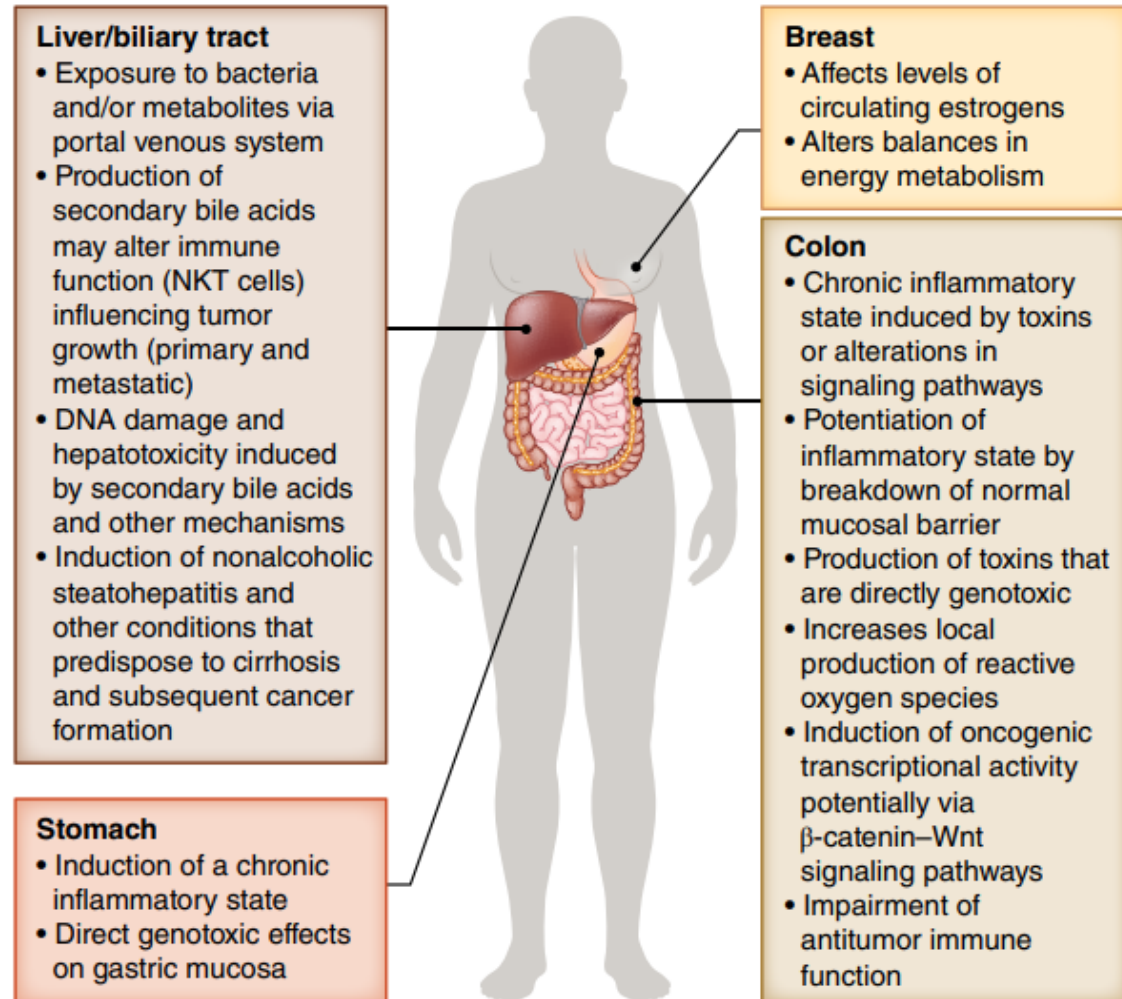
Gut Microbiome over the Human Lifecycle



Role of the gut microbiome and impact on oncologic therapies

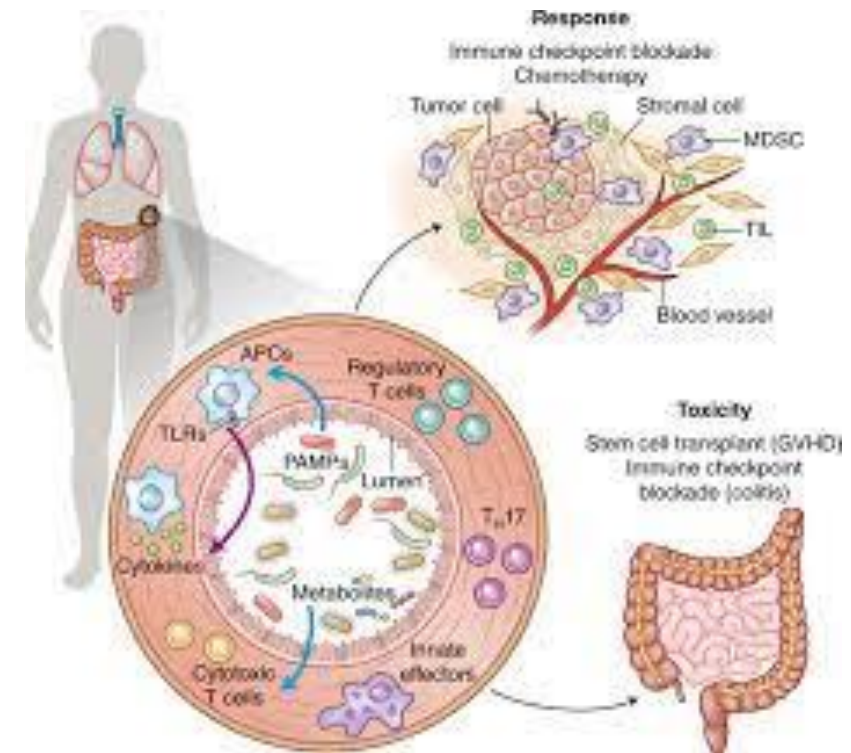
Gut Dysbiosis and Cancer Development

- Gut dysbiosis and cancer development is increasingly being recognized.
- Certain bacteria and viruses have been implicated in cellular dysplasia and carcinogenesis.
- Known oncogenic gut bacteria include *Salmonella* and *Helicobacter* in biliary cancer and *Helicobacter pylori* in gastric cancer.
- In most of these cases, carcinogenesis is believed to be secondary to the creation of a local chronic inflammatory state; however, some bacteria, including *H. pylori*, have direct genotoxic effects.
- ***Additionally, evidence supports the notion that generalized dysbiosis of the gut microbiota may contribute to carcinogenesis.***



Impact of gut microbiome during cancer therapy

- Highly diverse bacterial populations inhabit the GI tract and modulate host inflammation with promotion of immune tolerance
- During oncologic therapy, the GI tract may be damaged and colonizing bacteria are impacted, leading to an impaired intestinal microbiota with reduced diversity
- Gut microbes can impact both the response to various cancer therapies and associated toxicities
- **QUESTION: Does altering the gut microbiome improve patient outcomes?**



Helmk, 2019

Impact of gut microbiome: solid tumors

- **Breast cancer:**
 - Review paper evaluating the role of the gut microbiota dysbiosis in breast cancer patients summarized that women who consumed a vegetarian diet or Mediterranean diet had higher intestinal microbiome diversity with lower estrogen levels (*Ruo, 2021*)
- **Liver cancer:**
 - Patients with liver cancer treated with bacteriotherapy have been able to modify the microbiota composition, thereby reducing overall inflammation and fibrosis (*Moreno-Gonzalez, 2019*)
- **Melanoma:**
 - Higher dietary fiber intake associated with improved progression-free survival in 128 patients with metastatic disease maintained on immune checkpoint blockade treatment (*Spencer, 2021*)
- **NSCLC:**
 - Patients who responded to nivolumab therapy had a higher microbiome diversity with significantly prolonged progression survival when compared to those with low diversity (*Jin, 2019*)
- **Pancreatic cancer:**
 - Diverse gut microbiome associated with improved CD8 T-cell response and prolonged survival in patients with pancreatic ductal adenocarcinoma (*Riquelme, 2019*)

Diet recommendations for maintaining a healthy gut microbiome*.

***Work with a Registered Dietitian and medical provider to determine what diet modification and/or supplements may be indicated for your specific condition.**

***Do not start supplements without approval from your medical provider.**



We can all apply the motto
“BE THE BEST AT GETTING BETTER”
to our diet and lifestyle choices.

Diet and Gut Microbiome

Diet is critical.

Major role of the gut microbiome is food digestion + harvesting key nutrients that the host is not capable of metabolizing on its own.

BOTTOM LINE= Diverse microbiome associated with diets high in fruits, vegetables + fiber compared with a Western diet rich in fat, sugars + animal protein and depleted of fiber.

The American Gut Project

The number of plant types you eat plays a role in the diversity of their gut microbiome.

Participants who ate >30 different plant types/week had gut microbiomes that were more diverse than those who ate 10 or fewer plant types/week.

This is likely why fiber supplements do not have the same metabolic/microbiome effect.

To learn more or participate, visit AmericanGut.org.

BLUE & PURPLE

blackberries
blueberries
black currants
dates
eggplants
grapes

plums
prunes
purple figs
raisins

RED & PINK

beets
cherries
cranberries
pink grapefruit

pomegranates
radicchio
red radishes
red apples
red grapes
red peppers
red potatoes
rhubarbs
strawberries
tomatoes
watermelons



EAT MORE COLOR

The best way to get all of the vitamins, minerals and nutrients you need is to eat a variety of colorful fruits and veggies. Add color to your plate each day with the five main color groups.

GREEN

artichokes
asparagus
avocados
bok choy
broccoli
Brussels sprouts
celery
collard greens
cucumbers
green beans
green cabbage
green grapes
green onions
green peppers
kale

kiwis
leeks
limes
mustard greens
okra
pears
peas
romaine lettuce
snow peas
spinach
sugar snap peas
watercress
zucchini

WHITE

bananas
cauliflower
garlic
Jerusalem artichokes

mushrooms
onions
potatoes
parsnips
shallots

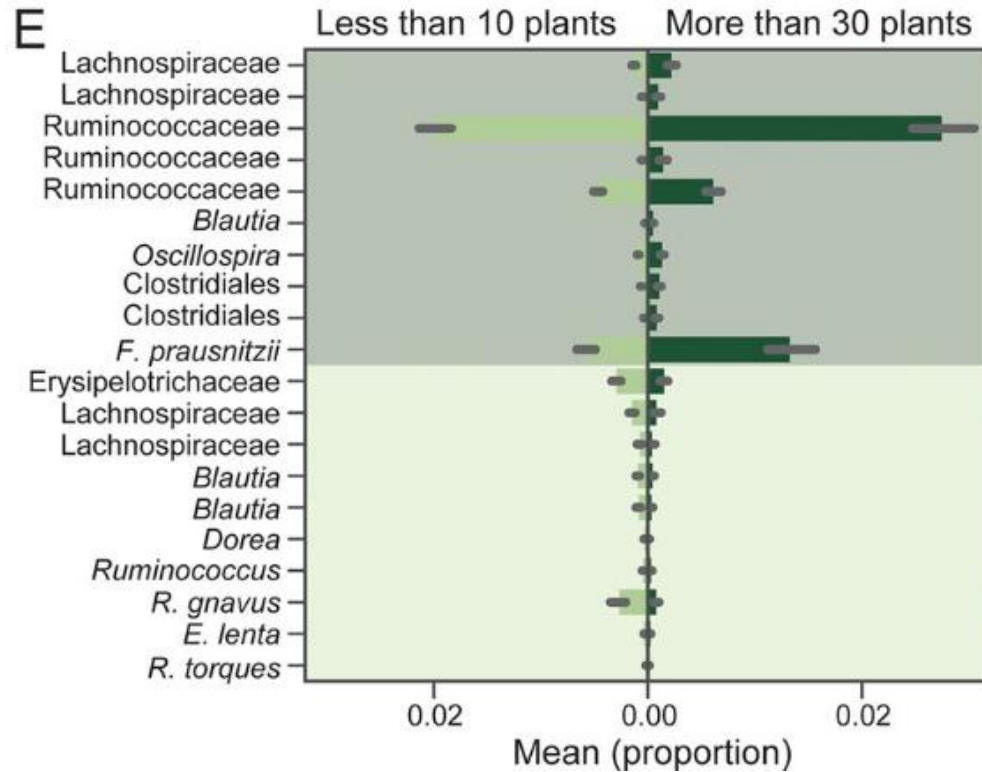
ORANGE & YELLOW

acorn squash
butternut squash
apricots
cantaloupes
carrots
corn
grapefruit
lemons
mangoes
nectarines
oranges

orange peppers
papayas
peaches
pineapples
pumpkins
summer squash
sweet potatoes
tangerines
yams
yellow apples
yellow peppers
yellow squash

Unique Fiber Types Matters for Diversity

N = 11,336



- # of unique plant species consumed is associated with microbial diversity, rather than self-reported categories (i.e. Vegetarian)
- Bacteria differ in CHO binding molecules and enzymes that hydrolyze diverse substrates in the gut, therefore a more diverse diet likely supports a more diverse microbiome
- Plant consumption also associated with a reduction in certain antibiotic resistance genes

Fiber and Phytochemicals

Goal

- 30 grams/day
- ~10% Americans meet this goal

Sources

- Fruit
- Vegetables
- Whole grains
- Nuts/seeds
- Beans

Serving

- ½ cup cooked whole grain
- 1 cup whole grain cereal
- 1 cup raw fruit/vegetable
- ½ cup cooked fruit/vegetable
- ~1.5 oz whole nuts
- 2 Tb nut butter



Nutrition Matters



Nutrition: *Part of Your Cancer Treatment Plan*



From day one, nutrition should be considered alongside treatment for improved outcomes.

After a cancer diagnosis, eating well can help you throughout chemotherapy, radiation therapy and surgery. Aim to follow AICR's Cancer Prevention Recommendations during treatment.*



NUTRITION CAN HELP YOU:

- Heal and recover faster from treatments
- Reduce your risk of infection
- Support your immune system
- Build your strength and energy
- Tolerate side effects from treatments
- Maintain a healthy weight
- Replace or retain nutrients that are stored in your body
- Reduce your risk of cancer recurrence

***Work with a dietitian and your cancer care team to set up your personal nutrition plan.**

Modes of gut microbiome modulation: Diet

Eat a diet rich in plant foods such as whole grains, vegetables, fruits, and legumes

- Endorsed by the American Institute of Cancer Research (AICR) for cancer prevention

Base diet around plant foods which contain fiber (to help diversify the gut microbiome) and other nutrients to reduce cancer risk

- Consumption of plant foods (lower in calories) also help maintain a healthy weight
- 2/3 of plate/bowl should include plant-based foods

Enjoy at least 4 cups (raw and cooked) daily

- Eat a “Rainbow of Colors”
- Don’t forget about beans, lentils, peas, nuts

AICR's Foods that Fight Cancer™

No single food can protect you against cancer by itself.

But research shows that a diet filled with a variety of vegetables, fruits, whole grains, beans and other plant foods helps lower risk for many cancers. In laboratory studies, many individual minerals, vitamins and phytochemicals demonstrate anti-cancer effects. By including more foods that fight cancer into your diet, you will help reduce your risk of developing cancer.



APPLES



ASPARAGUS



BLUEBERRIES



BROCCOLI AND
CRUCIFEROUS
VEGETABLES



BRUSSELS SPROUTS



CARROTS



CAULIFLOWER



CHERRIES



COFFEE



CRANBERRIES



FLAXSEED



GARLIC



GRAPEFRUIT



GRAPES



KALE



ORANGES



**PULSES: DRY BEANS,
PEAS, AND LENTILS
(LEGUMES)**



RASPBERRIES



SOY



SPINACH



SQUASH (WINTER)



STRAWBERRIES



TEA



TOMATOES



WALNUTS



WHOLE GRAINS

Modes of gut microbiome modulation: probiotics, prebiotics, postbiotics

- **Probiotics**

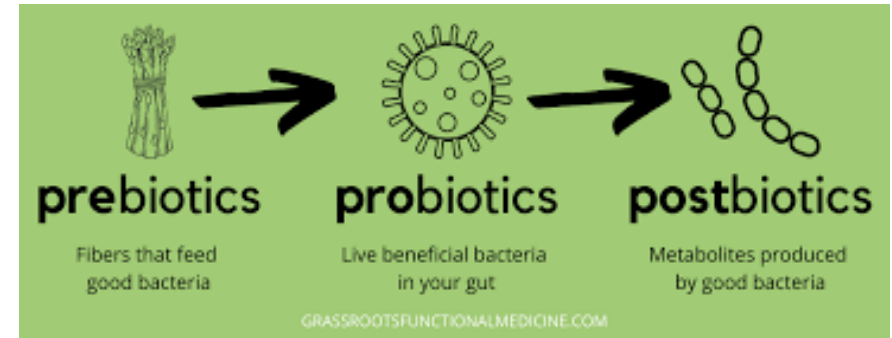
- Contain a defined amount of viable microorganism that upon administration, confer a benefit to the host
- Examples: pasteurized food sources (yogurt, kefir, acidophilus milk)

- **Prebiotics**

- Consist of nondigestible food ingredients (undigested fibers) that support the growth of beneficial bacteria
- Examples: whole grains, bananas, onions, garlic, artichokes

- **Postbiotics**

- “End game” of probiotics and prebiotics – released from gut bacteria that are fed nondigestible food ingredients (undigested fibers) that support the growth of beneficial bacteria
- Examples: Short-chain fatty acids, vitamins, amino acids, lipopolysaccharides (large molecules found on the outer layer of certain bacteria)



Putting Science on Your Plate

Tips to add Fiber and Phytonutrients



Add fruit to breakfast



Make a veggie omelet



Mix up the grains: quinoa, bulgar, etc



Mix seeds in salad, smoothies or oatmeal



Eat nuts as snacks and/or add to baked goods + salads



Include roasted vegetables w/lunch and dinner

Kitchen Strategy

Stay one step ahead:

When prepping a meal, consider future uses for those ingredients:

If cutting 4 carrots to roast with dinner, cut up a 5th one for a snack tomorrow

Roasting cauliflower for dinner? Roast 25% more and save for “leftover salad”

Mix any leftover cooked vegetables with eggs and serve with whole wheat toast for a plant-based breakfast

Cook whole grains (barley, quinoa, brown rice, etc) in large batches to use in a variety of ways

Strategies for Success

Pre-prepare your breakfast, lunch and snacks! (This can be done by someone else...)

Purchase premade salads and/or fruit bowls if too rushed to make your own.

Build a desk/locker “pantry” with self-stable items such as individually wrapped popcorn, nuts, and bars.

Consider a new lunch box or thermos with compartments.

Prep in advance a few days of salads layered in mason jars.

Plan your meals around high fiber, high protein foods for a long-lasting fuel.



Increasing Vegetable Intake: Greens



- Keep a container of greens in your fridge. They go with just about everything:
 - Add to eggs & sandwiches
 - Start each plate with greens
 - Leftover salad
 - Add to smoothies, soups and beans

- Kale fennel salad



- Dressings



Increasing Vegetable Intake: Carrots

Keep carrots in your fridge:

- They last a long time – great “backup” veggie
- Very versatile with many raw and cooked uses
- Inexpensive (<\$1/pound)

Baked root vegetables fries with Mexican crema





Increasing Bean Intake

The most economical protein source:

- Enjoy as a main dish, like chili, stuffed bell peppers, tacos/burritos, vegetarian curries, etc.
- Use bean spreads like humus on veggie sandwiches or as a snack/side with cut vegetables
- Roasted chickpeas are a great crunchy snack alternative to chips and/or a great salad topping

- Kale Caesar salad with chickpea croutons



Summary

- Diet is key for supporting a healthy gut microbiome.
- Encourage a whole-foods, plant-based diet with a focus on diversity.
- Modulation of the gut microbiome may play a role in improving outcomes for oncology patients as well as other chronic diseases.
- We can all embrace “Being the Best at Getting Better”!



Great Resources

- Cook for Your Life: <https://www.cookforyourlife.org/>



- American Institute for Cancer Research: <https://www.aicr.org/cancer-survival/>



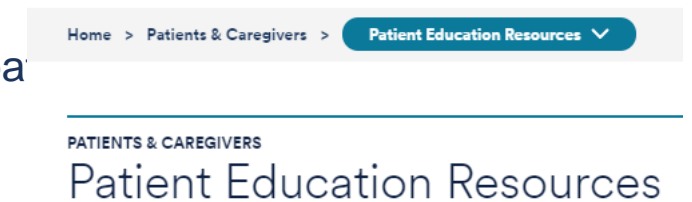
- American Cancer Society: <https://www.cancer.org/>



- Fred Hutch public facing website: <https://www.fredhutch.org/en/education-training/pa>



- Find a Registered Dietitian: <https://www.eatright.org/find-a-nutrition-expert>





Thank you



References:

- Cresci GA, Bawden E. Gut Microbiome: What We Do and Don't Know. *Nutr Clin Pract*. 2015 Dec;30(6):734-46. doi: 10.1177/0884533615609899. Epub 2015 Oct 8. PMID: 26449893; PMCID: PMC4838018.
- Sender R, et al. Revised Estimates for the Number of Human and Bacteria Cells in the Body. *PLoS Biol*. 2016 Aug 19;14(8):e1002533. doi: 10.1371/journal.pbio.1002533. PMID: 27541692; PMCID: PMC4991899.
- Qin J, et al. A human gut microbial gene catalogue established by metagenomic sequencing. *Nature*. 2010 Mar 4;464(7285):59-65. doi: 10.1038/nature08821. PMID: 20203603; PMCID: PMC3779803.
- Integrative HMP (iHMP) Research Network Consortium. The Integrative Human Microbiome Project: dynamic analysis of microbiome-host omics profiles during periods of human health and disease. *Cell Host Microbe*. 2014 Sep 10;16(3):276-89. doi: 10.1016/j.chom.2014.08.014. PMID: 25211071; PMCID: PMC5109542
- Rebersek M. Gut microbiome and its role in colorectal cancer. *BMC Cancer*. 2021 Dec 11;21(1):1325. doi: 10.1186/s12885-021-09054-2. PMID: 34895176; PMCID: PMC8666072.
- Zheng D, et al. Interaction between microbiota and immunity in health and disease. *Cell Res*. 2020 Jun;30(6):492-506. doi: 10.1038/s41422-020-0332-7. Epub 2020 May 20. PMID: 32433595; PMCID: PMC7264227.
- Geuking MB, et al. The interplay between the gut microbiota and the immune system. *Gut Microbes*. 2014 May-Jun;5(3):411-8. doi: 10.4161/gmic.29330. Epub 2014 Jun 12. PMID: 24922519; PMCID: PMC4153781.
- Matson V, et al. Cancer and the Microbiome-Influence of the Commensal Microbiota on Cancer, Immune Responses, and Immunotherapy. *Gastroenterology*. 2021 Jan;160(2):600-613. doi: 10.1053/j.gastro.2020.11.041. Epub 2020 Nov 28. PMID: 33253684; PMCID: PMC8409239.
- Ensm.eu [ESNM (European society of Neurogastroenterology + Motility)], accessed 1/8/23
- Aoun A, et al. The Influence of the Gut Microbiome on Obesity in Adults and the Role of Probiotics, Prebiotics, and Synbiotics for Weight Loss. *Prev Nutr Food Sci*. 2020 Jun 30;25(2):113-123. doi: 10.3746/pnf.2020.25.2.113. PMID: 32676461; PMCID: PMC7333005.
- Wastyk HC, et al. Gut-microbiota-targeted diets modulate human immune status. *Cell*. 2021 Aug 5;184(16):4137-4153.e14. doi: 10.1016/j.cell.2021.06.019. Epub 2021 Jul 12. PMID: 34256014; PMCID: PMC9020749.
- Bolte LA, et al. Long-term dietary patterns are associated with pro-inflammatory and anti-inflammatory features of the gut microbiome. *Gut*. 2021 Jul;70(7):1287-1298. doi: 10.1136/gutjnl-2020-322670. Epub 2021 Apr 2. PMID: 33811041; PMCID: PMC8223641.
- McDonald D, et al; American Gut Consortium; Knight R. American Gut: an Open Platform for Citizen Science Microbiome Research. *mSystems*. 2018 May 15;3(3):e00031-18. doi: 10.1128/mSystems.00031-18. PMID: 29795809; PMCID: PMC5954204.

References:

www.aicr.org. Accessed December 28, 2022.

Andermann TM, Rezvani A, Ghatt AS. Microbiota manipulation with prebiotics and probiotics in patients undergoing stem cell transplantation. *Curr Hematol Malig Rep*. 2016;11:19-28.

Boyle, NM, Podczervinski S, Jordan K, et al. Bacterial foodborne infections after hematopoietic cell transplantation. *Biol Blood Marrow Transplant*. 2014;20:1856-1861.
Fredricks DN. The gut microbiota and graft-versus-host disease. *J Clin Invest*. 2019;129:1808-1817.

Gong D, Yu X, Wang L, et al. Exclusive enteral nutrition induces remission in pediatric Crohn's disease via modulation of the gut microbiome. *Biomed Res*. Published online:2017;2017:8102589.

Gopalakrishnan V, Helmink BA, Spencer CN, et al. The influence of the gut microbiome on cancer, immunity, and cancer immunotherapy. *Cancer Cell*. 2018;33:570-580.

www.heart.org/HealthyForGood. Accessed January 2, 2023.

Helmink BA, Khan MAW, Hermann A, et al. The microbiome, cancer, and cancer therapy. *Nature Medicine*. 2019;25:377-388.

Jenq RR, Taur Y, Devlin SM, et al. Intestinal *Blautia* is associated with reduced death from graft-versus-host disease. *Biol Blood Marrow Transplant*. 2015;21:1373-1383.

Jin Y, Dong H, Xia L, et al. The diversity of gut microbiome is associated with favorable responses to anti-PD-1 immunotherapy in Chinese non-small cell lung cancer patients. *J Thorac Oncol*. 2019;14:1378-1389.

Lassiter M, Schneider SM. A pilot study comparing the neutropenic diet to a non-neutropenic diet in the allogeneic hematopoietic stem cell transplantation population. *Clin J Oncol Nurs*. 2015;19:273-278.

Ligibel JA, Bohlke K, May AMM. Exercise, diet, and weight management during cancer treatment: ASCO guideline. *J Clin Oncol*. 2022;40:2491-2507.

McDonald D, Hyde E, Debellus J, et al. American Gut: an open platform for citizen science microbiome research. *mSystems*. 2018;May 15;3(3):e00031-18.

Moreno-Gonzalez M, Beraza N. The role of the microbiome in liver cancer. *Cancers*. 2021;13:2330.

Muscaritoli M, Arends J, Bachmann P, et al. ESPEN practical guideline: Clinical nutrition in cancer. *Clin Nutr*. 2021;40:2898-2913.