

NUTRITION IN MEDICINE

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DISCLOSURES

- Royalties from general interest books about plant-based nutrition
- Research funding from Highland Hospital Foundation, with philanthropic donations from T. Colin Campbell Center for Nutrition Studies, the Ladybug Foundation, and multiple individuals

CASE

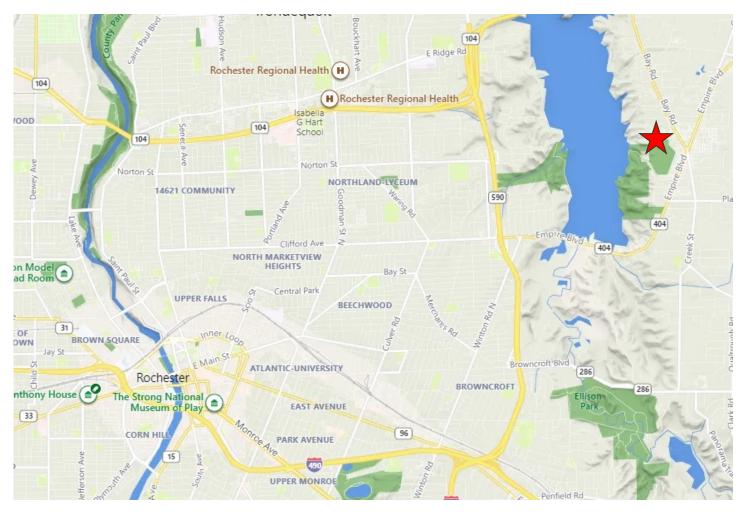
- 72-year old female patient with well controlled diabetes, hyperlipidemia, obesity, and worsening knee OA (starting to mildly limit her physical activity)
- Her slightly older sister was recently diagnosed with breast cancer and she is interested in lowering her cancer risks as much as possible, losing weight, and getting healthier to better enjoy time with her family and to maintain mobility and energy. Motivation is sky high.

Standard Food Fragments American (ex: white flour, oil, sugar) Vegetarian Vegan Low-Carb, Processing Mediterranean High Fat DASH Paleo Whole-food, Plant-Based Whole Foods More Animal Animal/Plant More Plant

DIETARY SPECTRUM

LOCATION











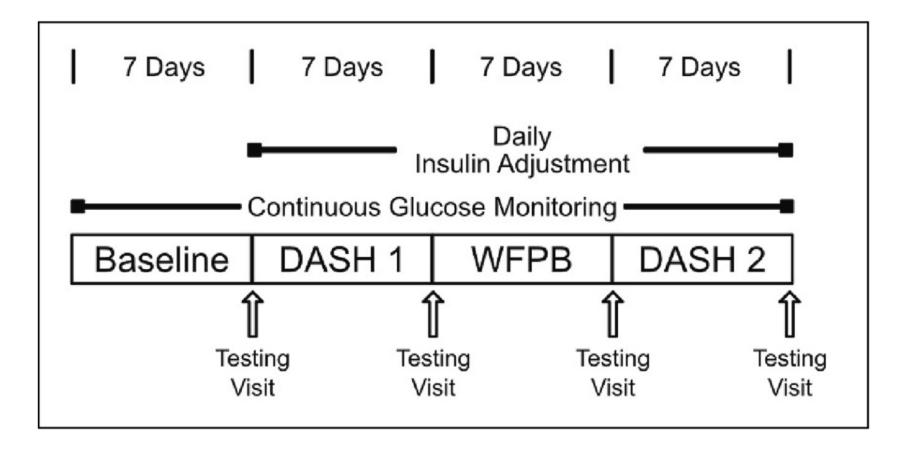


STAFF

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TWO PLANT-BASED DIETS IN TYPE 2 DIABETES

Campbell TM, Campbell EK, Attia J, et al. The Acute Effects of a DASH diet and Whole Food, Plant-Based diet on Insulin Requirements and Related Cardiometabolic Markers in Individuals with Insulin-Treated Type 2 Diabetes. *Diabetes Res Clin Pract.* 2023:110814.



BASELINE CHARACTERISTICS (n=15)

| Age | 56.7 Years |
|-----------------------------------|------------------------|
| BMI | 34.3 Kg/M ² |
| Years Since Diabetes Diagnosis | 12 Years |
| HbA1c | 8.4 % |
| Basal Insulin | 100 % |
| Prandial Insulin | 80 % |
| Total Daily Insulin Dose | 91 Units |





Breakfast:

Steel cut oatmeal with fruit salad, walnuts, flax seed

Lunch:

Turkey sandwich on whole wheat with lettuce, tomato, onion, mustard, and mayo Side Salad



Dinner:

Blackened Salmon Farro salad with spinach, cucumber, tomato, asparagus



Snacks:

Mozzarella stick Whole wheat crackers Yogurt parfait Walnuts

DASH MENU (DAY 1) – 1800 CALORIES

WFPB MENU (DAY 1) – 1800 CALORIES





Breakfast:

2 Banana Flax Muffins with fruit

Lunch:

Sweet Potato Enchiladas Side Salad

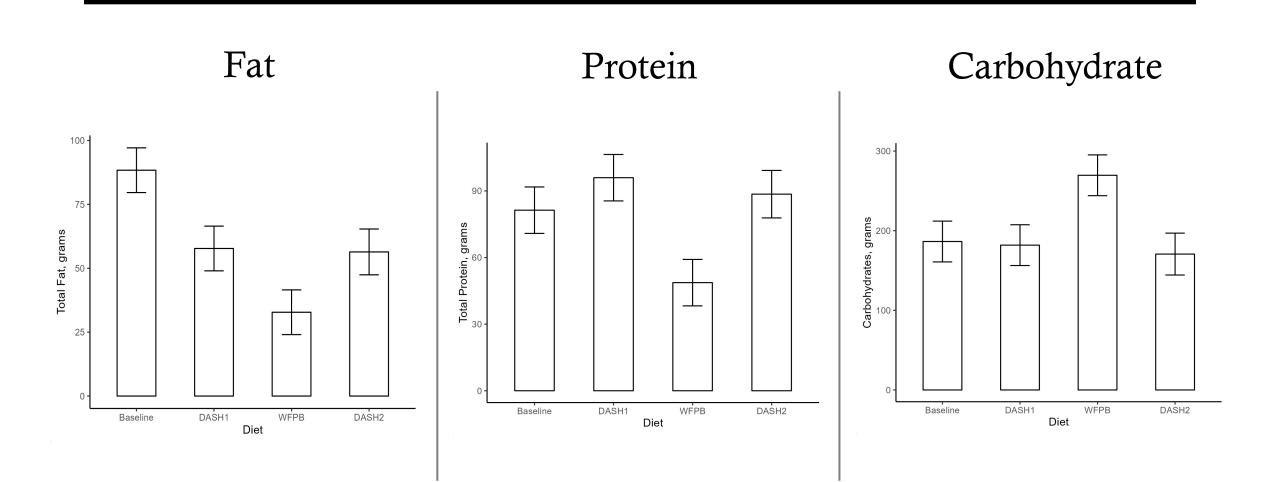
Dinner:

Mediterranean White Bean Soup Side Salad



Snacks:

Whole grain crackers (Mary's Gone Crackers) Hummus Fresh Pineapple Chunks



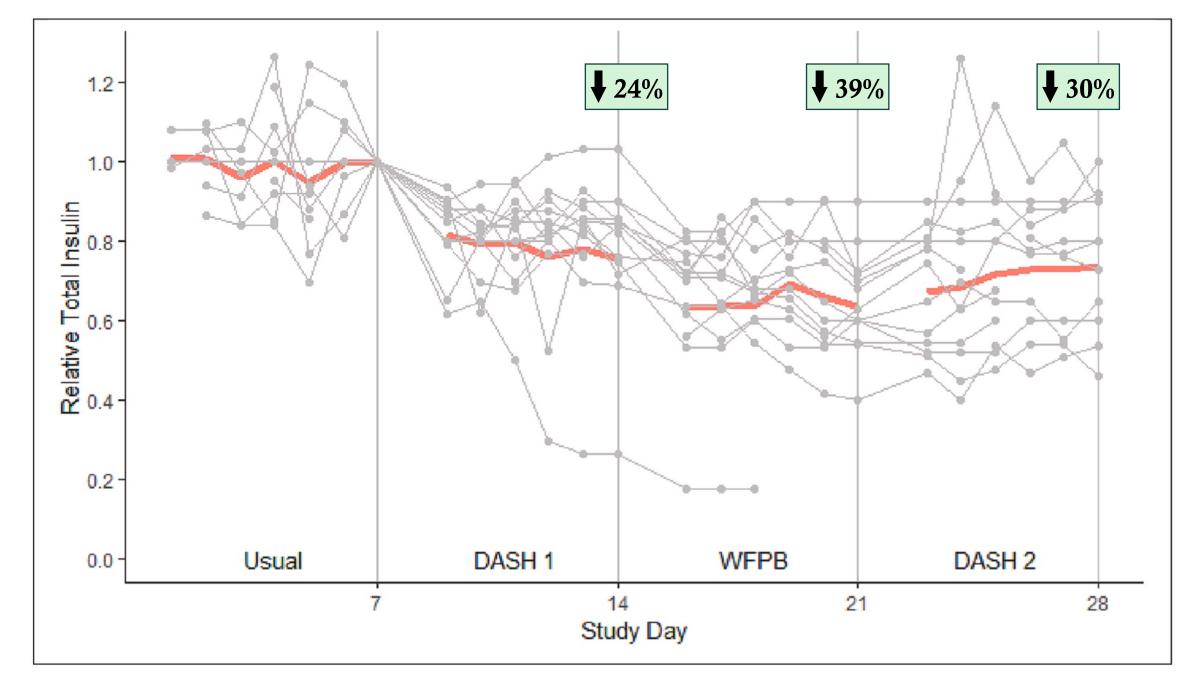
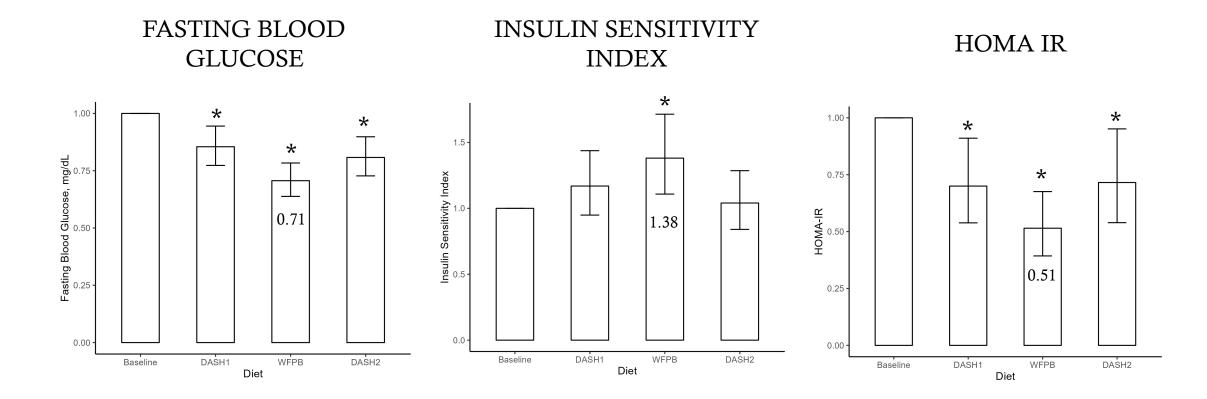


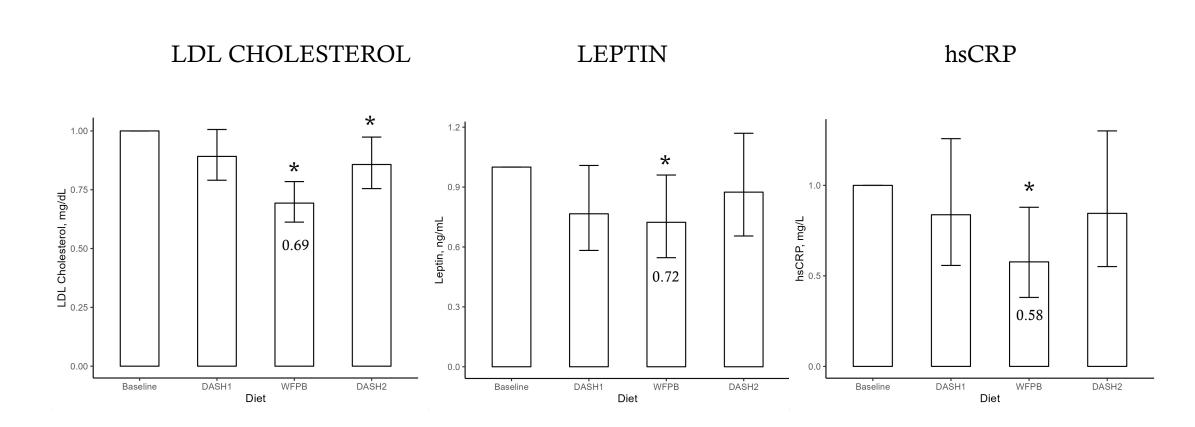
Fig. 4. Relative Daily Insulin Usage During Four Study Phases. Bold line denotes the geometric mean total daily insulin usage relative to the insulin dose at the and of the baseline week. Light gray lines with circles are trajectories of each individual participant. 1st day of each diet is emitted due to non dietary influence of

INSULIN SENSITIVITY



* p<0.05 compared to baseline

OTHER CARDIOMETABOLIC

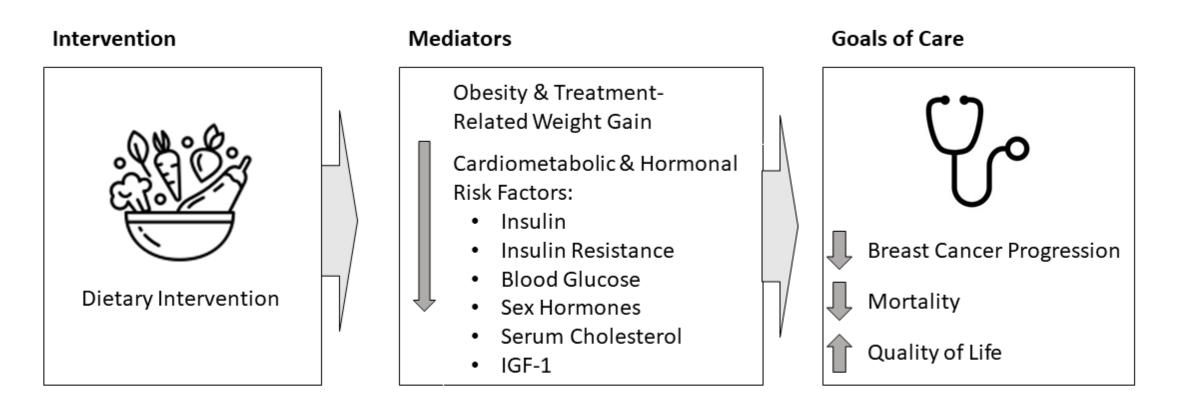


* p<0.05 when compared to baseline

Standard Food Fragments American (ex: white flour, oil, sugar) Vegetarian Vegan Low-Carb, Processing Mediterranean High Fat DASH Paleo Whole-food, Plant-Based Whole Foods More Animal Animal/Plant More Plant

DIETARY SPECTRUM

NOT JUST METABOLIC DISEASE... ...METASTATIC BREAST CANCER



WHOLE-FOOD, PLANT-BASED NUTRITION AMONG WOMEN WITH METASTATIC BREAST CANCER: A PILOT STUDY OF RECRUITMENT, RETENTION, AND PRELIMINARY CHANGES IN BIOMARKERS AND SYMPTOMS

Co-Investigators/Collaborators:

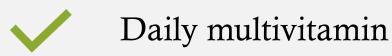
- Erin Campbell, MD (Public Health Sciences)
- Luke Peppone, PhD (Cancer Control)
- Eva Culakova, PhD (Cancer Control)
- James Fetten, MD (Oncology)
- Richard Moore (Gyn Onc)
- Alissa Huston, MD (Oncology)
- Michelle Shayne, MD (Oncology)

INTERVENTION

- 8 weeks of a whole-food, plant-based diet with meals provided.
- Subjects were encouraged to eat as much as necessary to be full, and could add their own food in place of, or in addition to, provided food.
- Weekly visits with TC and/or EC
- Daily Multivitamin
- Testing at weeks 0, 4, 8



CONTROL





Continue usual diet



2 weeks of food/education provided after all testing was complete as an incentive to continue participating.

SAMPLE MEALS

Breakfast: Muesli with ground flaxseed and fresh berries with unsweetened, nondairy milk

Lunch: Ocean Chickpea Sandwich with Baked Sweet Potato Fries **Dinner**: Mushroom Stew with Roasted Green Beans & Onions, Southern Kale

Breakfast: Banana Flax Muffin with 1 1/3 cup fresh fruit **Lunch**: Vegetable Barley Soup with Whole Grain Bread **Dinner**: Peanut Soba Noodles with Roasted Broccoli & Cauliflower

BASELINE CHARACTERISTICS

| Characteristics | Mean ± SD | Control | Intervention |
|--------------------------------------|----------------------------|-----------------|-----------------|
| | | (n=10*) | (n=21) |
| Age (years) | | 64.2 ± 8.9 | 59.1 ± 11.0 |
| Race | Black, n (%) | 0 (0) | 1 (4.8) |
| | White, n (%) | 10 (100.0) | 19 (90.5) |
| | No answer, n (%) | 0 (0) | 1 (4.8) |
| Ethnicity | Not Hispanic/Latino, n (%) | 10 (100.0) | 20 (95.2) |
| | No answer, n (%) | 0 (0) | 1 (4.8) |
| BMI at Study Baseline | Mean ± SD | 28.4 ± 4.4 | 30.2 ± 7.2 |
| (kg/m²) | | | |
| Age at First Breast Cancer Diagnosis | Mean ± SD | 52.9 ± 11.7 | 49.4 ± 10.9 |
| (years) | | | |
| Years Elapsed Since First Diagnosis | Mean ± SD | 11.2 ± 7.9 | 9.7 ± 6.4 |
| Years Elapsed Since Diagnosis of | Mean ± SD | 5.3 ± 6.0 | 2.2 ± 1.8 |
| Metastatic Breast Cancer | | | |
| Hormone Receptor Status | ER+, n (%) | 10 (100.0) | 20 (95.2) |
| | PR+, n (%) | 9 (90.0) | 17 (81.0) |
| | HER2+, n (%) | 3 (30.0) | 6 (28.6) |
| Location of Metastases | Bone, n (%) | 7 (70.0) | . , |
| | Lung, n (%) | 4 (40.0) | · / |
| | Brain, n (%) | 1 (10.0) | , , |
| | Liver, n (%) | 2 (20.0) | . , |
| | Other, n (%) | 6 (60.0) | 7 (33.3) |

WEIGHT (STARTING BMI 29.7)

Lost 1-2 pounds/week (177.5 to 165.7lb)



BREAST CANCER & WEIGHT

WEIGHT (STARTING BMI 29.7)

Lost 1-2 pounds/week (177.5 to 165.7lb)

CARDIOMETABOLIC

Within Group

| LDL Cholesterol | ↓ 21% (p<0.01) |
|---------------------------------|----------------|
| Insulin (fasting) | ↓ 33% (p<0.01) |
| HOMA-IR | ↓ 39% (p=0.01) |
| IGF-1 | ↓ 10% (p=0.01) |
| Sex Hormone Binding Globulin | ↑ 32% (p<0.01) |
| Free Testosterone | ↓ 35% (p=0.08) |

Campbell TM, Campbell EK, Culakova E, et al. A Whole-Food, Plant-Based Randomized Controlled Trial in Metastatic Breast Cancer: Weight, Cardiometabolic, and Hormonal Outcome. *Breast Cancer Research and Treatment.* 2024.

CANCER MARKERS: STABLE IN INTERVENTION GROUP INCREASED IN CONTROL GROUP (NS)

| Outcome | Intervention vs Control at week 8 (adjusted for baseline) | |
|------------------------------|---|---------|
| | Diff. | p-value |
| CA 27.29 (U/mL) ^c | -5.3 | 0.23 |
| CA 15-3 (U/mL) ^c | -5.2 | 0.53 |
| CEA3 (ng/mL) | -0.5 | 0.54 |

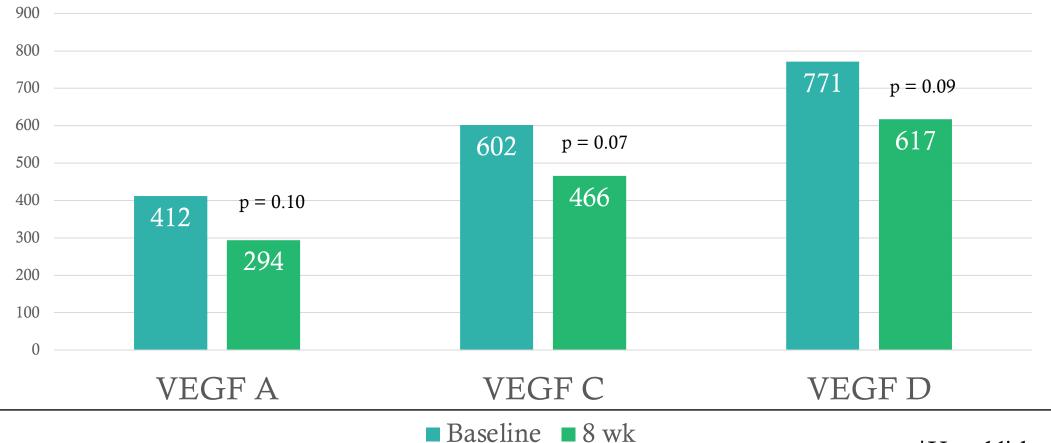
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Vascular Endothelial Growth Factor (VEGF) has 900 potent angio- and lymphangiogenic functions. Tumor 800 sections analyzed for VEGF-A, VEGF-C, and VEGF-D found that high levels of VEGF-A and VEGF-C 600 500 were associated with higher lymph vessel density, 400 microvessel density, lymph node metastasis, distant 300 metastasis, and shorter overall survival. 200 100

Mohammed, et al. "Prognostic significance of vascular endothelial cell growth factors –A, -C, and –D in breast cancer and their relationship with angio- and lymphangiogenesis." *British Journal of Cancer.* 2007

■ Baseline ■ 8 wk

Vascular Endothelial Growth Factor (within intervention group)



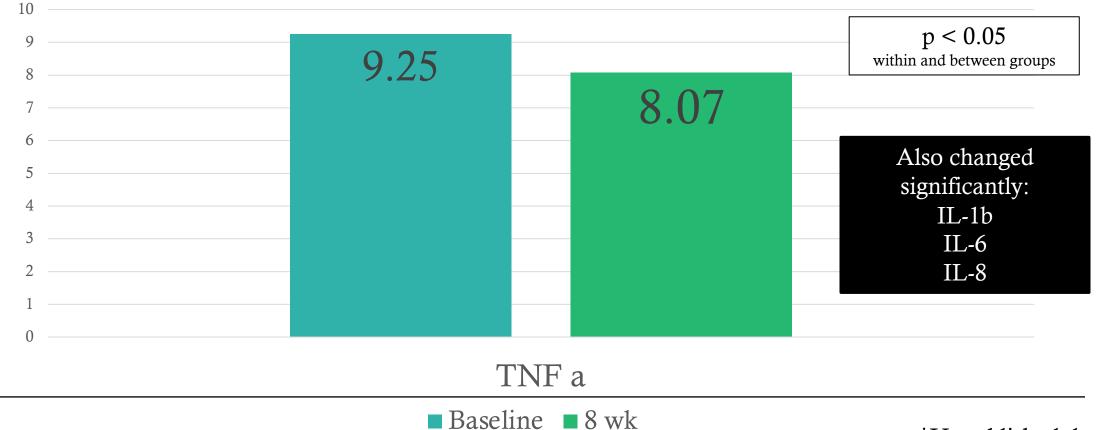
*Unpublished data

10

0

Tumor necrosis factor α (TNF α) is a pro-inflammatory cytokine (signaling molecules that control inflammation) whose expression is increased in a variety of cancers. In particular, in breast cancer it correlates with augmented tumor cell proliferation, higher malignancy grade, increased occurrence of metastasis and general poor prognosis for the patient. Florenxia Mecogliano, et al. "Tumor Necrosis Factor α Blockade: An Opportunity to Tackle Breast Cancer." Frontiers in Oncology. 2020

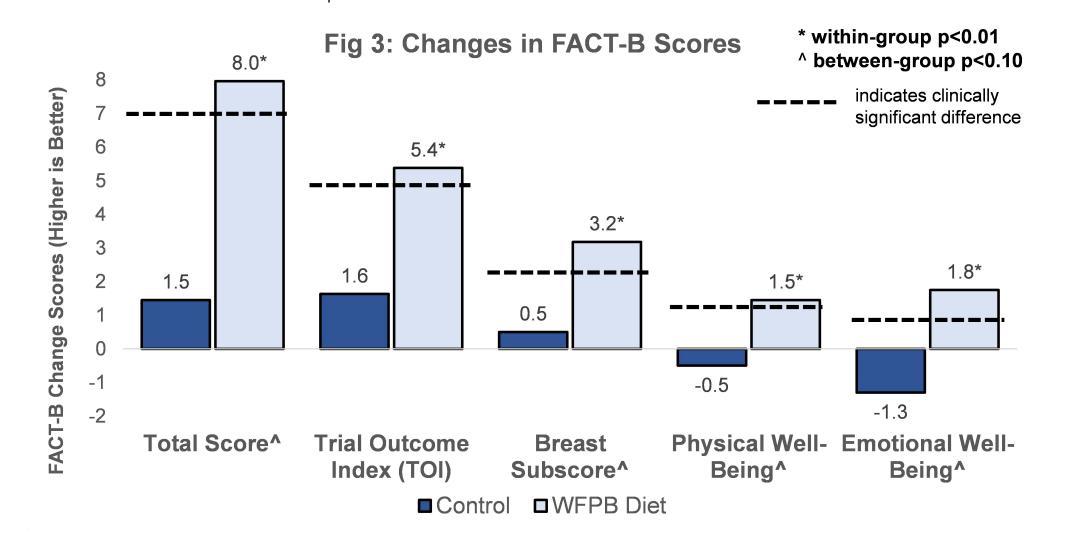
Tumor Necrosis Factor α (TNF α)



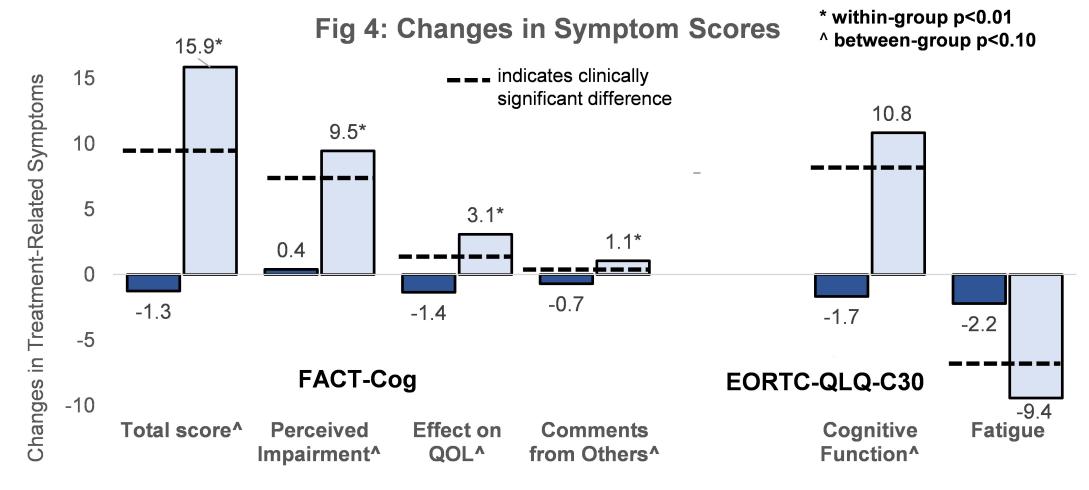
*Unpublished data

OVERALL QUALITY OF LIFE

Campbell EK, Campbell TM, Culakova E, et al. A Whole Food, Plant-Based Randomized Controlled Trial in Metastatic Breast Cancer: Feasibility, Nutrient, and Patient-Reported Outcomes. *Breast Cancer Research and Treatment.* 2024.



PERCEIVED COGNITIVE FUNCTION AND FATIGUE



Control

□ Intervention

FEASIBILITY

95% of intervention subjects were compliant with the diet

94.3% of all recorded calorie intake (from all intervention subjects) was 'on-plan'

100% completed all 3 testing visits

Minimal adverse events

DIETARY INTAKE (94% OF FOODS RECORDED WERE 'ON-PLAN')

| Nutrient Intake of the Intervention Group | | | |
|---|--------------|---------|--|
| | Week 8 Final | Percent | |
| | | Change | |
| Total weight (g) | 3192.5 | 16.0% | |
| Energy (kcal) | 1321 | ₽ 25.9% | |
| Fat (% of total kcal) | 20.4 | ♣ 43.0% | |
| Carbohydrate (% of total kcal) | 66.2 | | |
| Protein (% of total kcal) | 12.6 | ₽ 13.7% | |
| Dietary Fiber (g/1000 | 30.8 | | |
| kcal) | | | |

FEASIBILITY



Do you feel that your health has benefited from the study intervention?

19 of 19 respondents: Yes

RELATED TO REAL WORLD FEASIBILITY... ...WHAT ABOUT FOOD COST?

Previous research

Public perception





ANALYSIS OF T2DM STUDY FOOD RECORDS

3-day food records

Retailer prices







RESULTS

| Mean Daily Food Costs by Dietary Phase | | | |
|--|-----------------|-------------|-------------|
| | Baseline (n=14) | DASH (n=26) | WFPB (n=14) |
| US Dollar (\$)/day as consumed | \$15.72 | \$12.74* | \$9.78* |
| US Dollar (\$)/day adjusted to 1800 kcal/day | \$15.69 | \$14.92 | \$11.96* |

*p<0.05 when compared to baseline

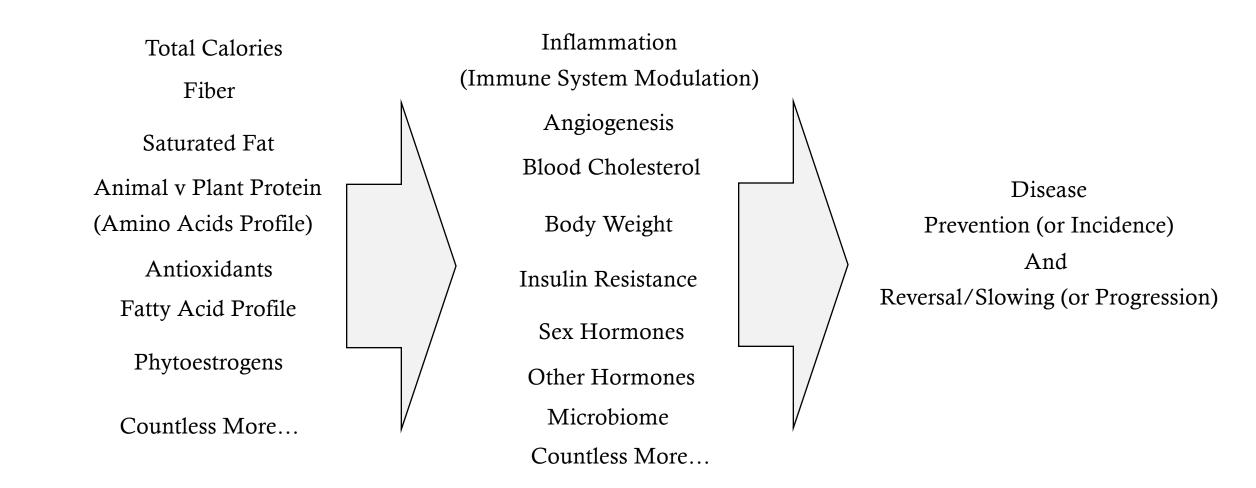
RESULTS

| Mean Daily Approximated <u>Ingredient Only</u> Food Costs by Dietary Phase | | | |
|--|-----------------|-------------|-------------|
| | Baseline (n=14) | DASH (n=26) | WFPB (n=14) |
| US Dollar (\$)/day as consumed | \$11.01 | \$11.81 | \$8.83* |
| US Dollar (\$)/day adjusted to 1800 kcal/day | \$10.84 | \$13.72* | \$10.67 |

*p<0.05 when compared to baseline

LIMITATIONS (AKA EDUCATION AND ADVOCACY OPPORTUNITIES)





NUTRITION: NETWORK IN.... NETWORK OUT

RESOURCES

- Lifestyle Medicine Clinic
- Meals from our research kitchen (coming soon!)
 - <u>NutritionInMedicineRocs.org</u>



THANK YOU

- Coinvestigators: Luke Peppone, PhD, Eva Culakova, PhD, Steven Wittlin, MD, Derick Peterson, PhD, Don Harrington, MBA, and many other collaborators/co-investigators across cancer control, oncology, and endocrinology
- **Study 'staff'**: Lisa Blanchard, Kelly Koch, Laurie Taillie, Nellie Wixom
- **Highland Hospital Foundation** and its donors that have supported research to learn about nutrition as medicine.