

# FROM MY HEART TO YOUR GUT: LET'S BEAT CANCER (AND OTHER DISEASES) BEFORE THEY EVEN START

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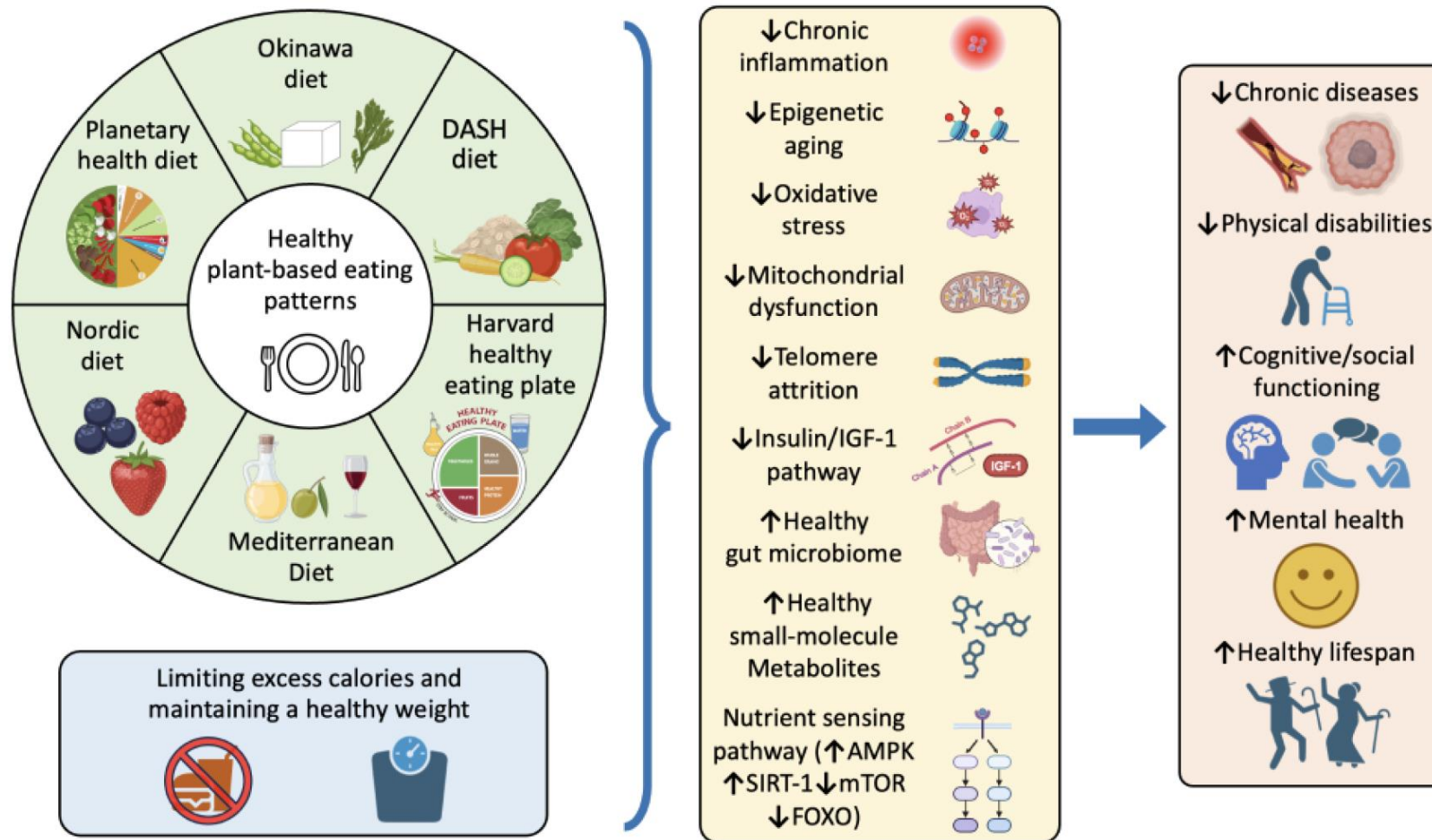
**Baptist Health Chief of Lymphoma Services**

## Stalina

- WWII survivor
- 85 yo
- 2 breast cancers
- 2 knee replacements
- Powered by lentils
- Plant based



# Diet strategies for promoting healthy aging and longevity



Hu FB J of Internal Medicine 2023

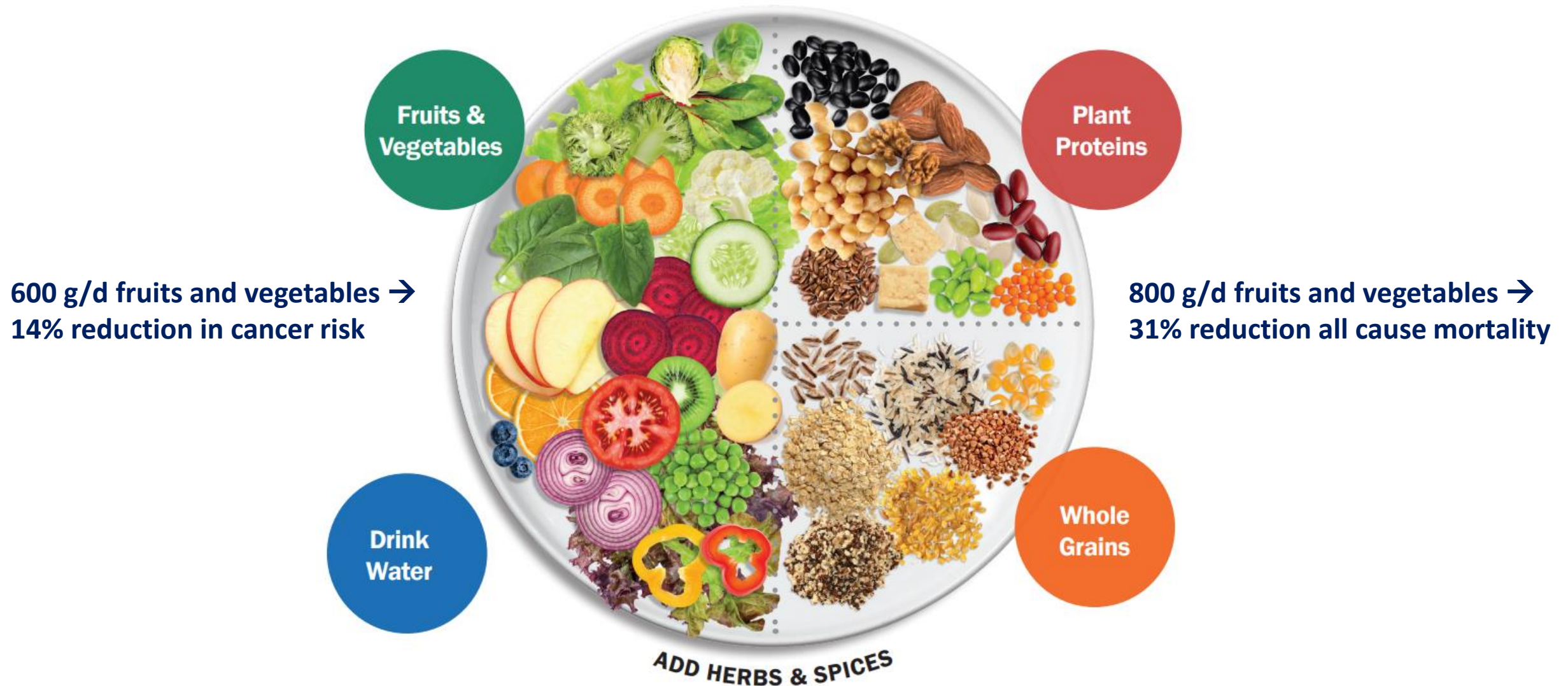
<https://onlinelibrary.wiley.com/doi/full/10.1111/joim.13728>



# A WHOLE FOOD, PLANT-BASED PLATE

## *Nutrition Prescription for Treating & Reversing Chronic Disease*

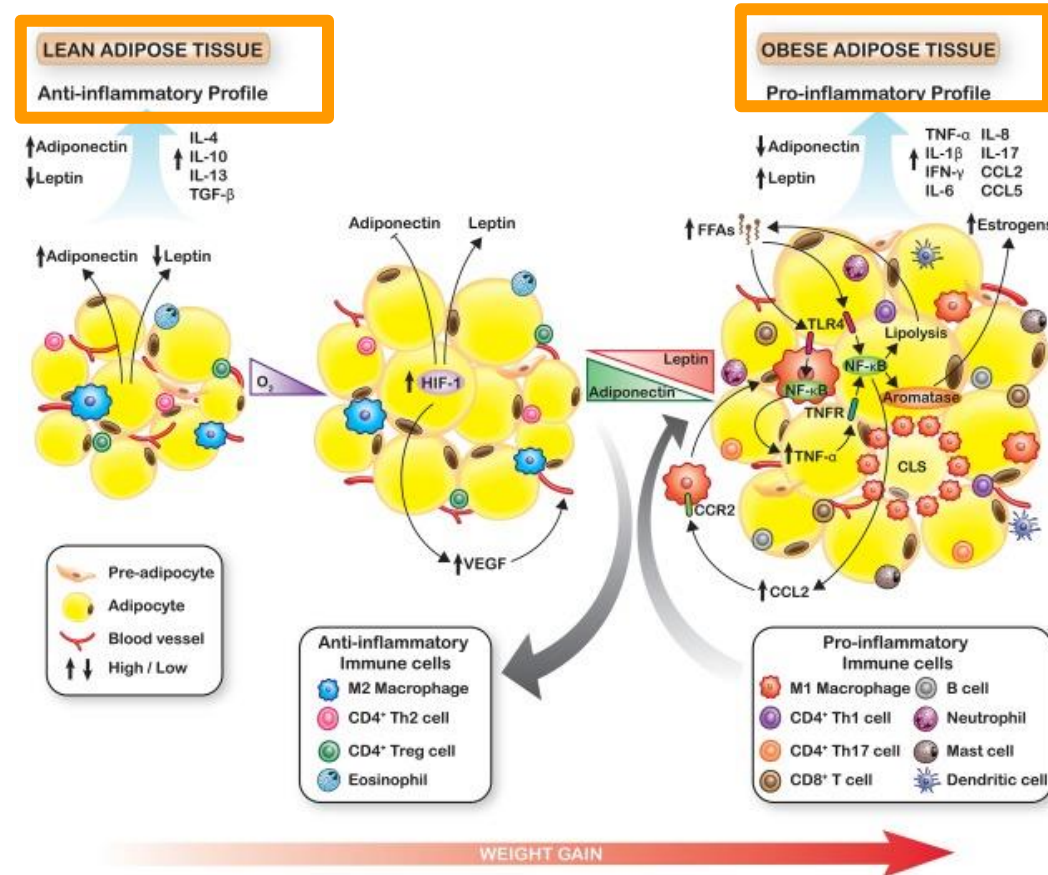
The American College of Lifestyle Medicine Dietary Lifestyle Position Statement for Treatment and Potential Reversal of Disease: ACLM recommends an eating plan based predominantly on a variety of minimally processed vegetables, fruits, whole grains, legumes, nuts and seeds.





# Obesity=Metabolic Disruption Increases the Risk of Most Common Solid and Blood Cancers

- Breast
- Ovarian
- Prostate
- Lung
- Esophageal
- Colon and rectal
- Pancreatic and liver
- Gallbladder
- Thyroid
- Kidney



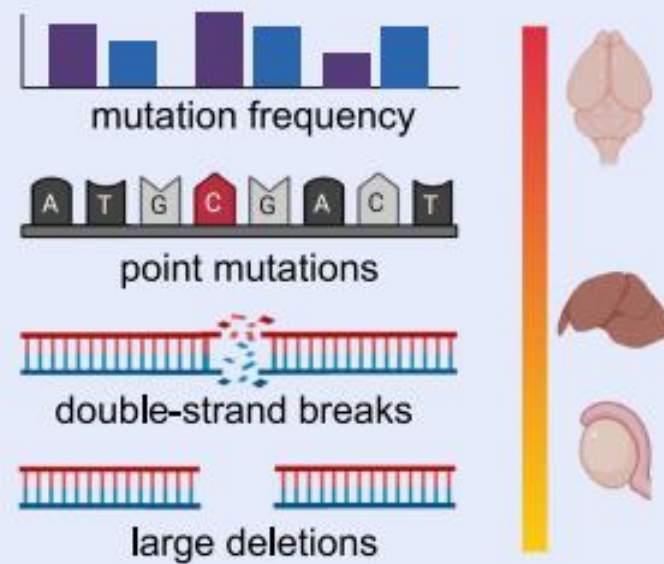
- Multiple myeloma
- Non-Hodgkin lymphoma
- Hodgkin lymphoma
- Acute myeloid leukemia/CHIP
- Acute lymphoblastic leukemia
- CLL

metabolic, endocrinologic, immunologic, and **inflammatory-like changes, oxidative stress** →  
**dysregulated gene function, DNA repair, epigenetic changes** → **CANCER**

## H-DNA mice on HFD



## Genomic instability



Tissue-specific mutagenic impact of obesity on H-DNA-forming sequence

# THE UNFORTUNATE CHAIN OF EVENTS

**POOR DIET**

**METABOLIC SYNDROME**

**Obesity**

**Microbiome change**

**INFLAMMATION**

**FREE RADICALS**  
**CHRONIC**  
**OXYDATIVE STRESS**

**MUTAGENESIS**

**Epigenetic disruption**  
**NF kappa B pathway**

**CANCER**

**Anthocyanins**



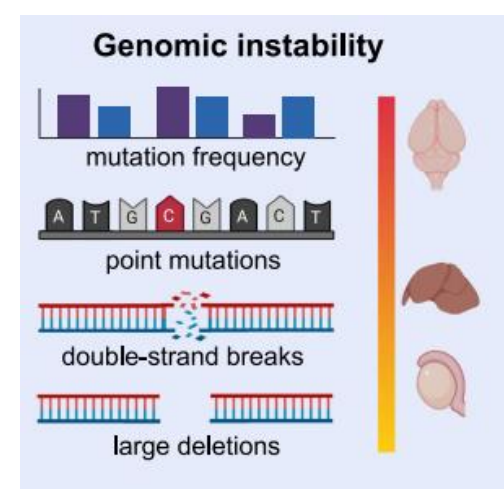
**Sulforaphane**



**Carotenoids**



**Fiber**

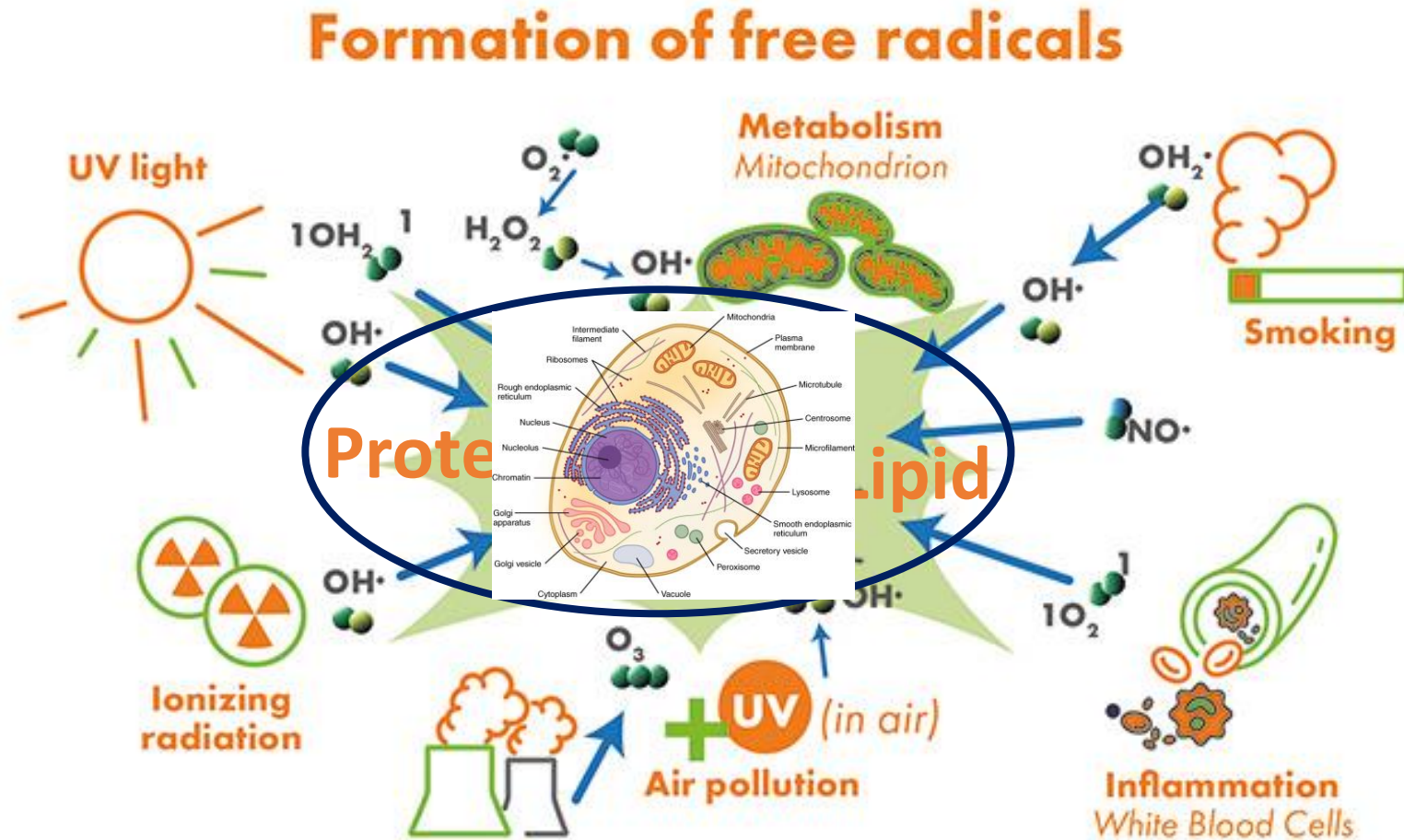




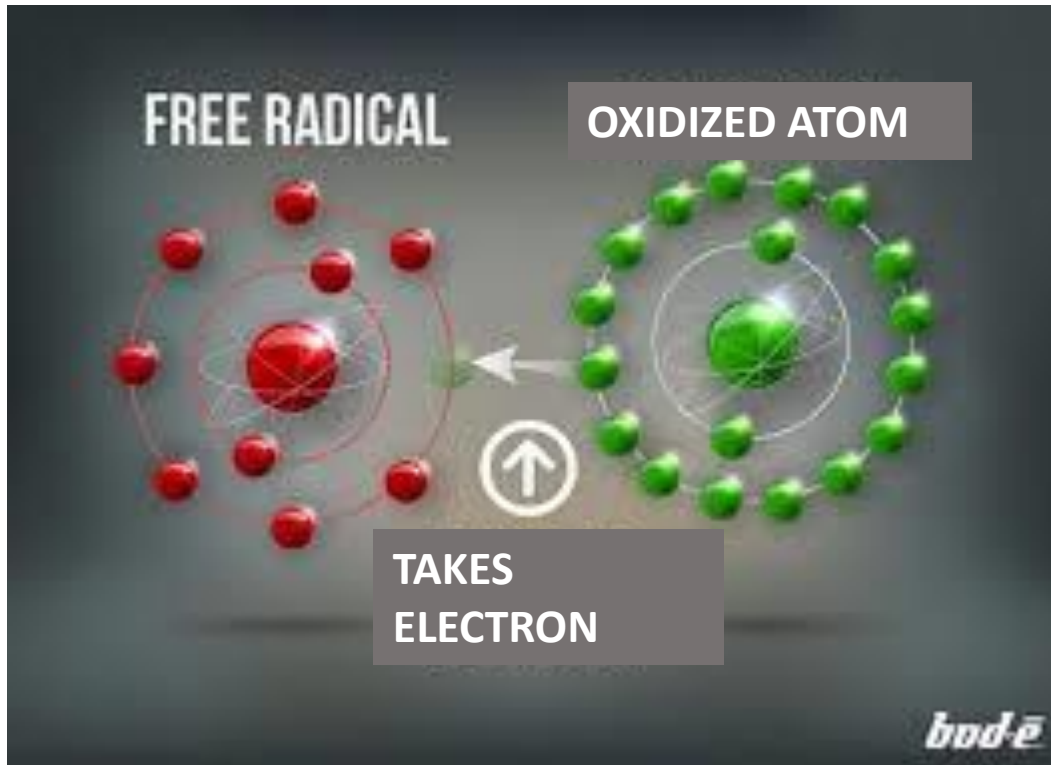
# FREE RADICALS- THEY DAMAGE EVERYTHING!!!!

$O_2^-$ - superoxide  
 $OH\cdot$ - hydroxyl

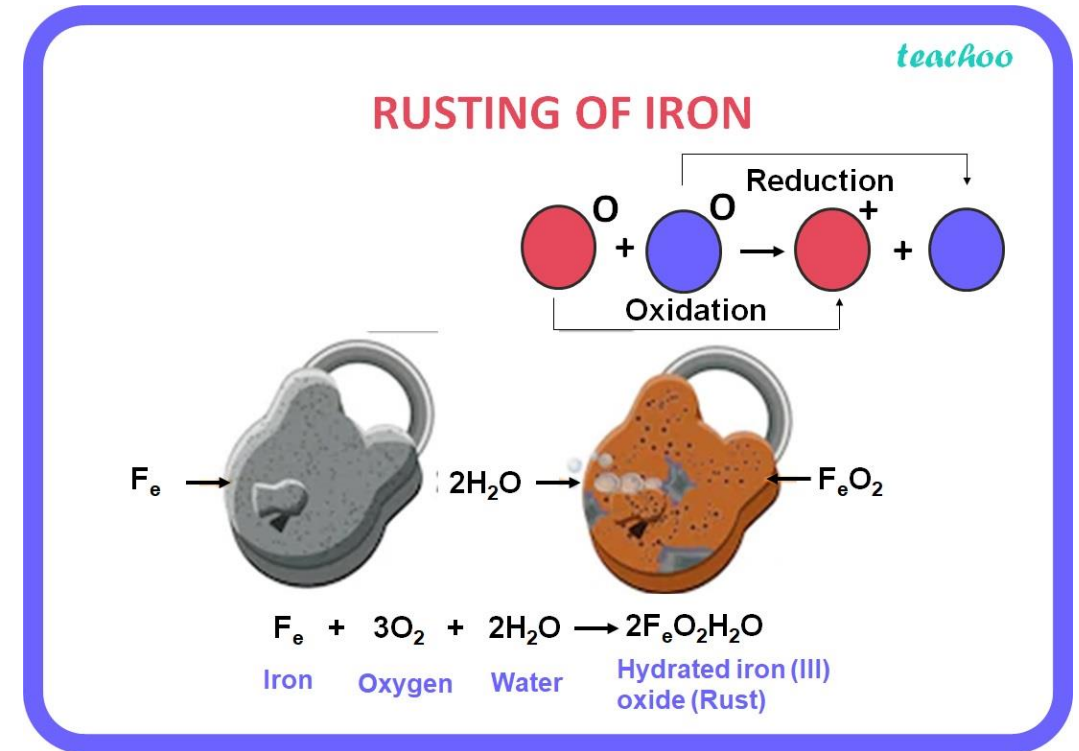
$NO\cdot$ - nitric oxide  
 $HO_2\cdot$ - hydroperoxyl



# FREE RADICALS- ELECTRON THIEVES!!!

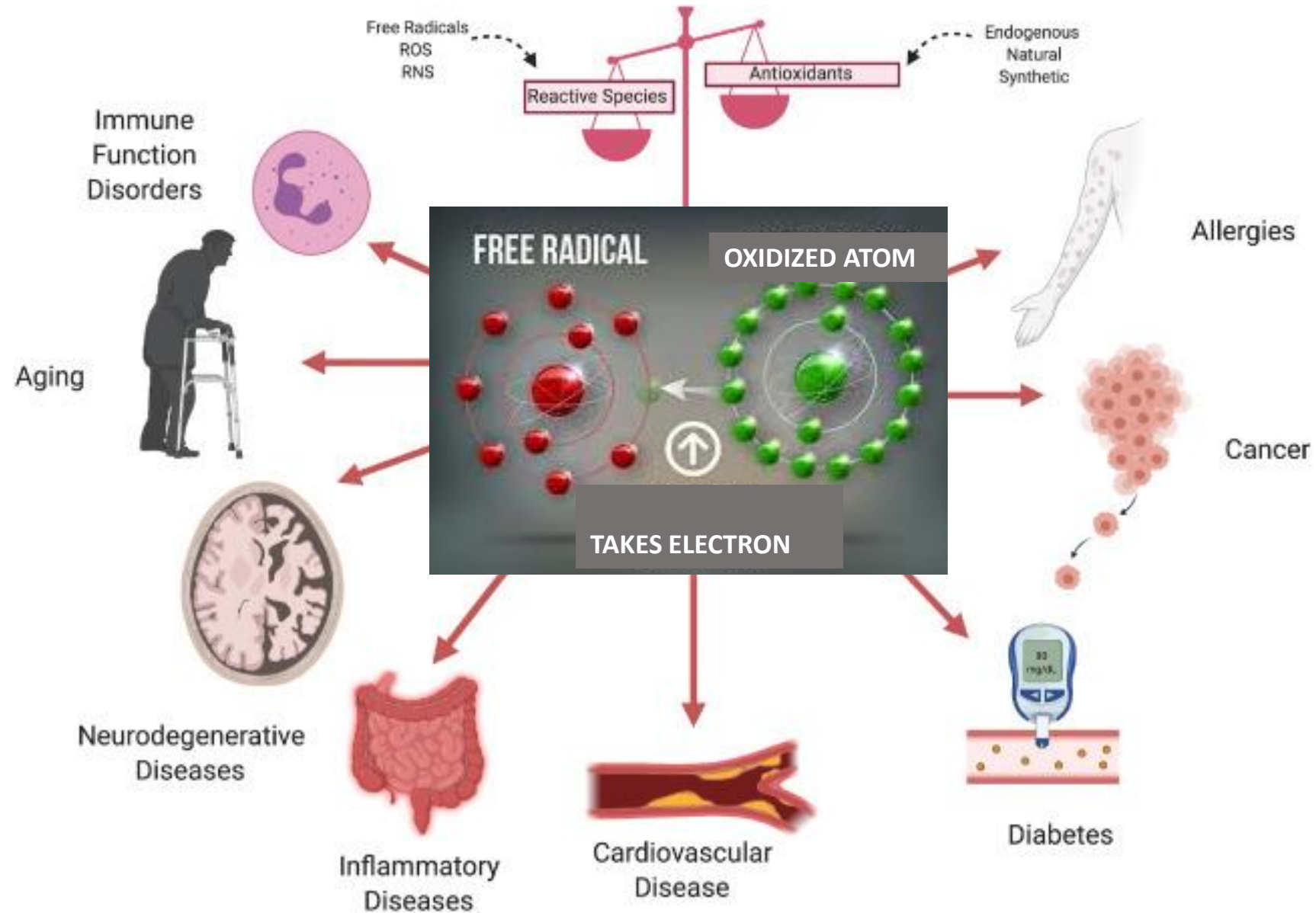


Free radicals:  
O<sub>2</sub>- superoxide  
OH- hydroxyl  
NO- nitric oxide  
HO<sub>2</sub>- hydroperoxyl



We get “rusty” as we go through life,  
but we can slow this process down!!!

# WHEN HUMANS GET RUSTY.....

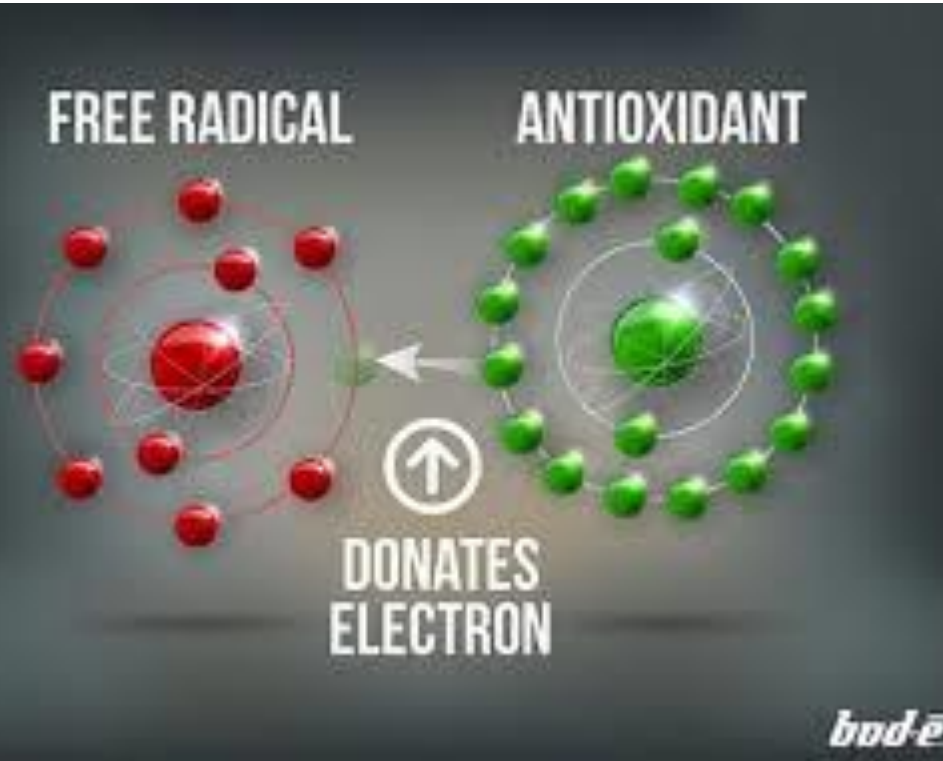




# WE NEED **DONORS** (ELECTRON DONORS😊 TO PROTECT US FROM ELECTRON THIEVES



**Anthocyanins**



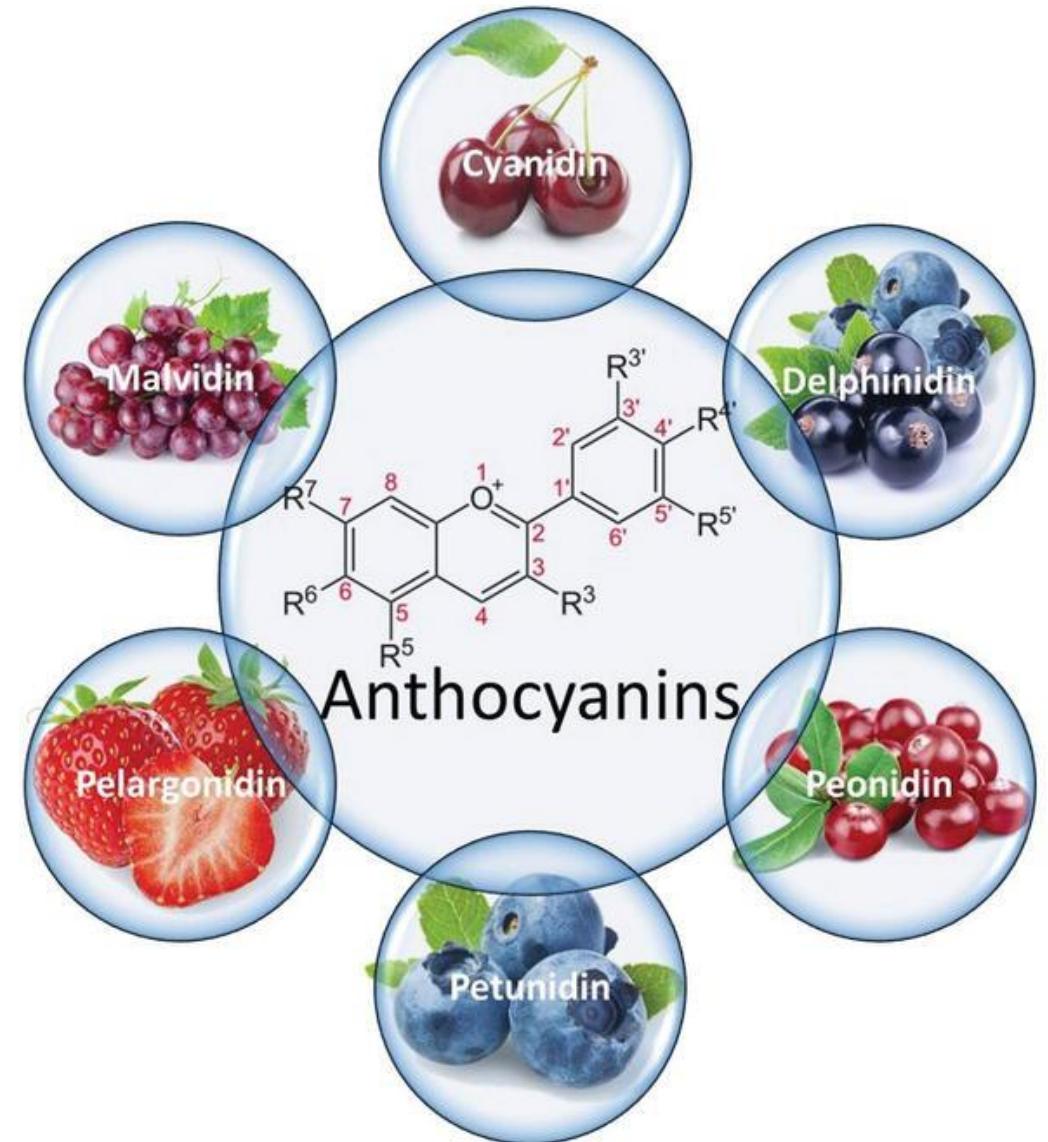
O<sub>2</sub><sup>-</sup> superoxide  
OH<sup>-</sup> hydroxyl  
NO<sup>-</sup> nitric oxide  
HO<sub>2</sub><sup>-</sup> hydroperoxyl



**Carotenoids**



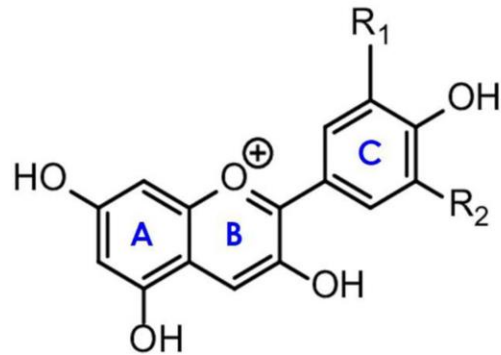
# Anthocyanins- our protectors





# Anthocyanins- Nature's Sunscreen

(a)

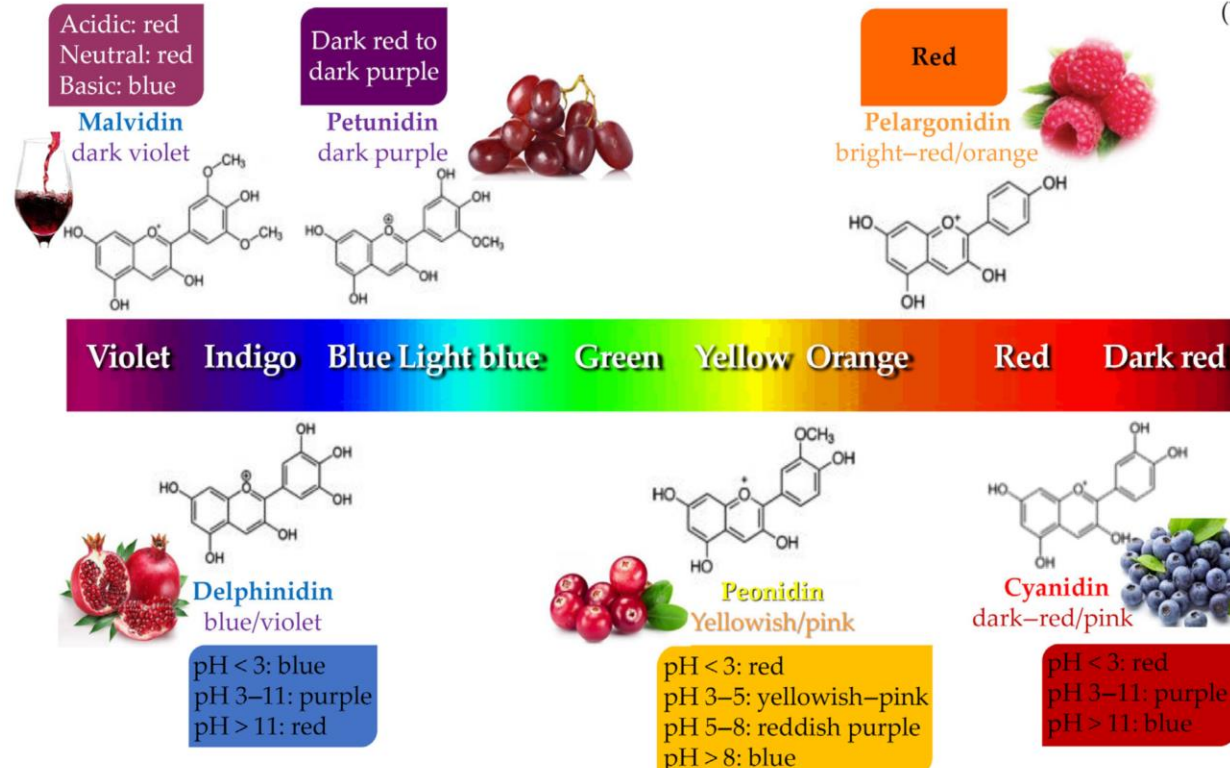


	R1	R2	$\lambda_{max}$
Pelargonidin	H	H	494 nm
Cyanidin	OH	H	504 nm
Delphinidin	OH	OH	508 nm
Peonidin	OCH <sub>3</sub>	H	506 nm
Petunidin	OCH <sub>3</sub>	OH	508 nm
Malvidin	OCH <sub>3</sub>	OCH <sub>3</sub>	510 nm

## PHOTOAGING REPAIR!

- **ABSORB AND DISSIPATE UV LIGHT (esp UVB)**
- **DONATE ELECTRONS TO PROTECT FROM UV OXIDATION (ANTIOXIDANTS)**
- **ANTIINFLAMMATORY**
- **PREVENT CELL DEATH UV INDUCED BY UV LIGHT**
- **INHIBIT COLLAGEN BREAKDOWN**
- **INHIBIT PIGMENTATION**

(b)



Red Cabbage pH Indicator | Kitchen Chemistry for Kids - Science Kiddo

Câmara JS, Locatelli M, Pereira JAM, et al. Behind the Scenes of Anthocyanins-From the Health Benefits to Potential Applications in Food, Pharmaceutical and Cosmetic Fields. *Nutrients*. 2022;14(23):5133. Oliveira H, Correia P, Pereira AR, et al. Exploring the Applications of the Photoprotective Properties of Anthocyanins in Biological Systems. *Int J Mol Sci*. 2020;21(20):7464.

[1] Li K, Zhang M, Chen H, et al. Anthocyanins from Black Peanut Skin Protect Against UV-B Induced Keratinocyte Cell and Skin Oxidative Damage Through Activating Nrf2 Signaling. *Food & Function*. 2019;10(10):6815-6828. doi:10.1039/c9fo00706g.

[2] Guo X, He L, Sun L, et al. Exploring the Potential of Anthocyanins for Repairing Photoaged Skin: A Comprehensive Review.  *Foods (Basel, Switzerland)*. 2024;13(2):3506. doi:10.3390/foods13213506.

[3] Wang Y, Zhang Q, Liu Y, et al. The Protective Effects of Berry-Derived Anthocyanins Against Visible Light Induced Damage in Human Retinal Pigment Epithelial Cells. *Journal of the Science of Food and Agriculture*. 2015;95(5):936-44. doi:10.1002/jsfa.6765.

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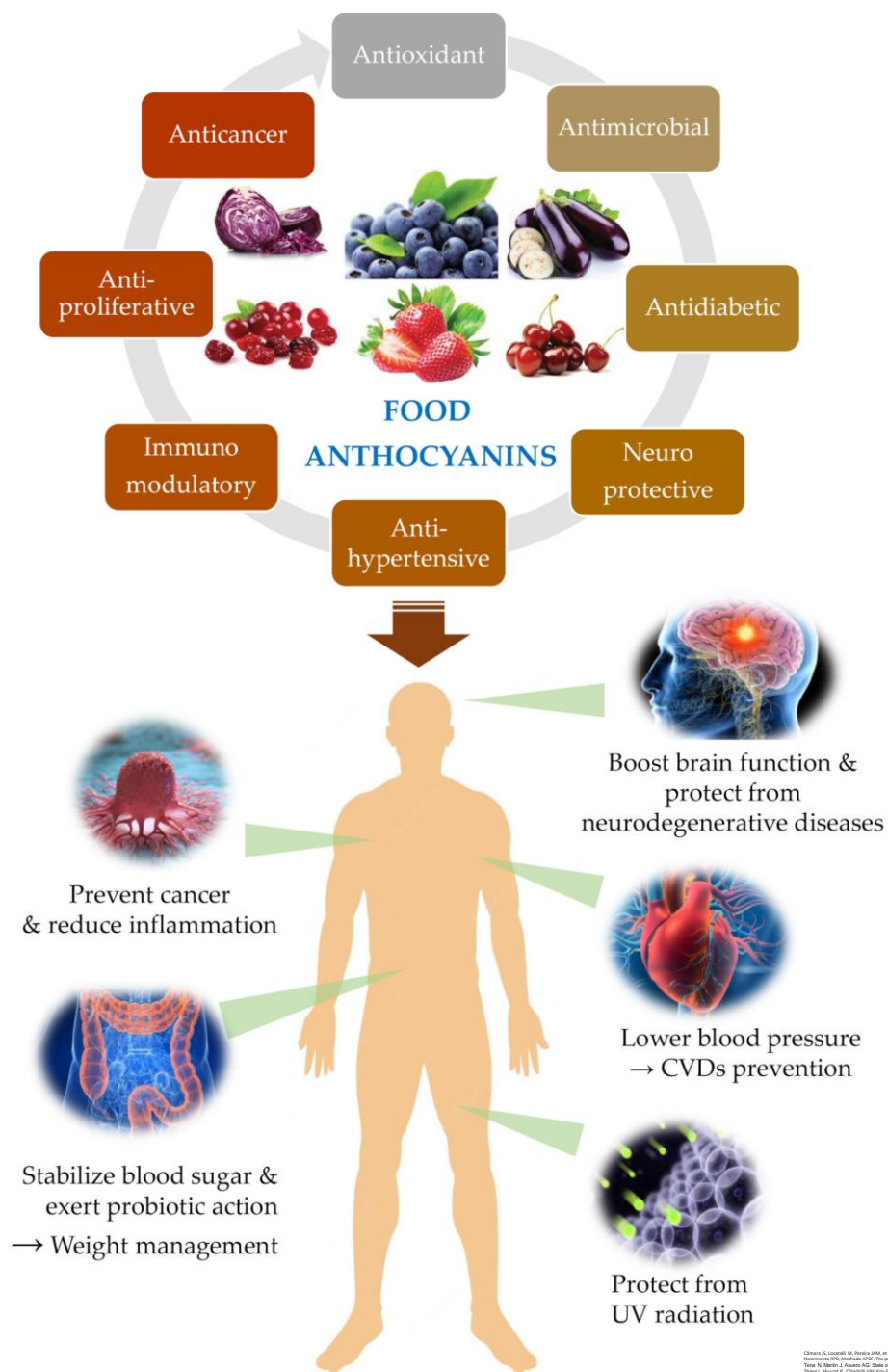
[5] Giampieri F, Alvarez-Suarez JM, Tulipani S, et al. Photoprotective Potential of Strawberry (Fragaria Ananassa) Extract Against UV-A Irradiation Damage on Human Fibroblasts. *Journal of Agricultural and Food Chemistry*. 2012;60(8):2322-7. doi:10.1021/jf202065a.

[6] Guo X, He L, Sun L, et al. Exploring the Potential of Anthocyanins for Repairing Photoaged Skin: A Comprehensive Review.  *Foods (Basel, Switzerland)*. 2024;13(2):3506. doi:10.3390/foods13213506.



# Anthocyanins- Our Most Important Donors

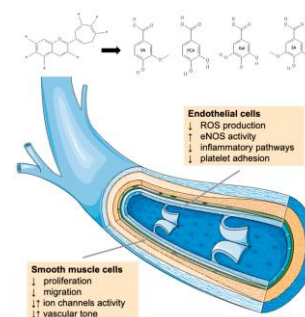
## Protect Us from Diseases Generated by Oxidative Stress



Reduce colorectal cancer risk

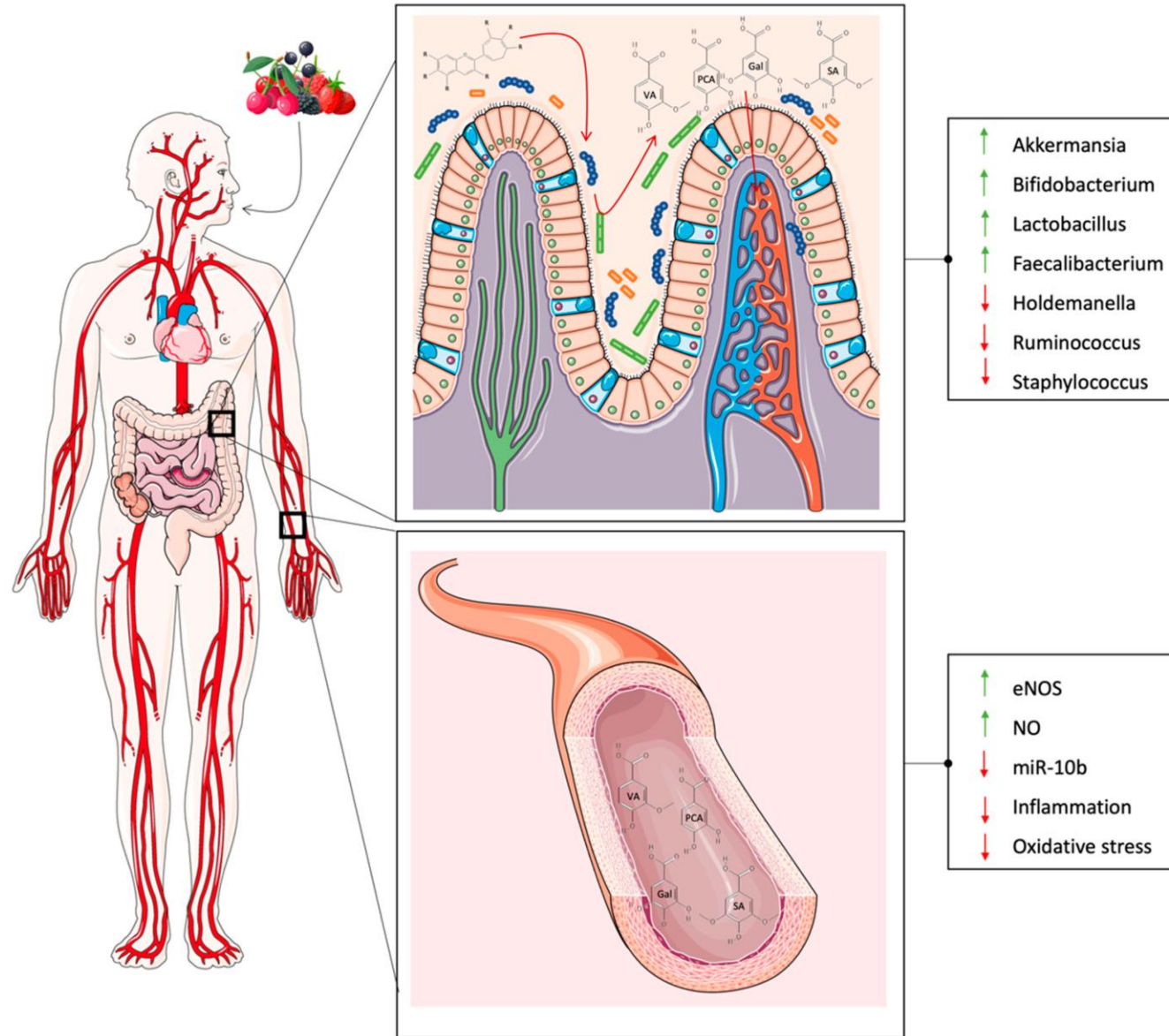


Berries- 20% reduction in all cause mortality  
Anthocyanins 30% reduction all cause mortality and stress related disease risk



40% reduction in cardiovascular disease

# ANTHOCYANINS- BOOST BENEFICIAL GUT BACTERIA







# PURPLE RAIN





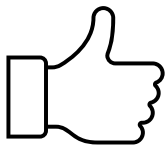
# MY DAILY MEDICINE



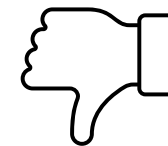
average American spends around \$146 per month on chronic illness medications



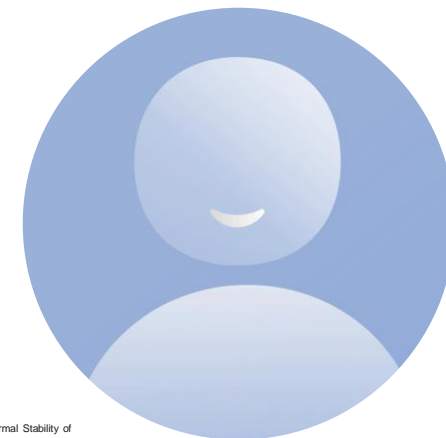




- Water soluble
- RAW
- Fast boiling
- Steaming
- Microwaving
- Save the broth



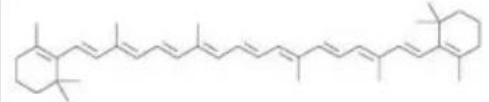



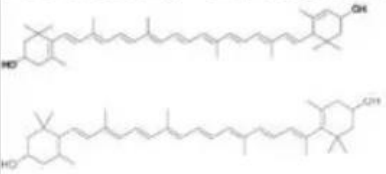

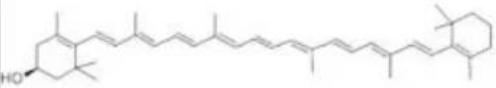



- Boiling x 30 minutes (60-80% decrease)  
11–45% decrease
- Baking
- Frying
- Air-frying
- Stir-frying







TYPE	CAROTENOID FOOD SOURCES
<b>ALPHA-CAROTENE</b> 	 <p>CARROTS, PUMPKIN, WINTER SQUASH, PLANTAINS, COLLARD GREENS</p>
<b>BETA-CAROTENE</b> 	 <p>CARROTS, LEAFY GREENS, SWEET POTATO, CANTALOUPE, PUMPKIN</p>
<b>LYCOPENE</b> 	 <p>TOMATOES, PAPAYA, GRAPEFRUIT, WATERMELON</p>
<b>LUTEIN/ ZEAXANTHIN</b> 	 <p>LEAFY GREENS, SUMMER/ WINTER SQUASH, BRUSSEL SPROUTS, YELLOW CORN</p>
<b>BETA-CRYPTOXANTHIN</b> 	 <p>PUMPKIN, PAPAYA, SWEET PEPPER, ORANGE, CARROT</p>



## ALZHEIMER'S, DEPRESSION, DEMENTIA- DISEASES OF INFLAMMATION

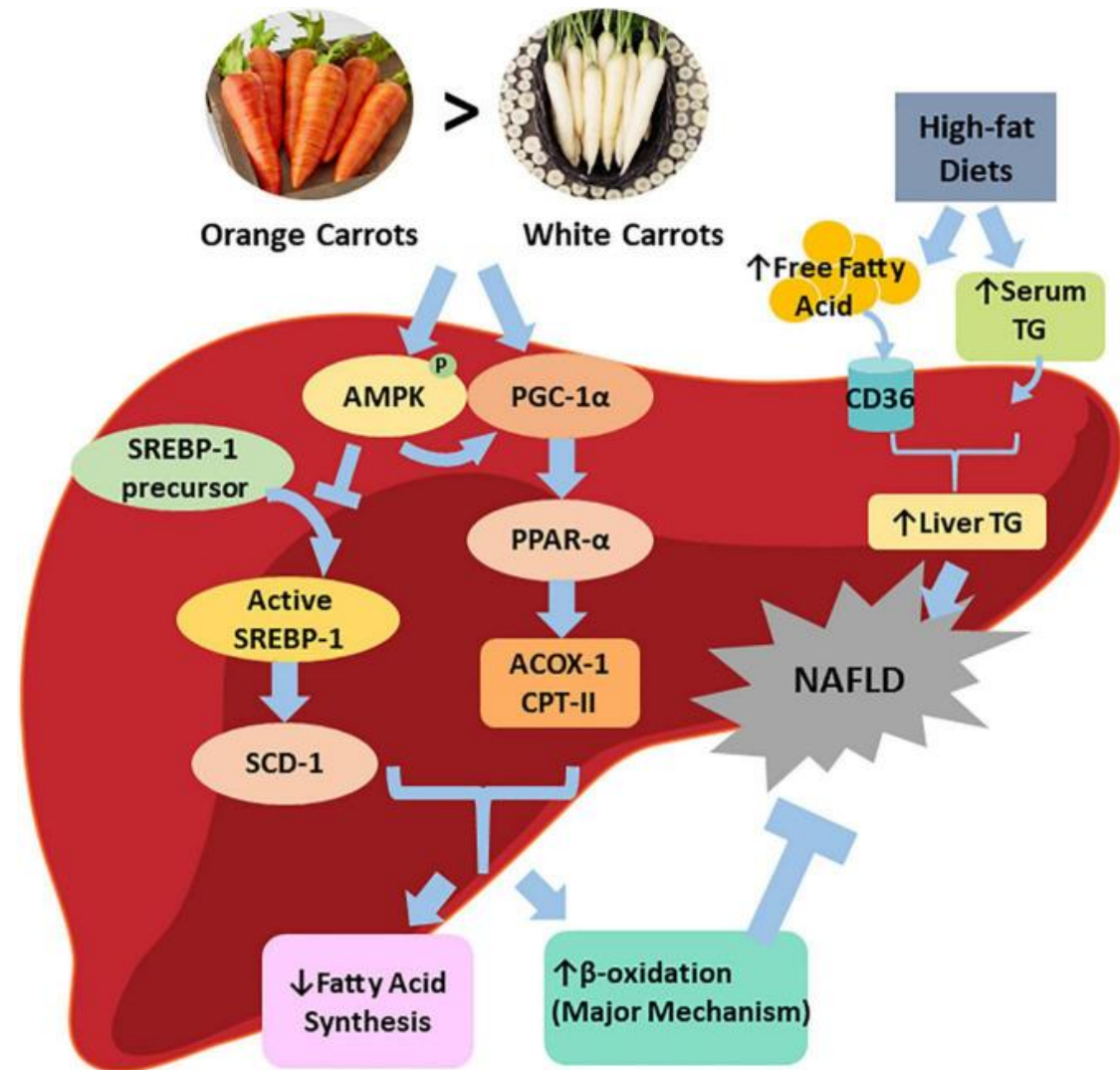
- beta-carotene, lutein, lycopene, and zeaxanthin → low levels in Alzheimer's patients
- higher serum concentrations of lutein, zeaxanthin, and beta-carotene ---> better processing speed, attention, and executive function
- lutein and zeaxanthin supplementation → decrease in cognitive decline and improvements in cognitive tasks; improved memory
- lower levels of carotenoids were associated with depressive symptoms and even predicted the development of new symptoms in older Italians



435 STUDIES ANALYZED  
CAROTENOID INTERVENTION → COGNITIVE FUNCTION IMPROVEMENT



# DON'T FEED YOUR GEESE CARROTS IF YOU WANT FOIS GRAS



# Carotenoids And Cancer Risk- GET THEM FROM FOOD!!!



## BREAST

decreased risk (14%)  
30% at the highest lutein level



## LUNG AND BLADDER

**Beta-carotene supplementation - increased risk**



## GASTROINTESTINAL

Increased tomatoe consumption-27%  
Reduction gastric cancer



## PROSTATE

$\alpha$ -carotene intake and lycopene 13-19% decrease  
1 serving of tomatoe sauce per week ~15% decrease



## ORAL AND PHARYNGEAL

Lycopene,  $\alpha$ -carotene, and  $\beta$ -cryptoxanthin 26% reduction

[4] Dehnavi MK, Ebrahimpour-Koujan S, Lotfi K, Azadbakht L. The Association Between Circulating Carotenoids and Risk of Breast Cancer: A Systematic Review and Dose-Response Meta-Analysis of Prospective Studies. *Advances in Nutrition* (Bethesda, Md.). 2024

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Eliassen AH, Hendrickson SJ, Brinton LA, et al. Circulating Carotenoids and Risk of Breast Cancer: Pooled Analysis of Eight Prospective Studies.

Peng C, Gao C, Lu D, et al. Circulating Carotenoids and Breast Cancer Among High-Risk Individuals.

Leoncini E, Nedovic D, Panic N, et al. Carotenoid Intake From Natural Sources and Head and Neck Cancer: A Systematic Review and Meta-Analysis of Epidemiological Studies. This meta-analysis found that dietary carotenoids are associated with a reduced risk of head and neck cancers

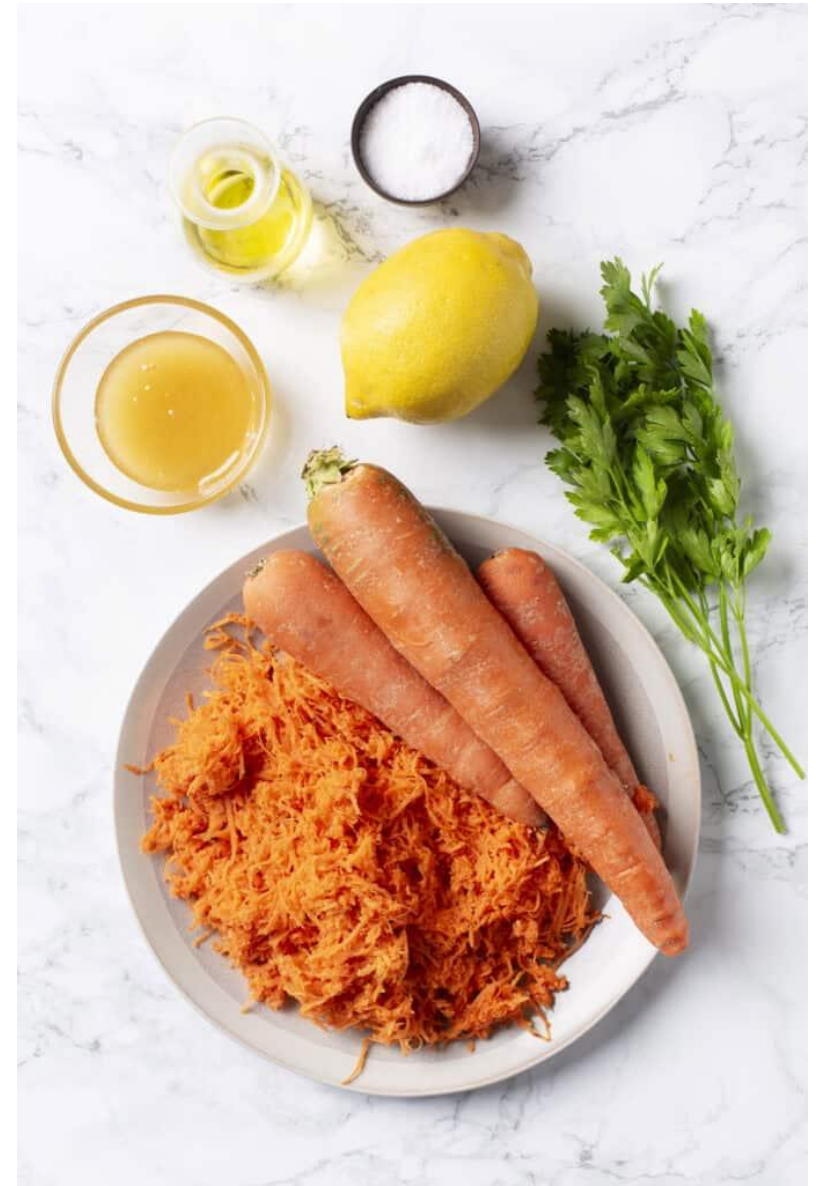
Rowles JL, Erdman JW. Carotenoids and Their Role in Cancer Prevention. This review summarizes the associations between dietary carotenoids and the risk of ten common cancers, suggesting potential benefits in reducing carcinogenesis.]





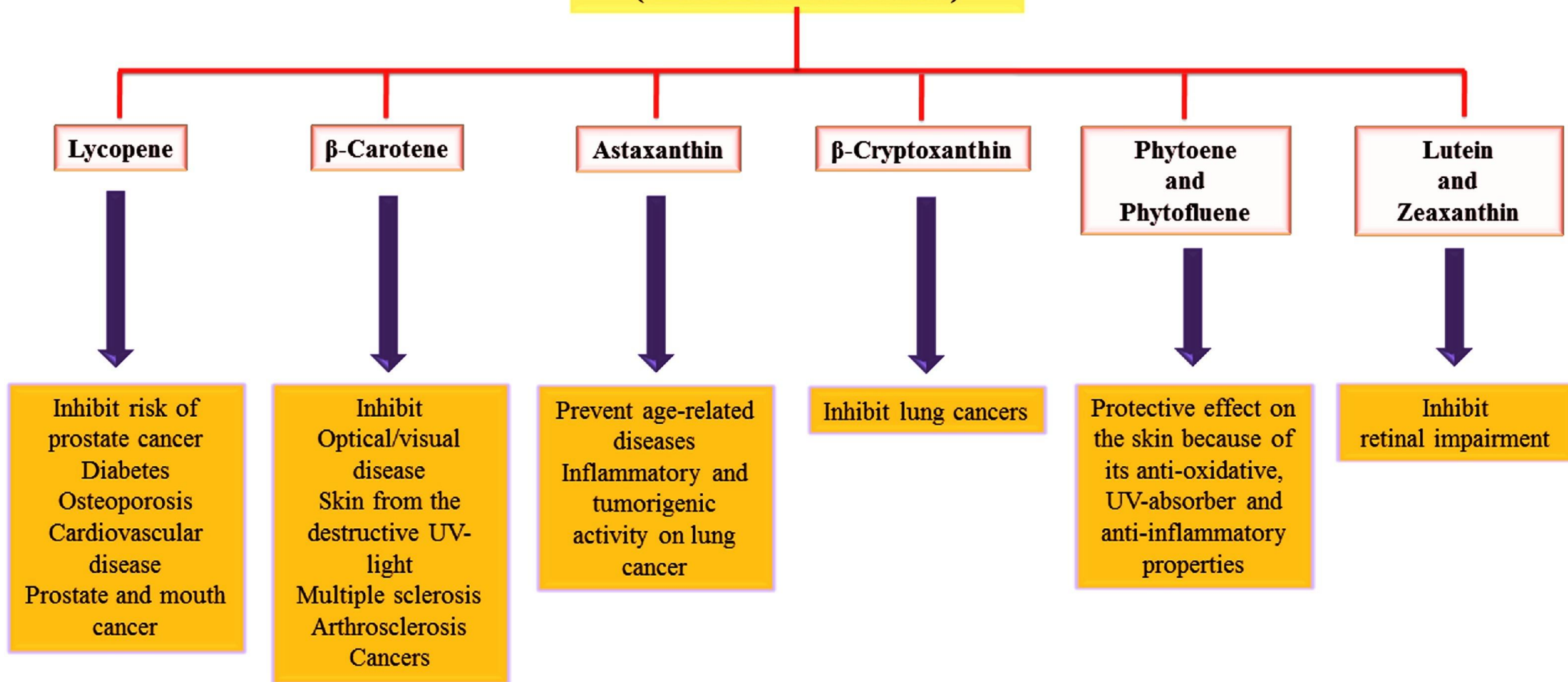
Heating and cooking lycopene-rich foods (e.g., tomatoes cooked into tomato sauce) increases bioavailability and absorption

HEAT AND FAT



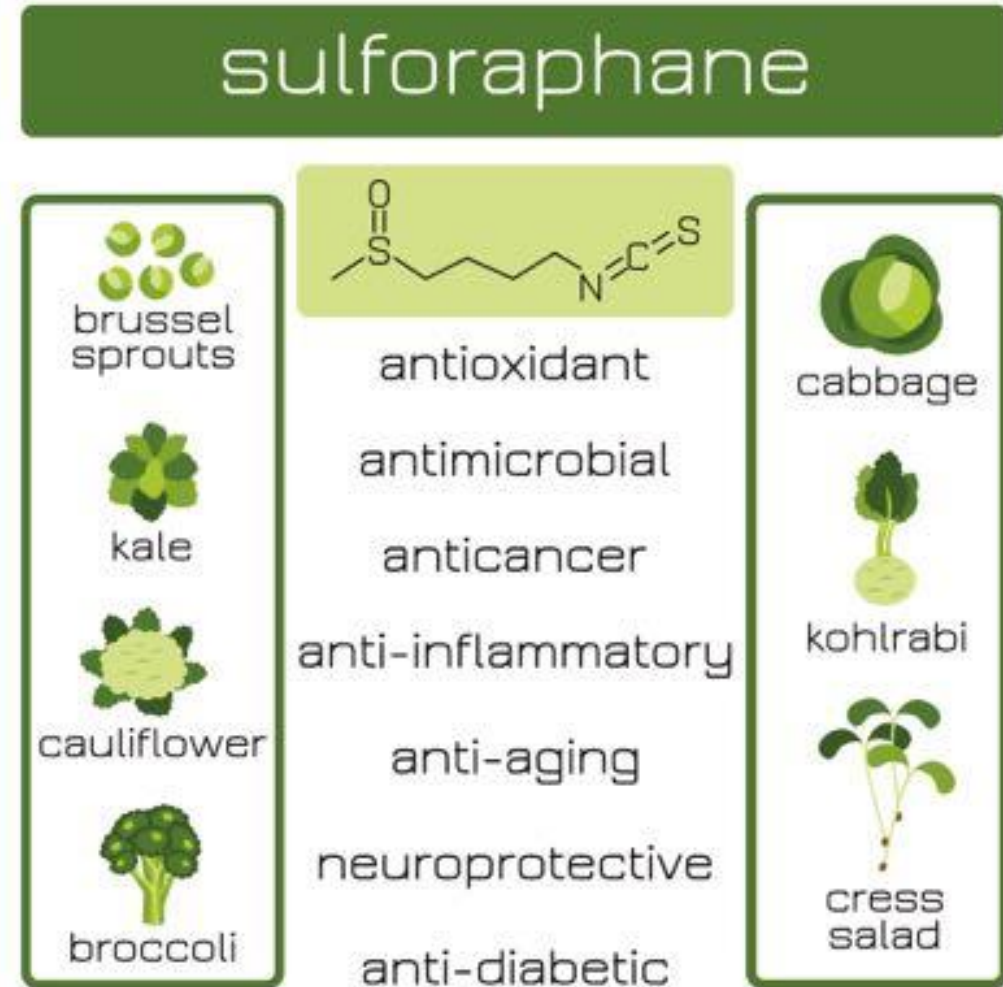
Carotenoids are fat soluble-best absorbed with fat

## **Carotenoids (Antioxidant Activities)**





# CRUCIFEROUS CRUSADE: HOW BRUSSEL SPROUTS PROTECT YOU FROM COLON CANCER



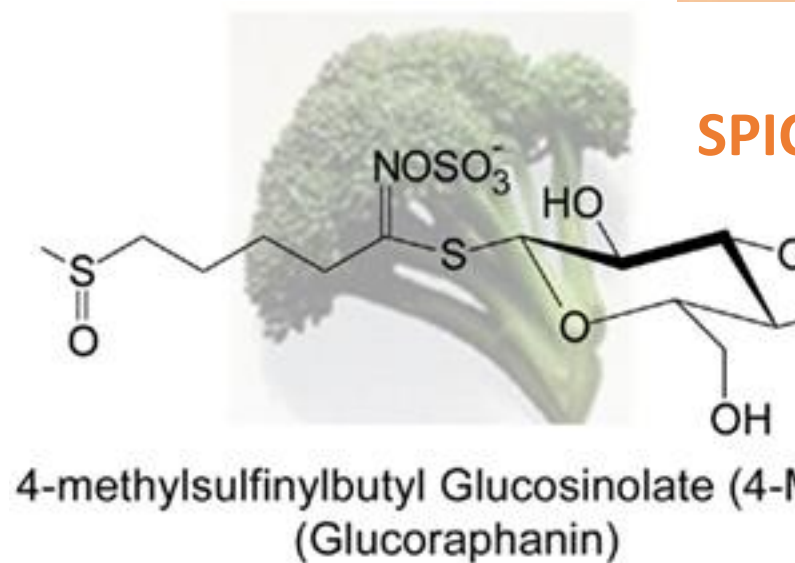




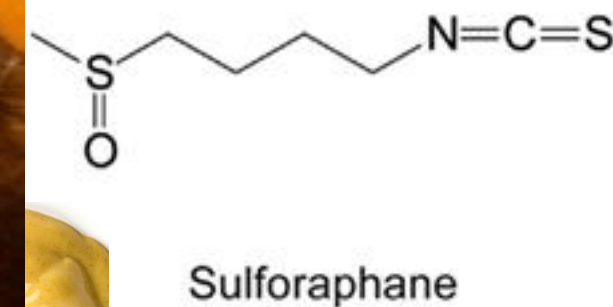
**RAW (chop, let sit x 40 min)**



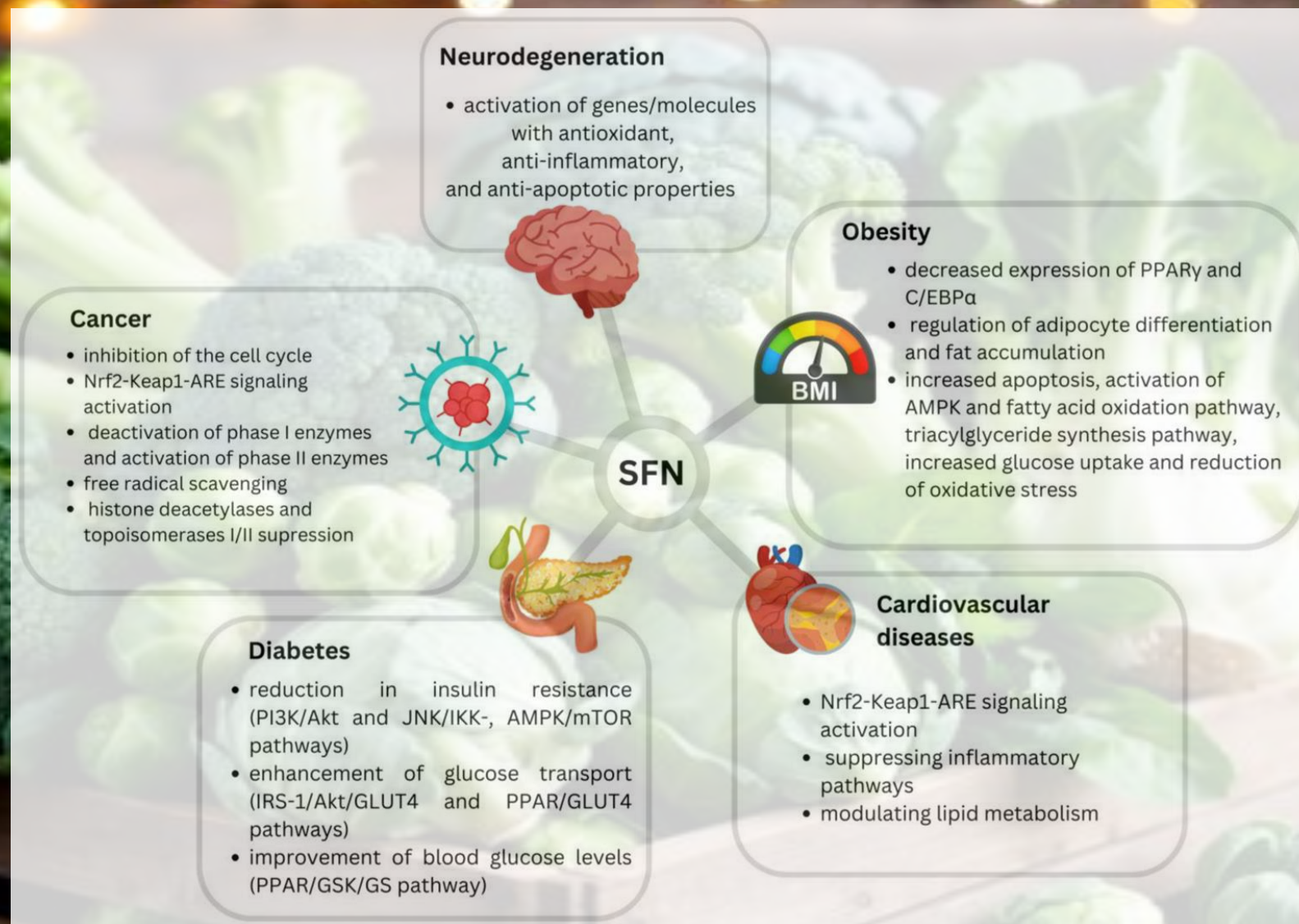
**STEAM <5 min**  
**Microwave < 2min**



**TARD!!!!**



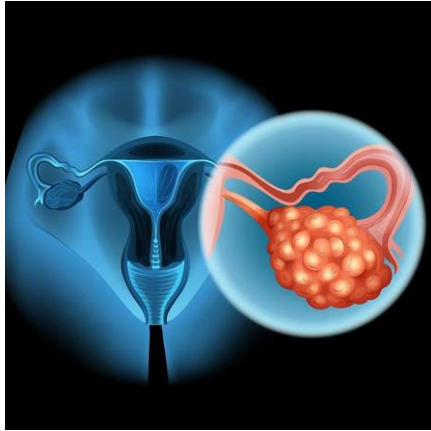




**Scientists analyzed data from the U.S. National Health and Nutrition Examination Survey. Those who ate broccoli 1-2 times a week had 30-40% lower risk of dying from any cause. Higher broccoli consumption is associated with lower risk of dying from cancer or cardiovascular disease. Cruciferous vegetable consumption in general is inversely associated with all cancer risk.**



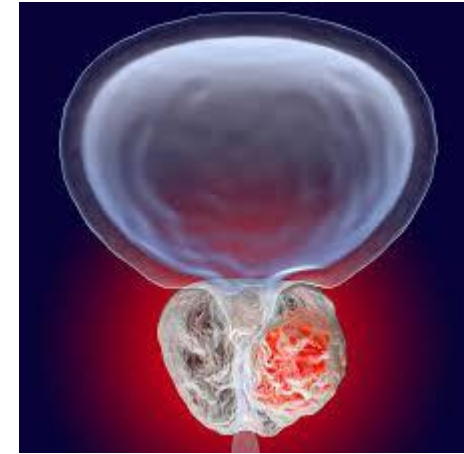
# CRUCIFEROUS CRUSAID TO FIGHT CANCER!!!



**OVARIAN  
ENDOMETRIAL**



**GASTROINTESTINAL**  
Esophageal  
gastric  
liver  
pancreatic  
colorectal cancers



**PROSTATE**

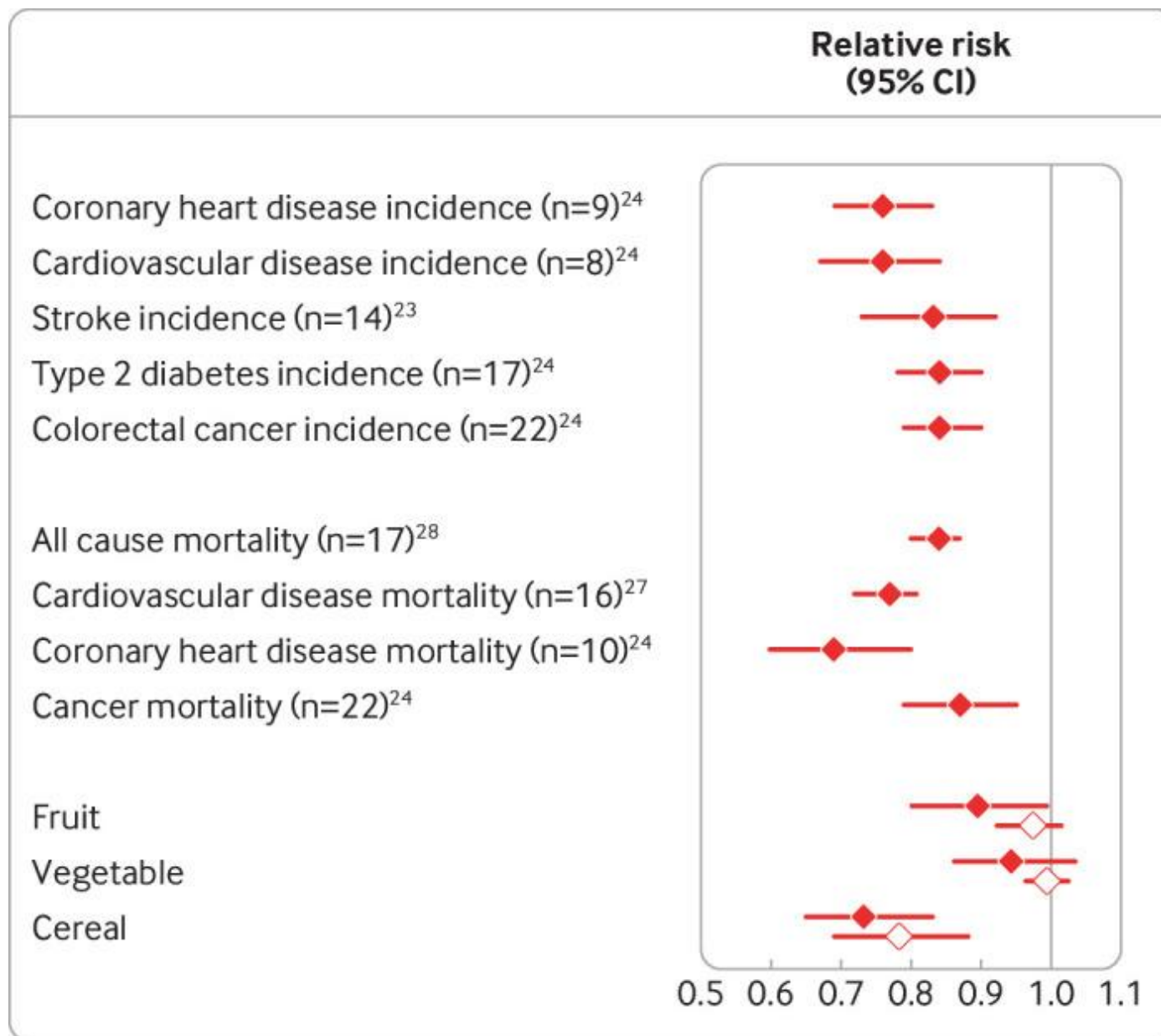


**LUNG**



**RENAL CELL AND BLADDER**

# FIBER FIBER FIBER !!!! FIBER INTAKE FOR OPTIMAL HEALTH

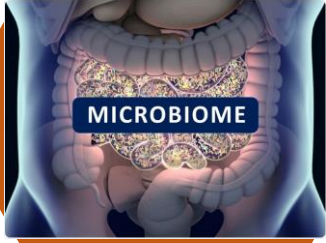


**Females > 25 g (best >40g)**  
**Men > 40g (best >45g)**

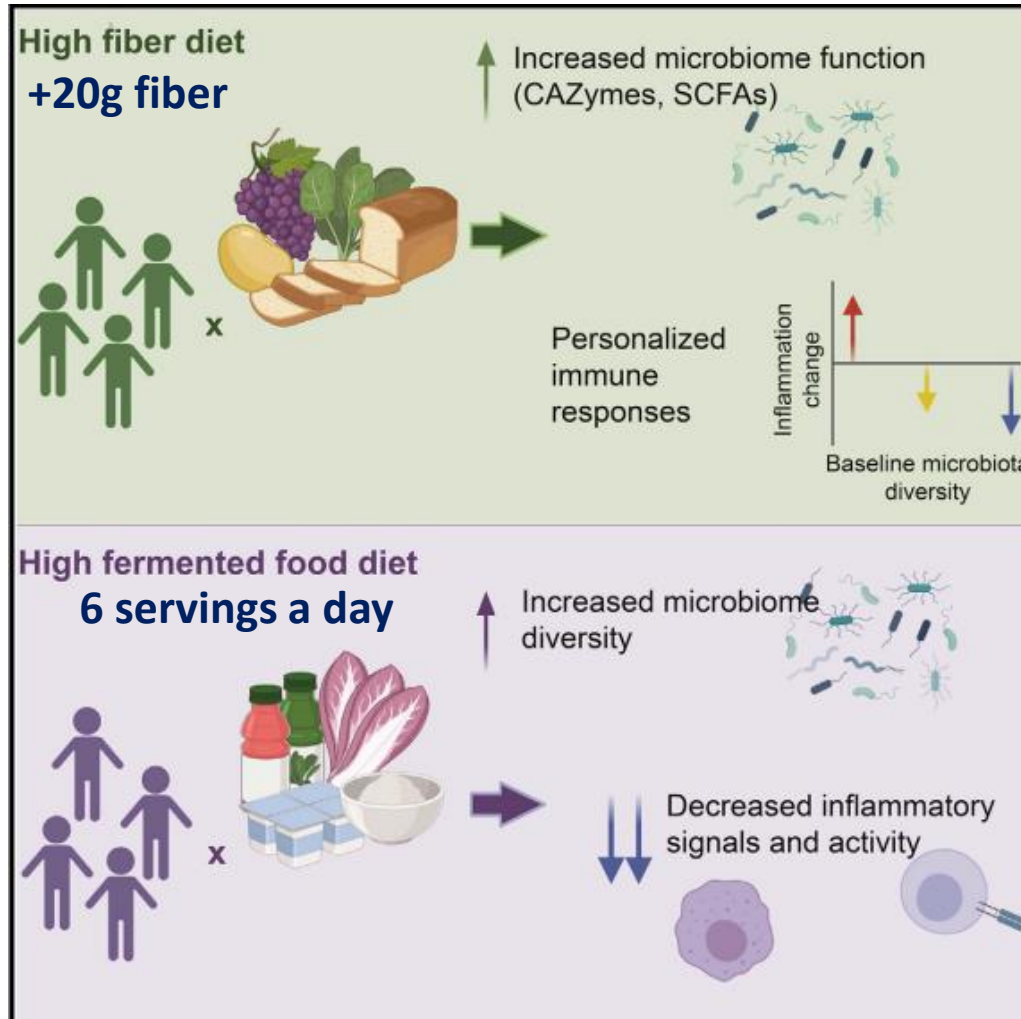
**800 g/d fruits and vegetables →  
31% reduction all cause mortality**



# FIBER FIBER FIBER and EVEN BETTER FERMENTED FIBER!!!



10 weeks

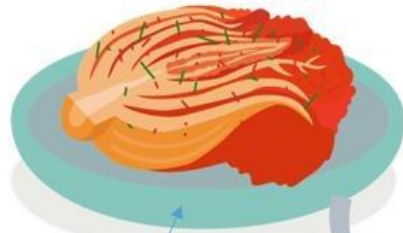


kombucha yogurt  
kefir buttermilk  
kvass = 6 oz  
kimchi sauerkraut  
other fermented  
veggies = 1/4 cup  
vegetable brine  
drink = 2 oz

- Increase in bacterial diversity
  - 19 markers of inflammation decreased
  - IL-6 decreased
- (key mediator of chronic inflammation, elevated in rheumatoid arthritis, type-2 diabetes, and chronic stress)

[Superfood-List.pdf \(lifestylemedicine.org\)](#)

## Kimchi



- A food rich in lactic acid bacteria, probiotics, and Phytochemical

- Partial degradation of phytochemicals

- Fermentation of dietary fiber
- Production of metabolites
- Maintenance of microbial balance
- Influence on host metabolism

## Effects of kimchi consumption on body fat & intestinal microbiota in overweight participants

### Alteration in the gut microbiota

- *A. muciniphila* ↑
- Proteobacteria ↓

Microbial metabolism in the colon

### Lower the risk of health condition

- Body fat mass ↓
- Triglyceride ↓





# NATURE'S OZEMPIC IN OUR KITCHEN--FOODS THAT INCREASE GLP-1

- **Anthocyanins**- all the purple fruits and vegetables
- **Beta glucan fiber**- (oat and barley bran)
- **Legumes**: soy, green lentils, peas
- **Quercetin**  
(apples with skin, berries, citrus, onions, capers, green tea)
- **Curcumin (turmeric)**
- **Sesquiterpenoids**  
(ginger, turmeric, black pepper, chamomile, green tea)
- **Berberine - barberry**
- **Resveratrol**- grapes, berries, dark chocolate
- **Cinnamon**
- **Whey protein (cheese water)**

## References:

- [1] Anghel SA, Badea RA, Chiritoiu G, et al. Novel Luciferase-Based Glucagon-Like Peptide 1 Reporter Assay Reveals Naturally Occurring Secretagogues. *British Journal of Pharmacology*. 2022;179(19):4738-4753. doi:10.1111/bph.15896.
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- [9] Obarakpo JU, Liu L, Zhang S, et al. In Vitro Modulation of Glucagon-Like Peptide Release by DPP-IV Inhibitory Polyphenol-Polysaccharide Conjugates of Sprouted Quinoa Yoghurt. *Food Chemistry*. 2020;324:126857. doi:10.1016/j.foodchem.2020.126857.
- [10] Shimizu Y, Hara H, Hira T. Glucagon-Like Peptide-1 Response to Whey Protein Is Less Diminished by Dipeptidyl Peptidase-4 in Comparison With Responses to Dextrin, a Lipid and Casein in Rats. *The British Journal of Nutrition*. 2021;125(4):398-407. doi:10.1017/S0007114520002834.
- [11] Hira T, Trakooncharoenwit A, Taguchi H, Hara H. Improvement of Glucose Tolerance by Food Factors Having Glucagon-Like Peptide-1 Releasing Activity. *International Journal of Molecular Sciences*. 2021;22(12):6623. doi:10.3390/ijms22126623.
- [12] Di Stefano E, Hüttmann N, Dekker P, et al. Solid-State Fermentation of Green Lentils by Lactiplantibacillus Plantarum Leads to Formation of Distinct Peptides That Are Absorbable and Enhances DPP-IV Inhibitory Activity in an Intestinal Caco-2 Cell Model. *Food & Function*. 2024;15(22):11220-11235. doi:10.1039/d4fo03326d.



# PROTEIN MYTH BUSTERS

**MYTH:** Most people do not get enough protein.

**FACT:** The average person consumes too much protein.

**How much protein do you need per day?**

19-51+ years old\*



\*The general recommendation for protein intake is 0.8g/kilogram or 0.36g/pound.

The average adult person consumes 80 grams of protein per day – far more than is necessary! Excess protein is stored as fat and can lead to weight gain or prevent weight loss. Our bodies benefit from consuming mostly fruits, vegetables, whole grains, beans, and legumes to provide healthy sources and amounts of carbohydrate, fat, and protein.

Sources: <https://health.gov/dietaryguidelines/2015/guidelines/appendix-7/>  
J Acad Nutr Diet. 2013 Dec; 113(12): 1610–1619.

# HAVE YOU BEEN LIED TO YOUR ENTIRE LIFE??

## 9 PLANT SOURCES OF 9 ESSENTIAL AMINO ACIDS



quinoa



amaranth



chia seeds



spirulina



buckwheat



soy (tofu, tempeh)



hemp seed



nutritional yeast





**cerabeta™**

BARLEY BETA-GLUCAN  
BETA-GLUCANE D'ORGE

LOWERS CHOLESTEROL NATURALLY  
ABAISSER LE CHOLESTEROL NATURELLEMENT

4g FIBER	30 CALORIES	0g SUGARS
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PLANT-BASED  
COLLAGEN · PREBIOTIC · FIBER

CLEANSE · ENERGIZE · FOCUS · FEEL FULL LONGER

**ZenBasil™**  
ORGANIC BASIL SEEDS

In a gram to gram comparison with Chia seeds, ZenBasil seeds have nearly:	2X THE FIBER	2X THE CALCIUM
	2X THE POTASSIUM	2X THE IRON

NATURALLY:  
GLUTEN FREE  
LECTIN FREE  
ORGANIC  
KETO  
PALEO

**15g  
FIBER**  
84% Daily Value  
per 30g Serv



14 oz | 390 g



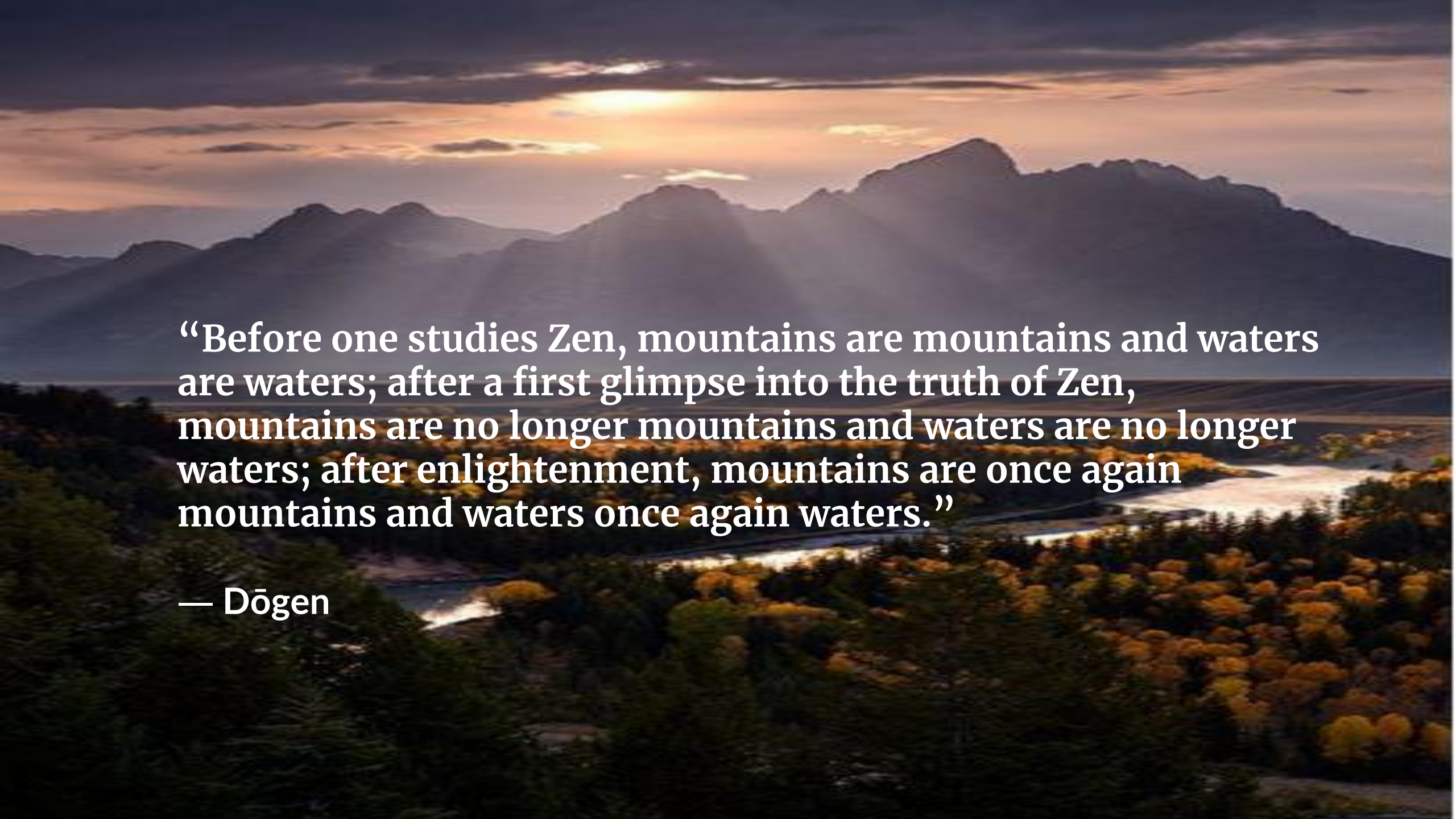
**ORGANIC  
HEMP  
HEARTS**



N WITH LESS GREENS AND MORE HEART

NET WT





**“Before one studies Zen, mountains are mountains and waters are waters; after a first glimpse into the truth of Zen, mountains are no longer mountains and waters are no longer waters; after enlightenment, mountains are once again mountains and waters once again waters.”**

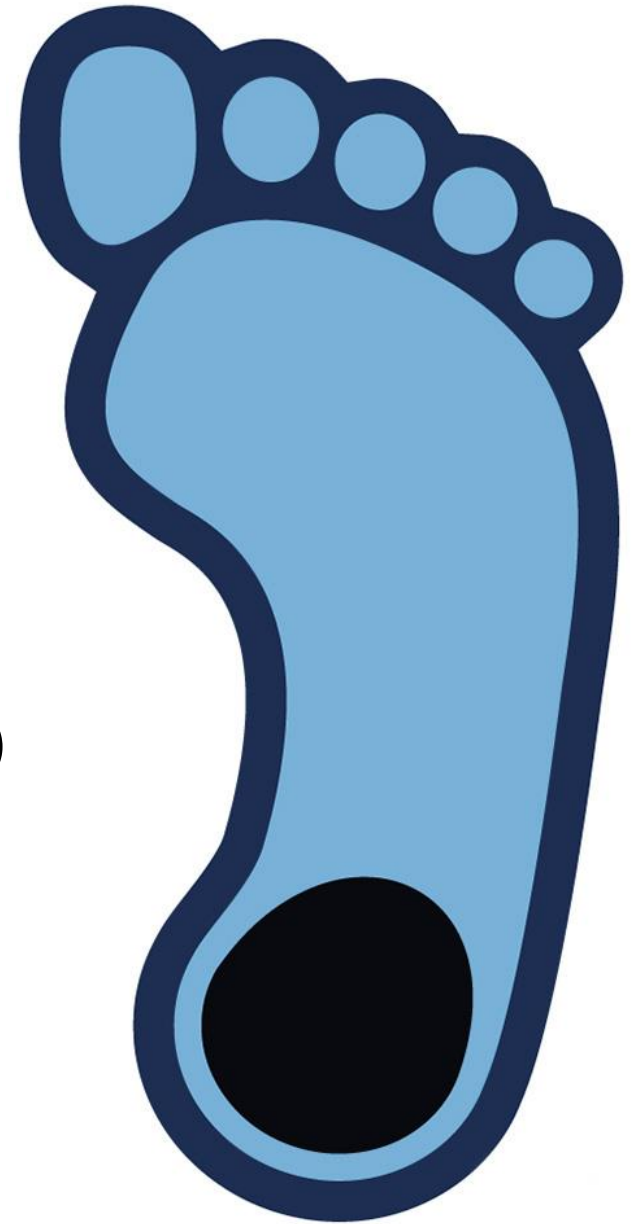
**— Dōgen**



# Yuliya Linhares, MD *with distinction*



- FIU Associate Professor, Translational Medicine
- Chief of Lymphoma at MCI 2019-
- Baptist Health- top 10% in lymphoma, myeloma and leukemia
- Principal Investigator (10 clinical trials)
- 10 publications, 23 abstracts (2019-2025)



# Yuliya Linhares, MD– SELCTED BIBLIOGRAPHY

## Brentuximab vedotin, nivolumab, doxorubicin, and dacarbazine for advanced-stage classical Hodgkin lymphoma.

Lee HJ, Ramchandren R, Friedman J, Melear J, Flinn IW, Burke JM, Linhares Y, Gonzales P, Peterson M, Raval M, Chintapatla R, Feldman TA, Yimer H, Islas-Ohlmayer M, Patel A, Metheny L, Dean A, Rana V, Gandhi MD, Renshaw J, Ho L, Fanale MA, Guo W, Yasenchak CA.

Blood. 2025 Jan 16;145(3):290-299. doi: 10.1182/blood.2024024681.

## OUTREACH: phase 2 study of lisocabtagene maraleucel as outpatient or inpatient treatment at community sites for R/R LBCL.

Linhares Y, Freytes CO, Cherry M, Bachier C, Maris M, Hoda D, Varela JC, Bellomo C, Cross S, Essell J, Fanning S, Terebelo H, Yimer H, Courtright J, Sharman JP, Kostic A, Vedal M, Ogasawara K, Avilion A, Espinola R, Yuan B, Mattar B.

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PMID: 37883980

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Brothers J, Castillo DR, Jeon WJ, Joung B, Linhares Y.

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Linhares Y, Kaganski A, Agyare C, Kurnaz IA, Neergheen V, Kolodziejczyk B, Kędra M, Wahajuddin M, El-Youssf L, Dela Cruz TE, Baran Y, Pešić M, Shrestha U, Bakiu R, Allard PM, Rybtsov S, Pieri M, Siciliano V, Flores Bueso Y.

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Rubens M, Ramamoorthy V, Saxena A, Das S, Bhatt C, Veledar E, McGranaghan P, Sundil S, Shehadeh N, Viamonte-Ros A, Linhares Y, Odia Y, Chuong M, Kotecha R, Mehta MP.

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## Bone marrow transplant-associated thrombotic microangiopathy without peripheral blood schistocytes: a case report and review of the literature.

Wirtschafter E, VanBeek C, Linhares Y.

Exp Hematol Oncol. 2018 Jun 22;7:14. doi: 10.1186/s40164-018-0106-9. eCollection 2018.

PMID: 29977661 Free PMC article.



## Brentuximab vedotin, nivolumab, doxorubicin, and dacarbazine for advanced-stage classical Hodgkin lymphoma

Hun Ju Lee,<sup>1</sup> Rod Ramchandren,<sup>2</sup> Judah Friedman,<sup>3</sup> Jason Melear,<sup>4</sup> Ian W. Flinn,<sup>5</sup> John M. Burke,<sup>4</sup> Yuliya Linhares,<sup>6</sup> Paul Gonzales,<sup>7</sup> Matthew Peterson,<sup>8</sup> Mihir Raval,<sup>4</sup> Rangaswamy Chintapatla,<sup>9</sup> Tatyana A. Feldman,<sup>10</sup> Habte Yimer,<sup>4</sup> Miguel Islas-Ohlmayer,<sup>4,11</sup> Ameet Patel,<sup>11</sup> Leland Metheny,<sup>12</sup> Asad Dean,<sup>4</sup> Vishal Rana,<sup>13</sup> Mitul D. Gandhi,<sup>4</sup> John Renshaw,<sup>4</sup> Linda Ho,<sup>14</sup> Michelle A. Fanale,<sup>14</sup> Wenchuan Guo,<sup>14</sup> and Christopher A. Yasechak<sup>15</sup>

## Effectiveness and Safety of Brentuximab Vedotin in Combination with Nivolumab, Doxorubicin, and Dacarbazine in Patients with Untreated Early-Stage Bulky and Advanced Stage Classical Hodgkin Lymphoma (cHL)

### Context of Research

- The combination of brentuximab vedotin with doxorubicin, vinblastine, and dacarbazine improved overall survival in patients with classical Hodgkin lymphoma (CHL)
- We hypothesized that adding the checkpoint inhibitor nivolumab and removing vinblastine (**AN+AD**) may improve both efficacy and safety

## Patients and Methods

- SGN35-027 (NCT03646123) was a phase 2, open-label, multi-part, multicenter study

Up to 6 cycles  
of **AN+AD**



- Stage I/II CHL with bulky mediastinal disease (>10 cm) or stage III/IV CHL
- Measurable disease per PET/CT imaging

**Primary end point**

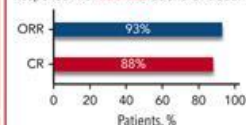
- Complete response (CR) at end of treatment
- Secondary end points**
  - Safety
  - Overall response rate (ORR)
  - Duration of response
  - Duration of CR
  - Event-free survival
  - Progression-free survival
  - Overall survival

**Secondary end points**

- Safety
- Overall response rate (ORR)
- Duration of response
- Duration of CR
- Event-free survival
- Progression-free survival
- Overall survival

### Main Findings

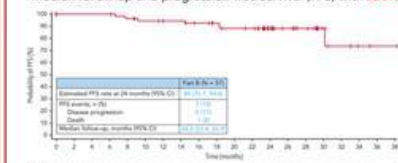
Response with **AN+AD** at end of treatment



No patients experienced febrile neutropenia.




Median follow-up and progression-free survival (PFS) with **AN+AD**



**Conclusions:** With a median follow-up of 24.2 months, the 2-year PFS was 88% in patients treated with AN+AD. ORR and CR rates at the end of treatment were 93% and 88%, respectively. AN+AD continues to show promising efficacy with a favorable safety profile in patients with early-stage bulky and advanced-stage CHL.

Lee et al. DOI: 10.1182/**blood**.2024024681

# OUTREACH: phase 2 study of lisocabtagene maraleucel as outpatient or inpatient treatment at community sites for R/R LBCL

 Clinical Trials & Observations

Yuliya Linhares, Cesar O. Freytes, Mohamad Cherry, Carlos Bachier, Michael Maris, Daanish Hoda, Juan C. Varela, Courtney Bellomo, Scott Cross, James Essell, Suzanne Fanning, Howard Terebello, Habte Yimer, Jay Courtright, Jeff P. Sharman, Ana Kostic, Min Vedal, Ken Ogasawara, Ariel Avilion, Ricardo Espinola, Brenda Yuan, Bassam Mattar

Outpatient CART administration and monitoring were safe and effective at MCI

**OUTREACH: Phase 2 study of lisocabtagene maraleucel as outpatient or inpatient treatment at community sites for R/R LBCL**  
(Linhares et al)

➤ **Primary endpoint:** Grade  $\geq 3$  TEAEs of special interest (CRS, NEs, infections, and prolonged cytopenia [grade  $\geq 3$  at day 29])

➤ **Key secondary endpoints:** Type, frequency, and severity of all AEs/lab abnormalities; ORR, CR rate, DOR, DOR in pts with CR, PFS, and OS

## Open-label, multicenter, phase 2 study at community centers

- Outpatient vs inpatient monitoring was per investigator discretion
- Patients monitored as outpatients received liso-cel in the outpatient facility or in the inpatient facility with same-day discharge

## Key site/patient requirements for outpatient monitoring

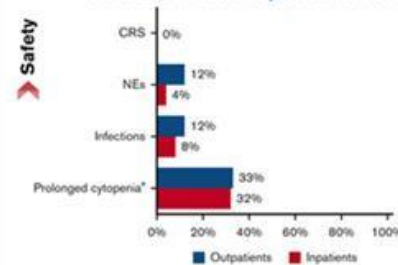
Liso-cel-treated, N = 82

Outpatient monitored, n = 57

Inpatient monitored, n = 25

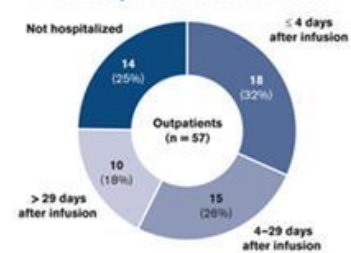
- Site has HSCT or phase 1 capabilities
- One designated hospital with 24/7 CAR T cell AE management and multidisciplinary team
- Patient/caregiver must stay < 1 hour of site x 30 days

## Grade $\geq 3$ TEAEs of special interest



\*Defined as grade  $\geq 3$  laboratory abnormalities of anemia, neutropenia, or thrombocytopenia at day 29.

## Initial hospitalization after liso-cel

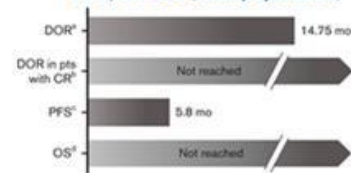


## ORR (total population)



Median on-study follow-up: 10.6 months

## Median DOR, DOR in pts with CR, PFS, and OS (total population)

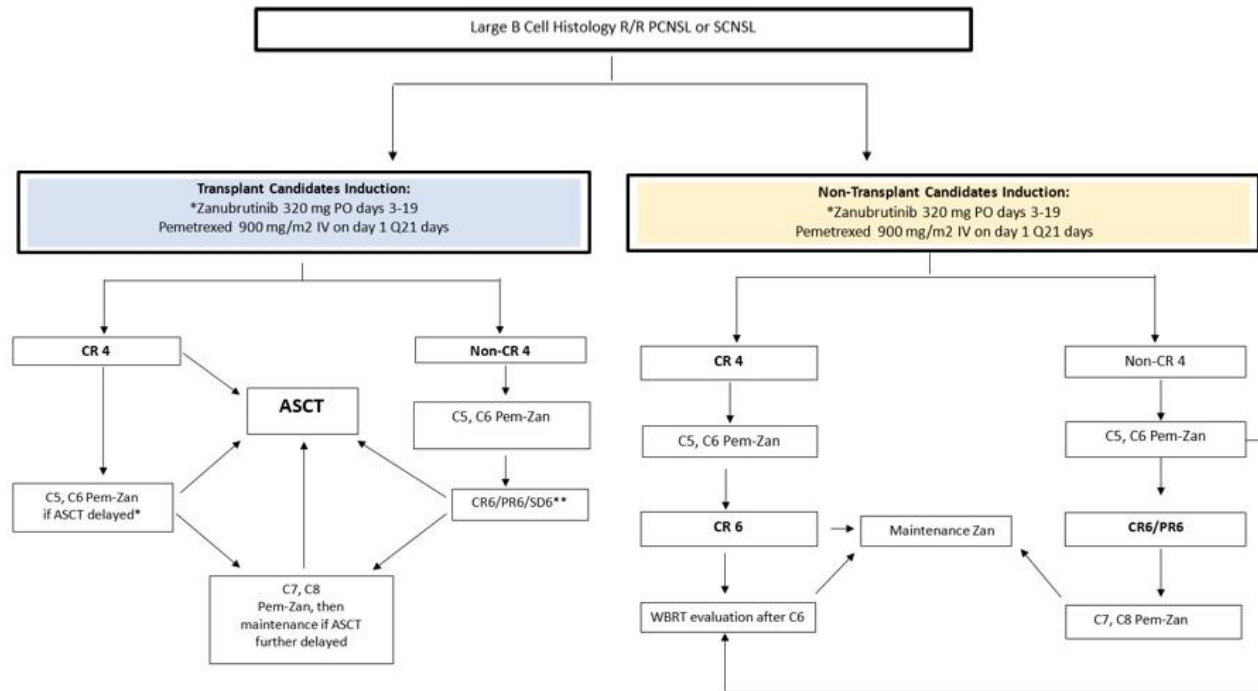


\*Median follow-up: 17.35 mo; \*Median follow-up: 17.7 mo; \*Median follow-up: 18.2 mo; \*Median follow-up: 22.0 mo.

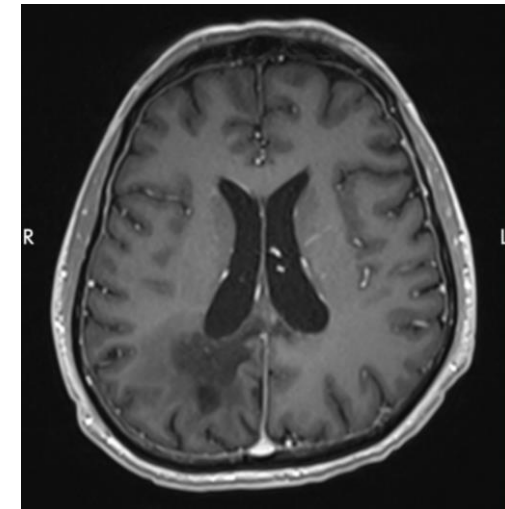
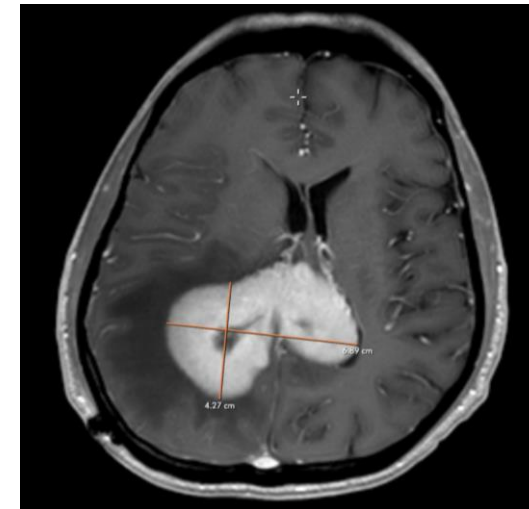


# Trial in Progress: Zanubrutinib with Pemetrexed for the Treatment of Relapsed/Refractory Primary and Secondary CNS Lymphomas: A Phase II Trial with a Safety Lead-in (P.I. Yuliya Linhares, MD; NCT05681195)

## Primary endpoint: best ORR



60 yo male with inadequate response to 2 cycles of standard therapy (high dose methotrexate) and progressive neurologic symptoms



Initial: 4.3 x 6.9 cm  
after 2 high dose methotrexates CR after 4 cycles Pem-Zan

- Fully outpatient regimen
- 2 patients enrolled and treated; both in CR, no significant toxicities
- Patient 1 underwent autologous stem cell transplant and continues to be in remission 5 months after transplant
- Complete resolution of all neurologic deficits in patient 1, patient 2 signed up for a gym

# HOW IS THIS TRIAL UNIQUE??

- Fully outpatient regimen, allows for quality of life and rehabilitation
- Reasonable and realistic low cost drug combination (as opposed to TEDDi-R, VIPOR)
- Real world patient population
- Accepts patients with low performance status, HIV, hepatitis and advanced age
- Allows for consent by proxy for those unable to consent due to neurological deficits
- No significant toxicities
- We analyze tumor tissue mutational profiles and microenvironment and correlate with treatment outcomes (BostonGene Tumor Portrait)
- We study the significance of circulating tumor DNA presence and quantity in blood and CSF (Adaptive-ClonoSeq, PhasEDSeq-Foresight)



# OUR TRIAL IS NEEDED



## Erin M Dunbar, MD

Neuro-Oncology, Medical Oncology

## Piedmont Physicians Neuro Oncology Atlanta

2001 Peachtree Road Northeast, Suite 645  
Atlanta, GA 30309



See Location



See on Map

Office Number

404-605-2050

Fax Number

833-605-2563



4.9 out of 5

1,599 ratings, 446  
comments

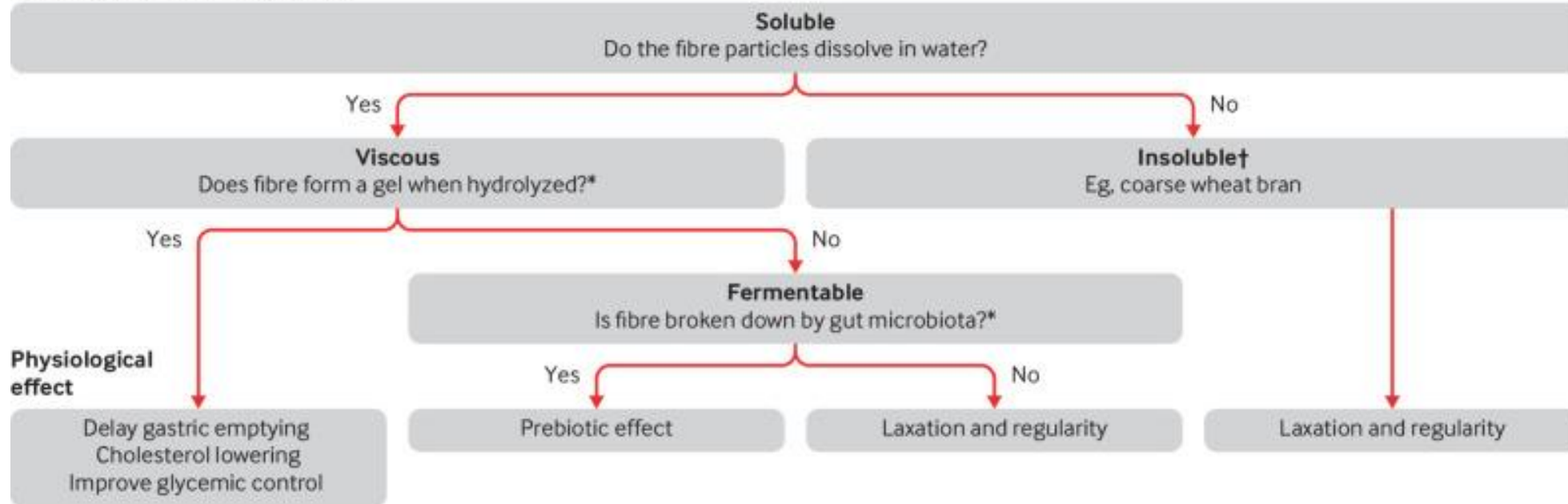
The Piedmont Brain Tumor Center provides a medical "home for life" for patients through multidisciplinary: clinical trials, specialty clinics, tumor boards, and hospital rounds. We are a not-for-profit hospital and rely on donations, volunteers, and resources to support the continual needs of our patients and caregivers.

# Additional Resources

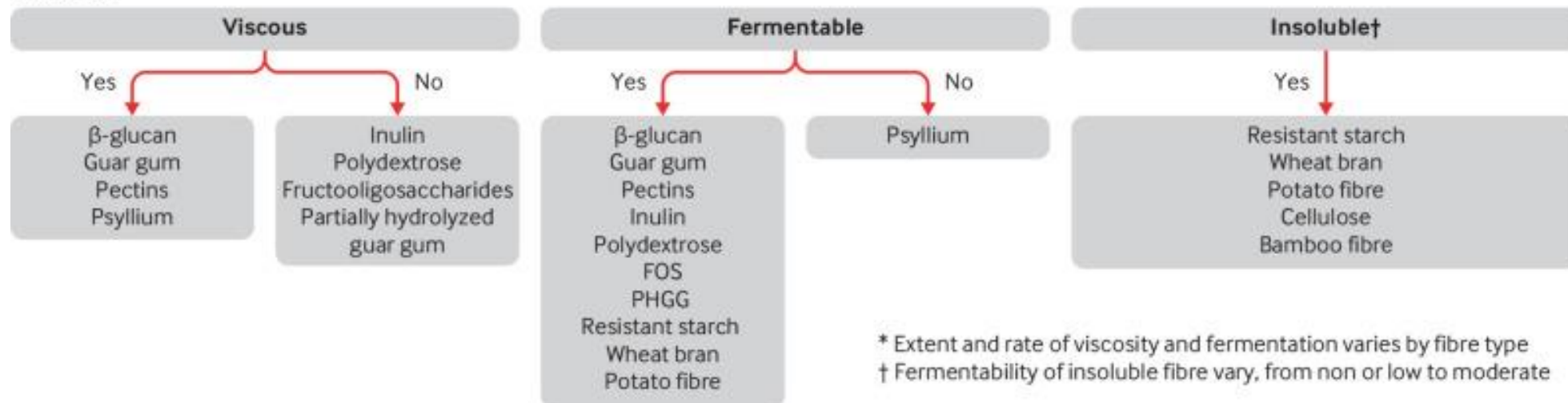


# NOT ALL FIBER IS CREATED EQUAL

## Physiochemical characteristics



## Examples



\* Extent and rate of viscosity and fermentation varies by fibre type

† Fermentability of insoluble fibre vary, from non or low to moderate

# Resources- longevity calculator, references

Lifespan calculator- current and if you make certain dietary changes- great resource to design healthier diet for longevity:

[Food4healthylife \(shinyapps.io\)](https://shinyapps.io/food4healthylife/)

Food for healthy life is based on the following paper: Fadnes LT, Økland J-M, Haaland ØA, Johansson KA (2022) Estimating impact of food choices on life expectancy: A modeling study. PLoS Med 19(1): e1003889.

Dietary Reference Intake Recommendations:

[DRI Calculator Results | National Agricultural Library \(usda.gov\)](https://www.nal.usda.gov/fnic/food-comp/dri-calculator/)

Which nutrients are you missing and nutritional information:

[Nutrient Optimiser](https://nutrientoptimiser.com/)

[Nutrition facts for buckwheat, recommended daily values and analysis.  
\(nutrientoptimiser.com\)](https://nutrientoptimiser.com/nutrition-facts-for-buckwheat-recommended-daily-values-and-analysis/)



# Resources- plant-based diet

**Excellent website to learn about healthy plant-based diet:**

fullplateliving.org

[Full Plate Living](https://fullplateliving.org)

Free “Food as Medicine” course:

[Food as Medicine - American College of Lifestyle Medicine](https://lifestylemedicine.org/nutrition-as-medicine/?gad_source=1&gclid=EALalQobChMIpdOcipyphQMV0LZaBR1zpQOfEAAYASAAEgKQAPD_BwE)

[https://lifestylemedicine.org/nutrition-as-medicine/?gad\\_source=1&gclid=EALalQobChMIpdOcipyphQMV0LZaBR1zpQOfEAAYASAAEgKQAPD\\_BwE](https://lifestylemedicine.org/nutrition-as-medicine/?gad_source=1&gclid=EALalQobChMIpdOcipyphQMV0LZaBR1zpQOfEAAYASAAEgKQAPD_BwE)

**Superfood list from American Academy of Lifestyle Medicine-PDF:**

[Superfood-List.pdf \(lifestylemedicine.org\)](https://lifestylemedicine.org/wp-content/uploads/2022/07/Superfood-List.pdf)

<https://lifestylemedicine.org/wp-content/uploads/2022/07/Superfood-List.pdf>

**Food as medicine curriculum –PDF:**

[ACLM-Food-As-Medicine-Jumpstart-8.5x11.pdf \(lifestylemedicine.org\)](https://lifestylemedicine.org/wp-content/uploads/2024/01/ACLM-Food-As-Medicine-Jumpstart-8.5x11.pdf)

<https://lifestylemedicine.org/wp-content/uploads/2024/01/ACLM-Food-As-Medicine-Jumpstart-8.5x11.pdf>

**Dr. Greger Books: incredible resources for everyone, highly recommend, buy on Amazon**

**How Not To Age   How Not To Die   How Not To Diet**

**Harvard Health Website- excellent resource for healthy living:**

[Signup for More Free Email Newsletters - Harvard Health](https://www.health.harvard.edu/healthbeat/co-reg)

<https://www.health.harvard.edu/healthbeat/co-reg>

## COMPLETE PROTEIN SOURCES

(g = grams of protein / per):

- Quinoa, cooked (8g /1 cup)
- Tofu, cooked (8-10g /100g)
- Tempeh, cooked (18g /100g)
- Buckwheat, raw groats (23g /100g)
- Rice & beans, cooked (10-15g /1 cup)
- Soybeans, raw (36g /100g)
- Hemp seeds (11g /30g)
- Chia seeds (4g /2 tablespoon)
- Spirulina (4g /1 tablespoon)

### Incomplete Protein Sources:

- Grains (approx. 5g /100g cooked)
- Nuts and seeds (average 6-9g /30g)
- Legumes/beans (average 7-9g /100g)
- Vegetables (Green Peas = 7g /1 cup. Spinach & Broccoli = 3-4g /1 cup)
- Nutritional Yeast (4g /1 tablespoon)

## COMBINATION IDEAS TO CREATE COMPLETE PROTEINS INCLUDE:

- Brown Rice and Beans
- Ezekiel Bread (Gluten- this bread is easier on the digestive track because the grains are sprouted.)
- A smoothie with Spirulina (Blue-Green Algae) and nuts
- Salads with any combination beans, nuts or grain
- Soup with any combinations beans or grains
- Get creative creating bowls of all types of grains, vegetables, beans or hummus, with nuts and hempseeds.
- Adding any item from the first list to any meal completes the protein.

Liquid Amino Acids can be used as a supplement to add you meal as well to be sure to get all the essential amino acids. It's flavor is similar to that of soy sauce.

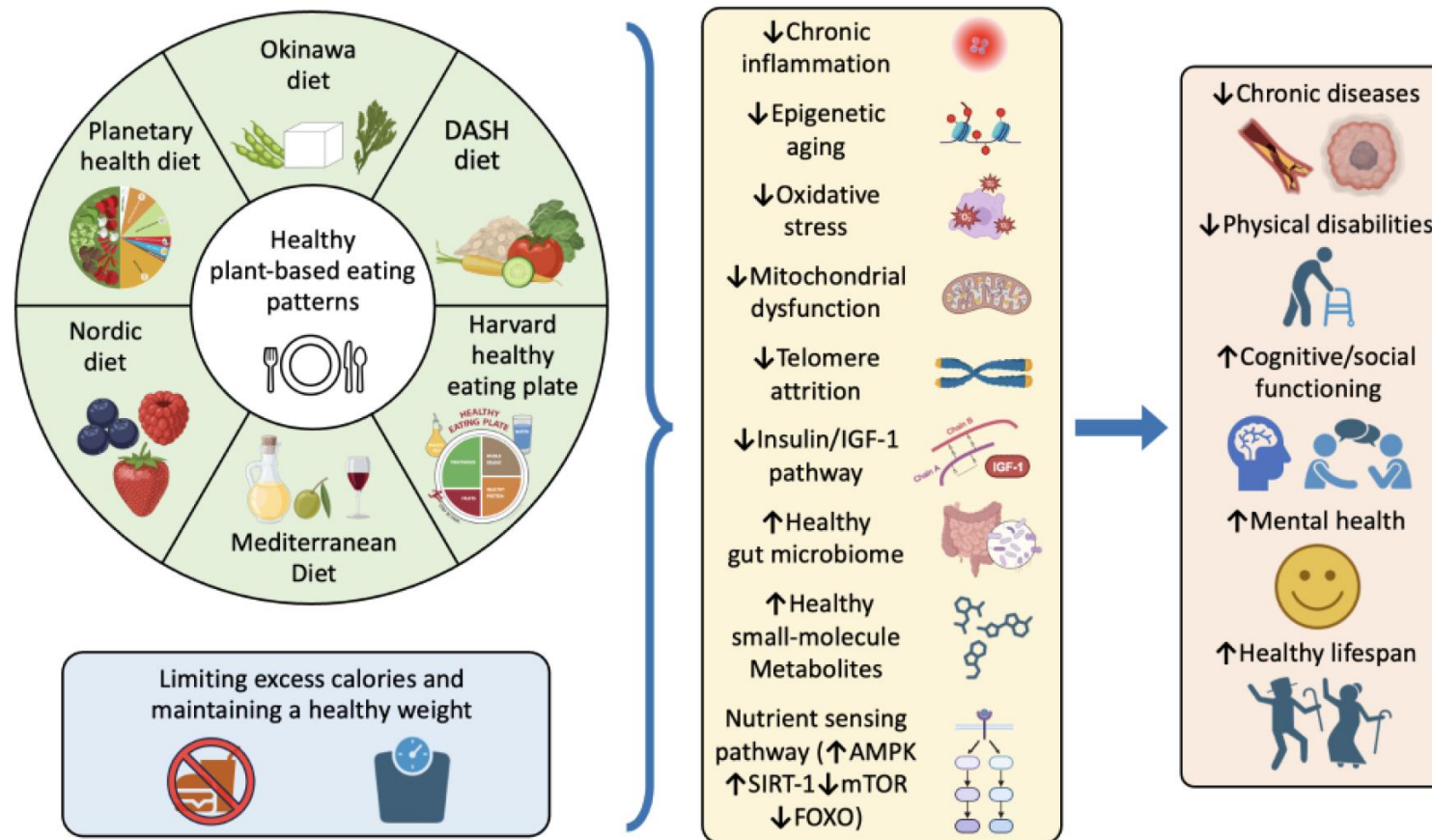


## Grain Nutrient Content Comparison- there is no reason to be eating white rice

In 100g	White rice	Black Rice (forbidden rice)	Pearl Barley	Buckwheat	Quinoa
Whole grain	No- no bran or endosperm	yes	No- bran is off but endosperm is preserved; Hulled barley is full grain	yes	yes
Calories per 100g	353	336	330	350	354
Fiber (g% daily requirement)	3g (10%)	4g	16g (55%)	10g/35%	7g/25%
Carbohydrates per 100 g (g/% daily requirement)	82g (28%)	71g	78g/26%	72g/24%	64g/22%
Protein	7g	10.5g	10g	13 g	14g
Essential amino acids				All 9 present	All 9
Iron	1.6 mg (8%)	1.5 mg	2.5 mg/13%	2.2 mg/12%	4.6mg/25%
Potassium	77 mg (2%)	0 mg	280mg/7%	460 mg/12%	563mg/14%
Special qualities	Incomplete protein	High in anthocyanins (antioxidants)	Rich in soluble fiber known as beta glucan, which is recognized for its cholesterol-lowering abilities	Complete protein	Complete protein

<https://nutrientoptimiser.com/>; table by Dr. Yuliya Linhares

## Diet strategies for promoting healthy aging and longevity



Hu FB J of Internal Medicine 2023

<https://onlinelibrary.wiley.com/doi/full/10.1111/joim.13728>



# PROTEIN MYTH BUSTERS

**MYTH:** You need to eat animal protein to meet your protein needs.

**FACT:** Plants foods such as beans, lentils, nuts, whole grains, and veggies provide ample protein, as well as fiber and other essential vitamins, minerals, and phytochemicals not found in animal products such as meat, fish, poultry, eggs, and dairy.

## PLANT PROTEIN

per serving

### ADVANTAGES

- Fiber
- Phytonutrients
- Vitamins & minerals
- Low or healthy fat profile
- No cholesterol

**18g**  
Red Lentils  
boiled, 1 cup



**17g**  
Edamame  
boiled, 1 cup



**15g**  
Black Beans  
cooked, 1 cup



**6g**  
Almonds  
1 oz



**5g**  
Peas  
cooked, 1 cup



**5g**  
Baked Potato  
1 medium



**5g**  
Spinach  
boiled, 1 cup



## ANIMAL PROTEIN

per serving

### DISADVANTAGES

- Cholesterol
- Saturated fat
- No fiber
- Higher in calories

**6g**  
Egg  
cooked, 1



**20g**  
Salmon  
cooked, 3 oz



**25g**  
Steak  
cooked, 3 oz



**25g**  
Chicken  
cooked, 3 oz



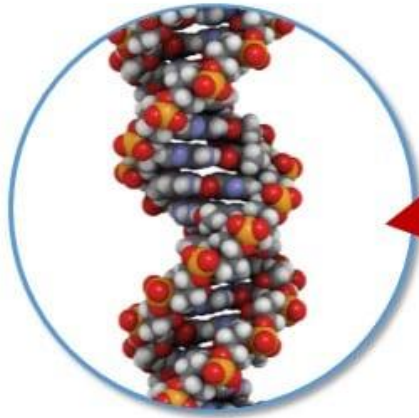
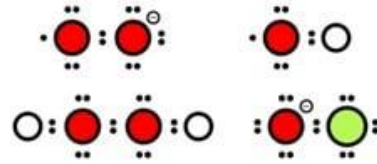
Eating minimally processed whole plant foods such as vegetables, fruits, whole-grains, legumes, and nuts lower the risk of diabetes, heart disease, cancer, and promote overall health.

Whole Grains	Fruits	Vegetables	Beans/Legumes	Nuts & Seeds
Oatmeal, cooked 1 cup <b>4.2 g fiber</b>	Avocado ½ fruit 4.6 g fiber	Spinach, raw 2 cups 1.3 g fiber	Chickpeas ½ cup <b>5.2 g fiber</b>	Flaxseed 1 tbsp. 2.9 g fiber
Quinoa, cooked ½ cup <b>2.2 g fiber</b>	Apple (with skin) 1 medium 4.4 g fiber	Cubed Sweet Potatoes (raw) 1 cup 4.0 g fiber	Lima beans ½ cup <b>4.5 g fiber</b>	Chia seeds 1 tbsp. 3.4 g fiber
Popcorn, plain 3 cups <b>3.5 g fiber</b>	Pear (w skin) 1 medium 5.6 g fiber	Broccoli, cooked 1 cup 5.2 g fiber	Black beans ½ cup <b>9 g fiber</b>	Almonds, whole ¼ cup (23 nuts) 3.8 g fiber
Barley, cooked ½ cup <b>4.3 g fiber</b>	Raspberries 1 cup 8.0 g fiber	Green peas 1/2 cup 3.6 g fiber	Soybean ½ cup <b>5.1 g fiber</b>	Walnuts ¼ cup 2.0 g fiber
Whole wheat spaghetti, cooked, ½ cup <b>2.7 g fiber</b>	Banana 1 medium 3.1 g fiber	Butternut squash, raw 1 cup cubed 3.9 g fiber	Kidney bean ½ cup <b>4.9 g fiber</b>	Peanut butter (chunky) 1 tbsp. 1.0 g fiber



Whole Grains	Fruits	Vegetables	Beans/Legumes	Nuts & Seeds
Oatmeal, cooked 234g 4.2 g fiber	Avocado ½ fruit (68g) 4.6 g fiber	Spinach, raw 60g 1.3 g fiber	Chickpeas 85g 5.2 g fiber	Flaxseed 10g 2.9 g fiber
Quinoa, cooked 194g 2.2 g fiber	Apple (with skin) 1 medium (182g) 4.4 g fiber	Cubed Sweet Potatoes (raw) 133g 4.0 g fiber	Lima beans 85g 4.5 g fiber	Chia seeds 10g 3.4 g fiber
Popcorn, plain 93g 3.5 g fiber	Pear (w skin) 1 medium (178g) 5.6 g fiber	Broccoli, cooked 156g 5.2 g fiber	Black beans 85g 9 g fiber	Almonds, whole 35g (23 nuts) 3.8 g fiber
Barley, cooked 24g 4.3 g fiber	Raspberries 1 cup (123g) 8.0 g fiber	Green peas 123g 3.6 g fiber	Soybean 85g 5.1 g fiber	Walnuts ¼ cup 30g 2.0 g fiber
Whole wheat spaghetti, cooked, 78g 2.7 g fiber	Banana 1 medium (118g) 3.1 g fiber	Butternut squash, raw, cubed 118g 3.9 g fiber	Kidney bean 85g 4.9g fiber	Peanut butter (chunky) 16g 1.0 g fiber

## ROS and RNS Production



### Nucleic Acid Oxidation

Oxidative modifications in DNA/RNA cause nucleotide alterations, strand breaks, base conversions, and the formation of adducts. Markers include: 8-OHdG, 8-oxoG, isoguanine, 8-oxoadenine, 5-hydroxycytosine, 5-chlorocytosine, and 5-chlorouracil



### Protein Oxidation

Reactive species cause modifications in amino acid residues, protein backbones and functional groups that affect protein structure. Markers include: protein carbonyls, nitrotyrosine, oxLDL, ischemia-modified albumin, 3-chlorotyrosine, and AGEs



### Lipid Peroxidation

The reaction of ROS with polyunsaturated fatty acids produces lipid hydroperoxides and reactive aldehyde end products. Markers include: malondialdehyde, 4-hydroxynonenal, F2-isoprostanes, isolevuglandins, and acrolein



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- Buckwheat, raw groats (23g /100g)
- Rice & beans, cooked (10-15g /1 cup)
- Soybeans, raw (36g /100g)
- Hemp seeds (11g /30g)
- Chia seeds (4g /2 tablespoon)
- Spirulina (4g /1 tablespoon)

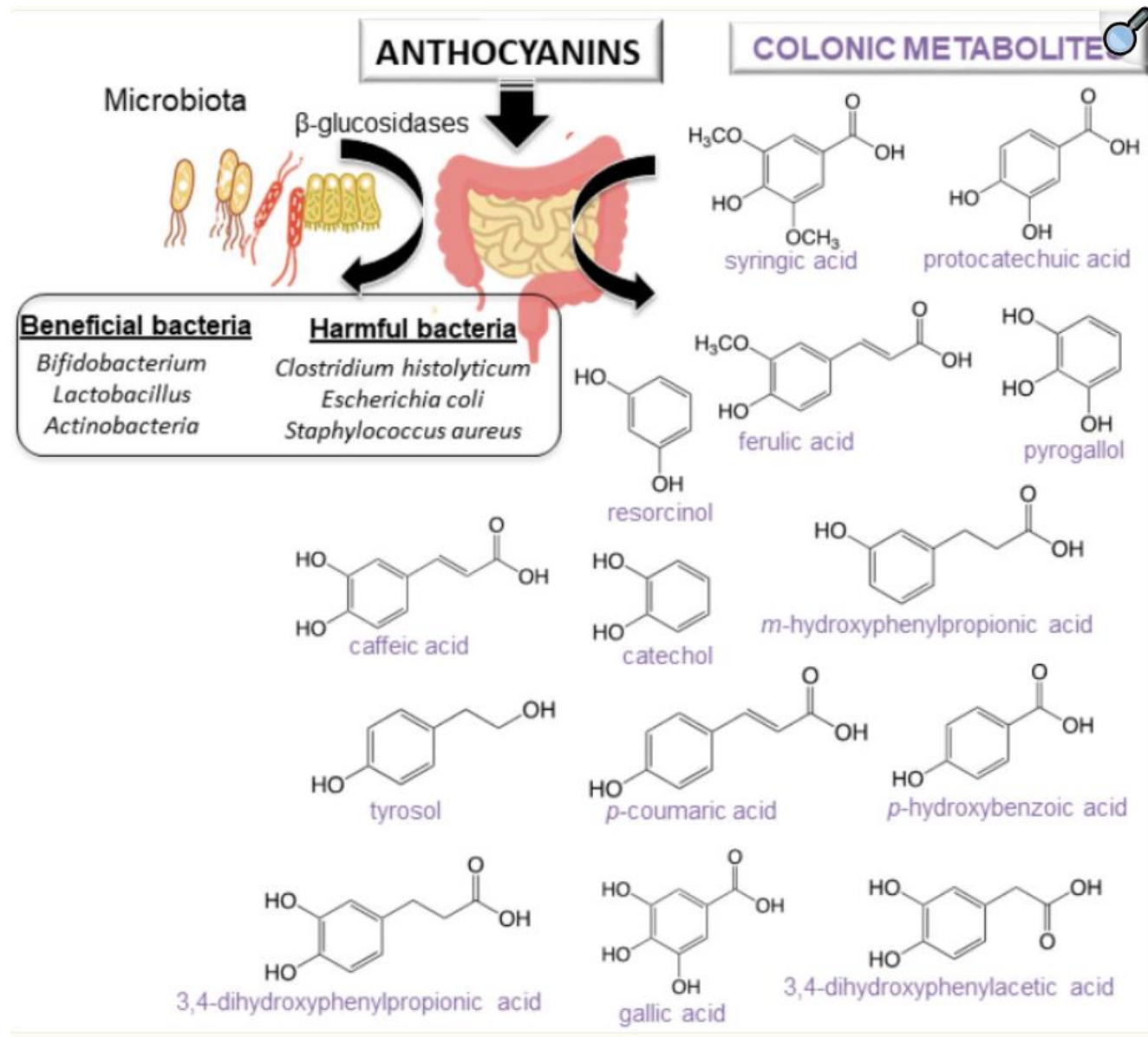
### Incomplete Protein Sources:

- Grains (approx. 5g /100g cooked)
- Nuts and seeds (average 6-9g /30g)
- Legumes/beans (average 7-9g /100g)
- Vegetables (Green Peas = 7g /1 cup. Spinach & Broccoli = 3-4g /1 cup)
- Nutritional Yeast (4g /1 tablespoon)

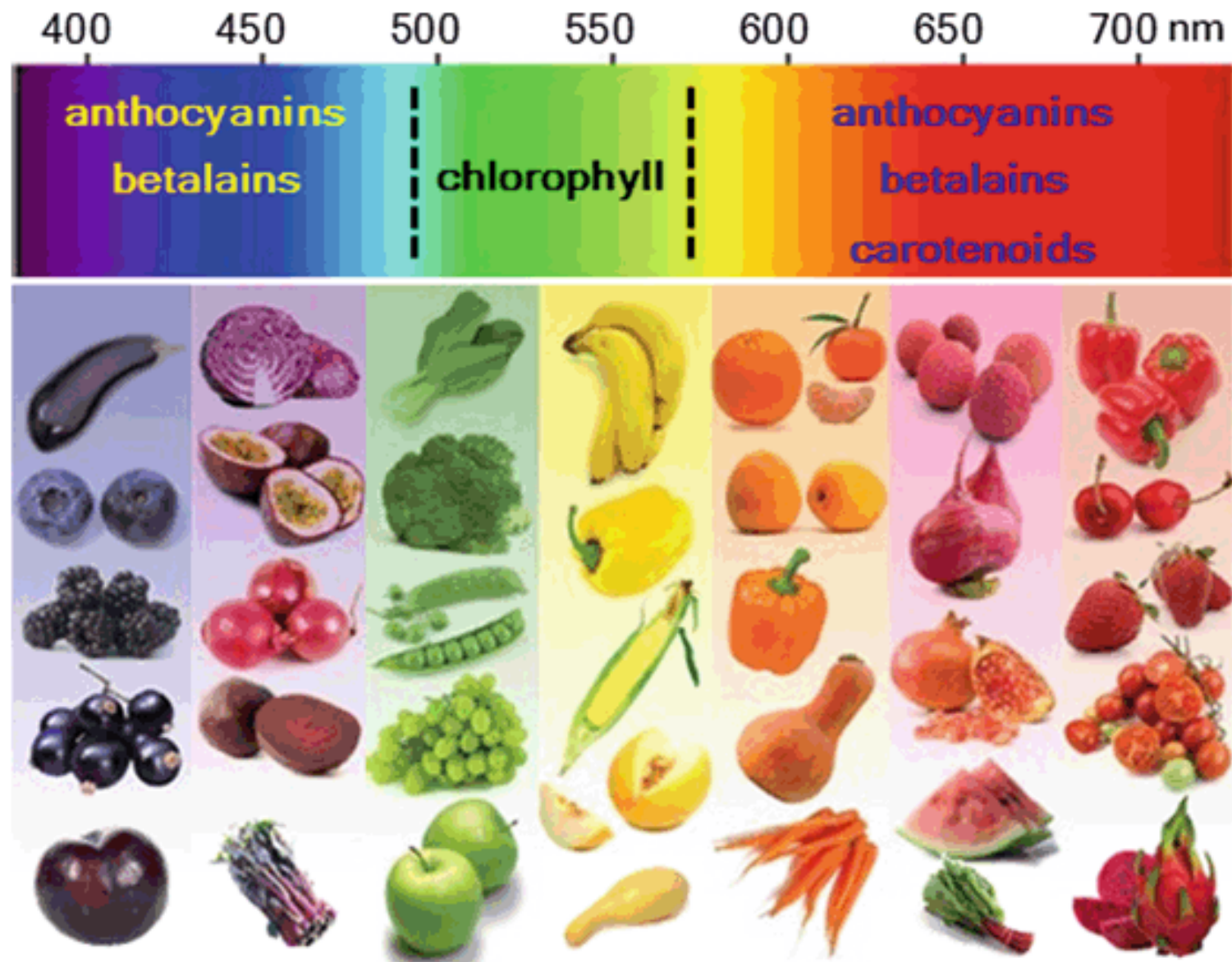
## COMBINATION IDEAS TO CREATE COMPLETE PROTEINS INCLUDE:

- Brown Rice and Beans
- Ezekiel Bread (Gluten- this bread is easier on the digestive track because the grains are sprouted.)
- A smoothie with Spirulina (Blue-Green Algae) and nuts
- Salads with any combination beans, nuts or grain
- Soup with any combinations beans or grains
- Get creative creating bowls of all types of grains, vegetables, beans or hummus, with nuts and hempseeds.
- Adding any item from the first list to any meal completes the protein.

Liquid Amino Acids can be used as a supplement to add you meal as well to be sure to get all the essential amino acids. It's flavor is similar to that of soy sauce.







## ADDDITIONAL REFERENCES

2. Sui J, Guo J, Pan D, et al. The Efficacy of Dietary Intake, Supplementation, and Blood Concentrations of Carotenoids in Cancer Prevention: Insights From an Umbrella Meta-Analysis. This umbrella meta-analysis found that dietary intake and blood concentrations of carotenoids are inversely associated with the risk of various cancers, although high doses of  $\beta$ -carotene supplements may increase the risk of lung and bladder cancers.[2]
7. Saini RK, Keum YS, Daglia M, Rengasamy KR. Dietary Carotenoids in Cancer Chemoprevention and Chemotherapy: A Review of Emerging Evidence. This review highlights the potential roles of various carotenoids in cancer chemoprevention and chemotherapy.[7]
11. Key TJ, Appleby PN, Travis RC, et al. Carotenoids, Retinol, Tocopherols, and Prostate Cancer Risk: Pooled Analysis of 15 Studies. This pooled analysis found that lycopene and  $\alpha$ -tocopherol are inversely associated with aggressive prostate cancer risk.[11]

# TYPES OF CAROTENOIDS

## Beta-Carotene ( $\beta$ -carotene) and A-Carotene ( $\alpha$ -Carotene)

- Closely related, as both are synthesized to form active A
  - Both found in foods like squash, spinach, sweet potatoes and carrots
  - Have anti-inflammatory, cancer-protective effects
- estrogen receptor positive breast cancer



## Lutein and Zeaxanthin

- The only two found in the retina and lens of the human eye
- Improve eye health and protect vision
- Best sources are dark leafy green vegetables and cruciferous veggies
- Lower age-related eye problems including macular degeneration and cataracts



## Lycopene

- Best source is tomato, especially cooked tomatoes
- Reduce risk for developing diseases like prostate cancer.





# The Second World Cancer Research Fund/American Institute for Cancer Research Expert Report. *Food, Nutrition, Physical Activity, and the Prevention of Cancer: A Global Perspective*

**Table 1.** Headline recommendations of the expert Panel convened to discuss and judge the evidence from the systematic literature reviews, to draw conclusions and to make recommendations

Category	Recommendation
Body fatness	Be as lean as possible within the normal range of body weight
Physical activity	Be physically active as part of everyday life
Foods and drinks that promote weight gain	Limit consumption of energy-dense foods
Plant foods	Avoid sugary drinks
Animal foods	Eat mostly foods of plant origin
Alcoholic drinks	Limit intake of red meat and avoid processed meat
Preservation, processing, preparation	Limit alcoholic drinks
Dietary supplements	Limit consumption of salt
Breast-feeding	Avoid mouldy cereals or pulses
Cancer survivors	Aim to meet nutritional needs through diet alone
	Mothers to breast-feed; children to be breast-fed
	Follow the recommendations for cancer prevention

Wiseman M. The second World Cancer Research Fund/American Institute for Cancer Research expert report. Food, nutrition, physical activity, and the prevention of cancer: a global perspective. *Proc Nutr Soc.* 2008;67(3):253-256. doi:10.1017/S002966510800712X.

Sustained Minimal Residual Disease Negativity in Multiple Myeloma is Associated with Stool Butyrate and Healthier Plant-Based Diets

Urvi A. Shah<sup>1,2</sup>, Kylee H. MacLachlan<sup>1,2</sup>, Andriy Derkach<sup>3</sup>, Meghan Salcedo<sup>1</sup>, Kelly Barnett<sup>1</sup>, Julia Caple<sup>1</sup>,



Original Investigation | Oncology

Association of Short-Chain Fatty Acids in the Gut Microbiome With Clinical Response to Treatment With Nivolumab or Pembrolizumab in Patients With Solid Cancer Tumors

Motom Nomura, PhD; Bunzoku Nagatomo, MDPhD; Kaitaro Doi, MD; Ikuo Shimizu, MDPhD; Kishiro Baba, MD; Tomoki Saito, MD; Shinomi Matsumoto, PhD;

The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Microbiota as Predictor of Mortality in Allogeneic Hematopoietic-Cell Transplantation

J.U. Peled, A.L.C. Gomes, S.M. Devlin, E.R. Littmann, Y. Taur, A.D. Sung, D. Weber,

RESEARCH PAPER

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Microbial metabolite butyrate promotes anti-PD-1 antitumor efficacy by modulating T cell receptor signaling of cytotoxic CD8 T cell

Xinhai Zhu<sup>a#</sup>, Ke Li<sup>b#</sup>, Guichao Liu<sup>c,d#</sup>, Ruan Wu<sup>e</sup>, Yan Zhang<sup>a</sup>, Siying Wang<sup>f</sup>, Meng Xu<sup>a\*</sup>, Ligong Lu<sup>g\*</sup>, and Peng Li<sup>g,h\*</sup>

- Feed your gut bacteria to enhance butyrate production
- *Bifidobacterium*, *Faecalibacterium*, *Eubacterium*, and *Roseburia*
- fiber -whole grains, legumes, fruits, vegetables, nuts, and seeds, dark chocolate

IMMUNOTHERAPY

Dietary fiber and probiotics influence the gut microbiome and melanoma immunotherapy response

Christine N. Spencer<sup>1†‡</sup>, Jennifer L. McQuade<sup>2†</sup>, Vancheswaran Gopalakrishnan<sup>1†§</sup>,

Zhu X, Li K, Liu G, et al. Microbial metabolite butyrate promotes anti-PD-1 antitumor efficacy by modulating T cell receptor signaling of cytotoxic CD8 T cell. *Gut Microbes*. 2023;15(2):2249143. doi:10.1080/19490976.2023.2249143. Shah, U. A., MacLachlan, K. H., Derkach, A., Salcedo, M., Barnett, K., Caple, J., Blaslov, J., Tran, L., Ciardiello, A., Burge, M., Shekarshand, T., Adinolfi, P., Cross, J., Planko, M. J., Hosse, K., McVoy, D., Mailankody, S., Kord, N., Hultcrantz, M., Hassoun, H., ... Lesokhin, A. M. (2022). Sustained Minimal Residual Disease Negativity in Multiple Myeloma is Associated with Stool Butyrate and Healthier Plant-Based Diets. *Clinical cancer research: an official journal of the American Association for Cancer Research*, 28(23), 5149–5155. <https://doi.org/10.1158/1078-0432.CCR-22-0723> Lee SY, Jhun J, Woo JS, et al. Gut microbiome-derived butyrate inhibits the immunosuppressive factors PD-L1 and IL-10 in tumor-associated macrophages in gastric cancer. *Gut Microbes*. 2024;16(1):2300846. doi:10.1080/19490976.2023.2300846; Nomura M, Nagatomo R, Doi K, et al. Association of Short-Chain Fatty Acids in the Gut Microbiome With Clinical Response to Treatment With Nivolumab or Pembrolizumab in Patients With Solid Cancer Tumors. *JAMA Netw Open*. 2020;3(4):e202895. Published 2020 Apr 1. doi:10.1001/jamanetworkopen.2020.2895; Butyrate in microbiome abates a host of IIS studies find - [UW Medicine | Newsroom](http://www.uwmedicinejournal.com); Peled JU, Gomes AL, Devlin SM, et al. Microbiota as Predictor of Mortality in Allogeneic Hematopoietic-Cell Transplantation. *N Engl J Med*. 2020;382(9):822-834. doi:10.1056/NEJMoa1900623; Spencer CN, McQuade JL, Gopalakrishnan V, et al. Dietary fiber and probiotics influence the gut microbiome and melanoma immunotherapy response. *Science*. 2021;374(6575):1632-1640. doi:10.1126/science.aba7015

