# What To Know About Diet and MS





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## SEPTEMBER PROGRAMS

Thursday, September 14
JUMPSTART

Nutritious, Delicious, and Doable Cooking with MS

Tuesday, September 19
YOUR QUESITONS ANSWERED
Nutrition for You

Thursday, September 21
BLACK COMMUNITY MEETUP

Wednesday, September 27 MS MOVES MEETUP



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5 in-person attendees will have the chance to be gifted a one-on-one coaching session!



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Type in your questions using the Questions Box











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Provide comments and engage with the speakers and audience using the **Chat Box** 

**NEW!** Closed captioning

# YOUR SPEAKER



Ilana Katz Sand, MD

Neurologist

New York, NY

# LEARNING OBJECTIVES

Understand what we know about diet and MS

Learn how MS symptoms impact eating and how what we eat can impact MS symptoms

Practical tips and strategies

# Why Are We Interested in Diet?



#### People living with MS are asking!

- Desire to be proactive
- "Natural" approach to complement traditional DMT

Guidance should be based on evidence

Healthcare providers need to advocate for resources to help with access and implementation



# Why Are We Interested in Diet?



Strong environmental component to MS + considerable variability in outcomes

Continued search for modifiable factors that impact prognosis

There is increasing scientific evidence that diet can help manage MS symptoms and may act as a disease modifier

- Mechanistic studies
- Observational studies
- Early clinical trials

# Potential Mechanisms for Dietary Effects



#### Indirect effects mediated by comorbidities associated with worse outcomes

- Obesity
- Cholesterol levels
- Other vascular risk factors

#### Effects directly related to diet

- Direct effects of dietary metabolites
- Effects mediated through the gut microbiota
  - Effects on microbial composition
  - Induction of metabolite production by gut microbiota

# Foods Have Different Molecular Components With Varying Structures

#### Brocolli Sulphoraphane)



#### Garlic Diallyl sulphide



#### Green Tea Epigallocatechin-3gallate



#### Honey Caffeic acid phenethyl ester



#### Soybeans Genistein



#### Tumeric Curcumin



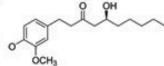
#### Cabbage SIndole-3carbinol





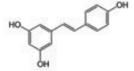
#### Ginger Gingerol





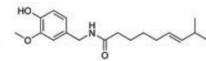
#### Grapes Resveratrol





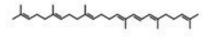
#### Chilli peppers Capsaicin



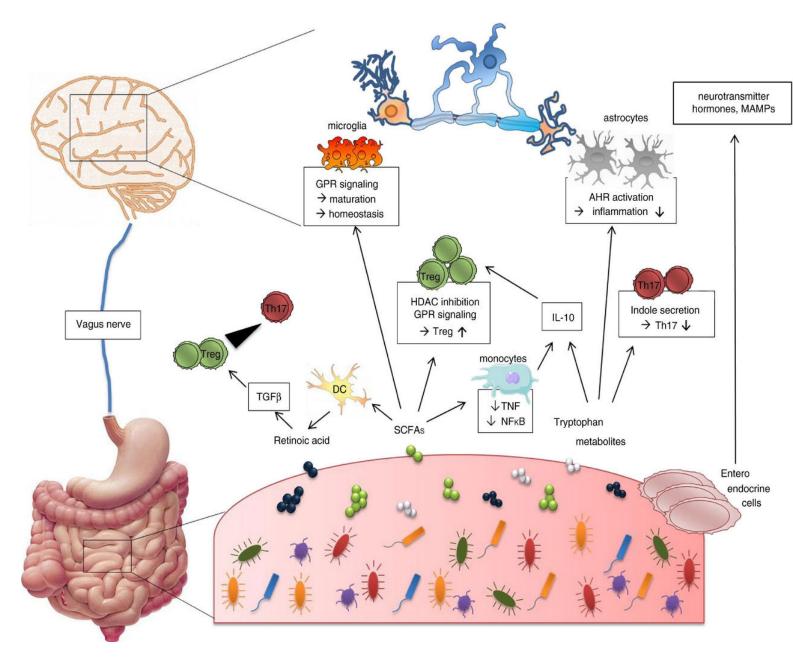


#### Tomatoes Lycopene





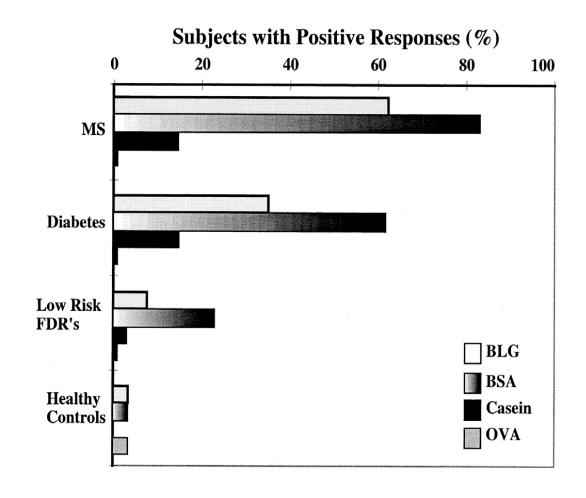
# Communication Pathways of the Gut-Brain Axis



# Evidence For Dietary Components in MS

# Dairy

- A study of T cell reactivities in MS patients showed abnormally heightened responses to multiple milk antigens
- The milk protein butyrophilin has been implicated through antigenic mimicry with myelin Increased inflammation may also be mediated through the intestinal microbiome
- Registry studies with mixed results



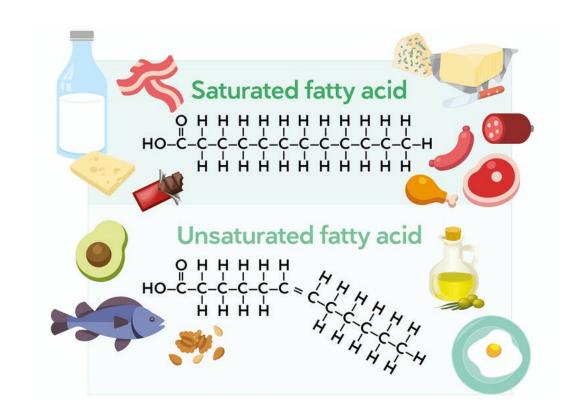
## Fats Have Different Chemical Structures

Saturated fats: fatty acid chain is "saturated" with hydrogen atoms (no double bonds)

Polyunsaturated fats: fatty acid chain has multiple double bonds

 Of particular interest: omega-3 fatty acids (double bond at 3<sup>rd</sup> carbon of FA chain)

Monounsaturated fats: fatty acid chain has one double bond



# Clinical Studies of Saturated Fat Intake In MS

#### **Swank studies**

- Observational studies regarding MS incidence
- Interventional study of low saturated fat diet (Swank, Lancet 1990)

#### Prospective pediatric MS study

- 219 children followed for an average of nearly 2 years
- For every 10% increase in energy intake from saturated fat the risk of relapse was increased 3.37 times

#### CLINICAL DETAILS

	Good dieters	Poor dieters
Minimum disability (grade 1)		
Number (M:F)	23 (14:9)	6 (3:3)
Mean (SD) age (yr)	31.0 (8.4)	30-8 (7-3)
Mean (SD) duration of MS	31-0 (5-8)	25-9 (7-1)
Before trial	2-4 (2-1)	3.5 (2.4)
Diet period	28-6 (5-0)	23.8 (5.9)
Mean (SD) final neurological		
grade [change]	1.9 (2.2) [0.9]	5.3 (1.6) [4.3]
Deaths		,.
All causes	5 (21%)	5 (83%)
MS only	1 (5%)	4 (80%)
Mean lipid intake	' '	, , , , , , , , , , , , , , , , , , , ,
Fats	17.1 (2.4)	35.7 (11.5)
Oils	16.3 (4.3)	11-0 (2-2)
Moderate disability (grade 2)		
Number (M:F)	25 (9:16)	33 (16:17)
Mean (SD) age (yr)	31.8 (9.3)	34-4 (8-2)
Mean (SD) duration of MS (yr)	32-0 (7-2)	28.0 (9.0)
Before trial	4.9 (5.2)	5.3 (4.6)
Diet period	27.1 (6.8)	22.7 (8.0)
Mean (SD) final neurological		
grade [change]	3.6 (2.4) [1.6]	5.3 (1.3) [3.4]
Deaths		
All causes	10 (40%)	25 (76%)
MS only	8 (34%)	16 (66%)
Lipid intake		
Fats	15.4 (3.4)	46-1 (17-0)
Oils	18.2 (3.5)	10.2 (4.1)
Severe disability (grades³-5,		
mean 3·21 [0·4])		
Number (M:F)	24 (7:17)	33 (17:16)
Mean (SD) age (yr)	34.2 (10.2)	37-1 (7-5)
Mean (SD) duration of MS (yr)	33-8 (9-5)	29-9 (10-7)
Before trial	6.2 (7.10)	10.4 (7.8)
Diet period	27-6 (8-5)	19.5 (9.6)
Mean (SD) final neurological		
grade [change]	4.0 (1.8) [0.8]	5-6 (1-0) [2-4]
Deaths		
All causes	8 (33%)	28 (85%)
MS only	5 (21%)	25 (83%)
Lipid intake		
Fats	15.8 (2.6)	36.5 (10.5)
Oils	18-1 (7-9)	10.5 (6.9)

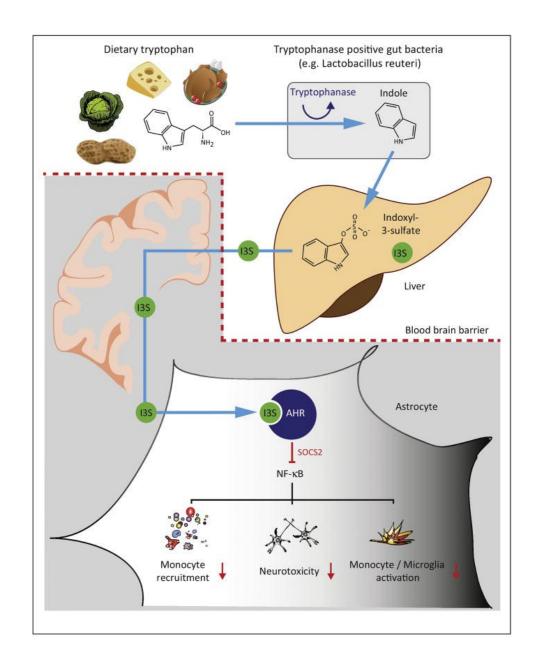
## Polyunsaturated Fats



- Found in fish, walnuts, flax seeds
- Animal model effects on:
  - Immunomodulation
  - Neuroprotection
  - Remyelination and repair
- Epidemiologic studies in MS with conflicting results
  - Nurses Health studies suggest link between MS incidence and intake of alpha linolenic acid
     (ALA)
  - Several other studies demonstrate link between intake of fish and omega-3 fatty acids, but not clearly ALA
- Clinical trials of PUFA supplements with mixed results

## Fruits and Vegetables

- Fiber intake: induction of production of short chain fatty acids by gut microbiota
- Flavonoids (brightly colored) effects on:
  - Immunomodulation
  - Neuroprotection/Repair
- Example: tryptophan derived from the diet can cross the blood brain barrier and have important effects inside the brain



# Fruits and Vegetables (Continued...)

- Pediatric MS study: a one-cup equivalent increase in vegetable intake decreased the risk of relapse by 50% (HR 0.50, p=0.024)1
- Registry-based study found link between higher intake of fruits and vegetables and patient-reported disability and disease activity2



### **Grains and Gluten**



- Studies specifically evaluating the role of gluten, including both animal model studies and those in MS patients, have had mixed results<sup>1</sup>
- NARCOMS study found an association between higher intake of whole grains and lower level of MS-related disability<sup>2</sup>
- Theoretically, high fiber content may be of benefit due to induction of SCFA production

#### Salt

- High salt intake induces the development of proinflammatory immune cells (T<sub>H</sub>17)<sup>1</sup>
- T<sub>H</sub>17 cells that develop in a high salt environment demonstrate a more pathogenic phenotype and mice fed a high salt diet exhibit worse disease<sup>2</sup>
- 70 RRMS patients stratified by sodium intake
  - Medium and high intake with higher relapse rates over 2 years
  - Also, with increased number of T2 lesions

#### Multiple sclerosis

RESEARCH PAPER

### Sodium intake is associated with increased disease activity in multiple sclerosis

Mauricio F Farez, <sup>1</sup> Marcela P Fiol, <sup>1</sup> María I Gaitán, <sup>1</sup> Francisco J Quintana, <sup>2</sup> Jorge Correale <sup>1</sup>

Table 2 Association between sodium intake and exacerbation rate in a regression analysis

	IRR	95% CI	p Value
IRR of exacerbation (univariate model)			
Sodium intake (g/day)			
<2	1 (baseline)	_	-
2-4.8	2.56	1.3 to 4.9	0.005
>4.8	3.37	1.5 to 9.55	0.001
IRR of exacerbation (adjusted model)			
Sodium intake (g/day)			
<2	1 (baseline)	_	-
2-4.8	2.75	1.3 to 5.8	0.008
>4.8	3.95	1.4 to 11.2	0.01
Age (1-year increment)	0.992	0.96 to 1.02	0.59
Gender (male)	1.09	0.49 to 2.42	0.82
Disease duration (1-year increment)	0.99	0.98 to 1.01	80.0
Vitamin D (1 ng increase)	1	0.96 to 1.04	0.85
Smoking (smoker)	1.13	0.56 to 2.28	0.73
BMI (1 unit increase)	0.97	0.87 to 1.07	0.58
Treatment (immunosuppressant vs immunomodulators/untreated)	1.46	0.79 to 2.73	0.22

#### Additional Studies Have Not Confirmed These Effects



- Pediatric case control study of 170 MS patients and 331 controls using food frequency questionnaire (FFQ) to estimate dietary sodium found no link1
- Pediatric study of 174 RRMS participants followed for 1.8 years found no link between relapse rates and sodium intake by FFQ2
- BENEFIT trial data3

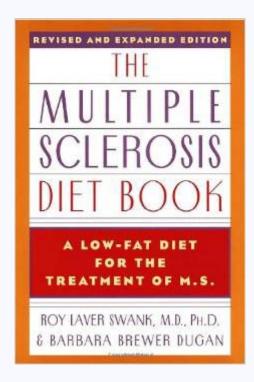
  - Over 400 patients in the trial, followed for 2 years then additional 3 year extension
  - No link between dietary sodium and clinical relapses or MRI lesions

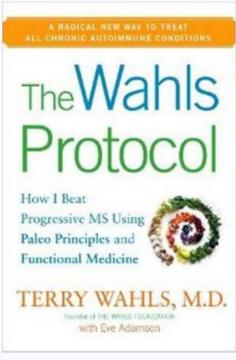


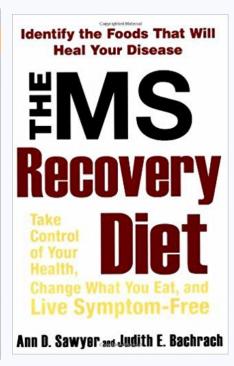
# Evidence For Dietary Patterns in MS

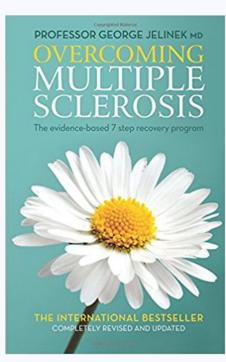


### Many Popular MS Diets To Choose From, BUT...









- Each recommends something different: is one of them "RIGHT?"
- What is the evidence?

## **Overall Dietary Quality**



- Within the HOLISM (Health Outcomes in a Sample of people with MS) study, 2047 participants completed the Diet Habits Questionnaire (DHQ)
  - 10-point increase on the DHQ (higher scores indicating higher quality diet) was associated with a 30% less likelihood of higher disability level.
  - Higher DHQ scores were also significantly associated with better physical and mental health-related quality of life
- North American Research Committee on MS (NARCOMS) Registry, 6989 participants completed a dietary screener questionnaire (DSQ)
  - Participants with top diet quality scores were at 20% lower odds of higher disability scores compared to those in the bottom range.

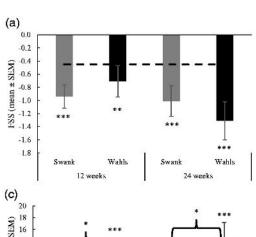
# Dietary Patterns Under Investigation In MS

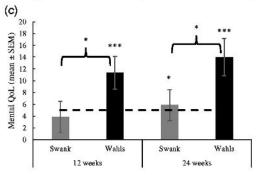


- Swank diet (low saturated fat)
- Modified paleolithic (Wahls protocol)
- McDougall diet (plant-based, very low fat)
- Caloric restriction/intermittent fasting
- Ketogenic diet
- Mediterranean/MIND diet

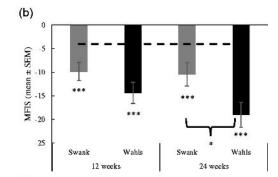
# Modified Paleolithic (Wahls) Diet: WAVES Diet

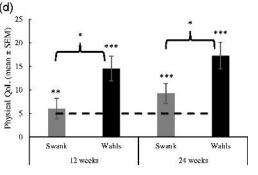
- Swank diet vs. Paleolithic elimination (Wahls) diet
- Paleolithic elimination diet: include meat, fish, certain vegetables and fruits, eliminate gluten, casein, and lectins
- 87 participants with significant fatigue randomized 1:1, followed 24 weeks











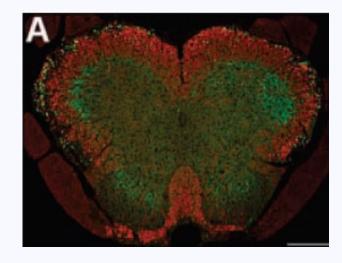
## McDougall Diet



- Based on intake of starchy plant-based foods, with addition of non-starchy vegetables and fruits
  - 10% of energy intake from fats
  - No animal products or oils (including olive oil)
- One study randomized 61 MS patients to either follow the diet or participate as wait-list control x 1 year
  - Primary endpoint (new T2 lesions on MRI) not satisfied
  - No difference in clinical relapse rate
  - Significant reduction in fatigue, moderated by weight loss
- Current study aims to enroll just over 100 participants, focused on fatigue

# Caloric Restriction and Intermittent Fasting

- Chronic caloric restriction of benefit in animal models of MS
- "Fasting mimicking diet" with multiple benefits in animal models of MS
  - Immunomodulation
  - Oligodendrocyte protection
  - Promotion of oligodendrocyte differentiation and repair
- A initial study in MS patients shows that mild chronic caloric restriction and intermittent fasting
  - Are relatively well-tolerated without evidence of harm
  - Result in weight loss, improved emotional well-being



Control

Myelin integrity in spinal cord in mouse MS model



Caloric restriction

# **Ketogenic Diet**



- Shift metabolism by using fatty acids as the primary energy source 
  increase in oxidative phosphorylation and production of ketone bodies
- Decrease in pro-inflammatory cytokines
- Ketogenic diet of benefit in animal models of MS
  - Suppressed expression of inflammatory cytokines and reactive oxygen species
  - Associated with improvements in memory and motor disability
- Open-label single arm studies of KD in MS showed that it was:
  - Relatively well-tolerated
  - Associated with weight loss and improved body composition
  - Resulted in reductions in fatigue, depression, and disability

# Rationale For Pursuing Mediterranean-Style Diet Investigation



- General health benefits are well-established
- Data in cognitive aging
- Combines limited available information about dietary components that might be important
- Reasonable to aim for long-term adherence
  - Lifestyle change rather than a "diet"
  - Budget friendly
  - Household involvement



#### Multiple Sclerosis and Related Disorders

MULTIPLE SCLEROSIS

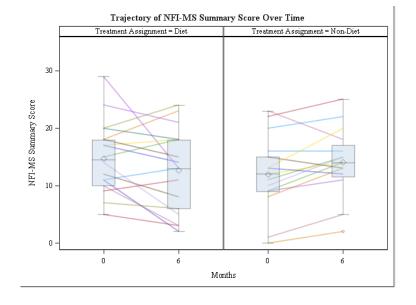
journal homepage: www.elsevier.com/locate/msard

Randomized-controlled trial of a modified Mediterranean dietary program for multiple sclerosis: A pilot study



Ilana Katz Sand<sup>a,\*</sup>, Emma K. T. Benn<sup>b</sup>, Michelle Fabian<sup>a</sup>, Kathryn C. Fitzgerald<sup>c</sup>, Elise Digga<sup>a</sup>, Richa Deshpande<sup>b</sup>, Aaron Miller<sup>a</sup>, Samantha Gallo<sup>d</sup>, Lenore Arab<sup>e</sup>

- Recruitment and adherence went great!
- Dietary intervention participants lost a small amount of weight (approximately one pound per month)
- There was a benefit seen in the intervention group compared to the non-intervention group with respect to:
  - Fatigue
  - Impact of MS symptoms on daily life (MSIS-29)
  - Disability score



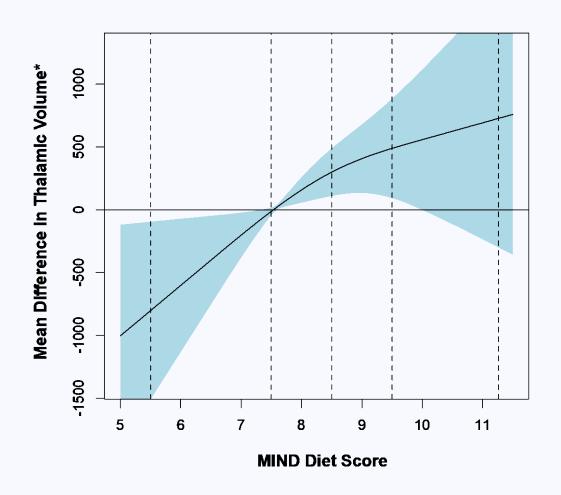
# Mind Diet: Mediterranean-Dash Intervention for Neurodegenerative Delay

MIND diet component servings and scoring

	0	0.5	1
Green Leafy Vegetables <sup>a</sup>	≤2 servings/wk	> 2 to <6/wk	≥6 servings/wk
Other Vegetables <sup>b</sup>	<5 serving/wk	5 – <7 wk	≥1 serving/day
Berries <sup>C</sup>	<1 serving/wk	1 /wk	≥2 servings/wk
Nuts	<1/mo	1/mo - <5/wk	≥5 servings/wk
Olive Oil	Not primary oil		Primary oil used
Butter, Margarine	>2 T/d	1-2/d	<1 T/d
Cheese	7+ servings/wk	1-6/wk	< 1 serving/wk
Whole Grains	<1 serving/d	1-2/d	≥3 servings/d
Fish (not fried) <sup>d</sup>	Rarely	1-3/mo	≥1 meals/wk
Beans <sup>e</sup>	<1 meal/wk	1-3/wk	>3 meals/wk
Poultry (not fried)	<1 meal/wk	1 /wk	≥2 meals/wk
Red Meat and productsg	7+ meals/wk	4-6/wk	< 4 meals/wk
Fast Fried Foods <sup>h</sup>	4+ times/wk	1-3 /wk	<1 time/wk
Pastries & Sweets <sup>1</sup>	7+ servings/wk	5-6/wk	<5 servings/wk
Wine	>1 glass/d or never	1/mo - 6/wk	1 glass/d
TOTAL SCORE			15

- Scored based on brain "healthy" and "unhealthy" foods
- Literature demonstrating slowing in cognitive decline with aging and in progression of Parkinson's disease
- We hypothesized a potential benefit on neurodegeneration in MS, in addition to potential immunomodulatory benefits

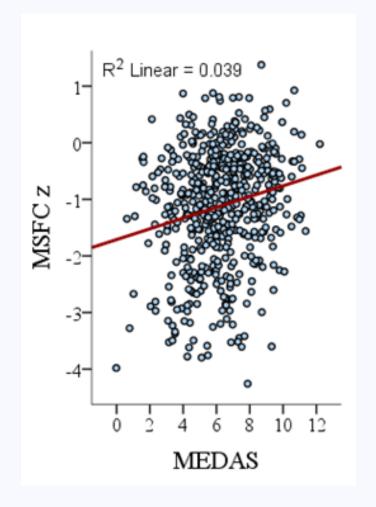
# Association Between MIND Diet Score and Thalamic Volume in MS



## Primary Analysis: MEDAS Score Predicts MSFC

# Multiple Regression Adjusting for All Demographic and Health-Related Covariates

<b>Predictor</b>	В	95%	CI	β	р
Constant	-1.655	-2.368	-0.941		< 0.001
AGE	-0.046	-0.054	-0.038	-0.43	< 0.001
SEX	0.381	0.194	0.569	0.14	< 0.001
RACE	0.422	0.209	0.634	0.16	< 0.001
ETHN	0.151	-0.095	0.396	0.05	0.228
SES	0.017	0.012	0.022	0.26	< 0.001
BMI	0.004	-0.011	0.020	0.02	0.570
MVPA	0.002	-0.002	0.006	0.04	0.235
SLEEP	-0.187	-0.358	-0.017	-0.08	0.032
HTN	0.212	-0.034	0.458	0.07	0.091
DM	-0.090	-0.531	0.350	-0.02	0.687
HLD	0.135	-0.099	0.368	0.04	0.258
<b>SMOKE</b>	-0.142	-0.327	0.043	-0.05	0.131
MEDAS	0.095	0.055	0.134	0.18	< 0.001



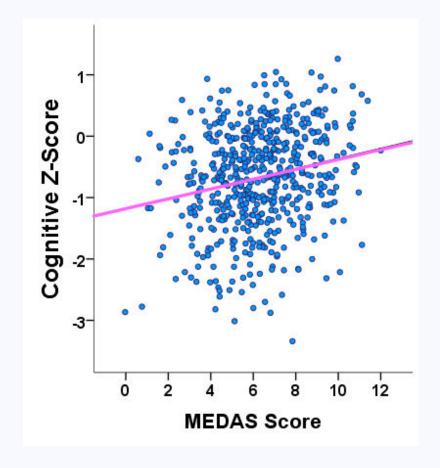
## Secondary Analysis: MSFC Components & PROs

Outcome	В		95% CI	β	p
MSFC	0.095	0.055	0.134	0.18	< 0.001
SDMT	0.869	0.458	1.279	0.17	< 0.001
NHPT	-0.392	-0.584	-0.199	-0.17	< 0.001
T25FW	-0.077	-0.121	-0.033	-0.14	0.001
MSIS-20	-0.993	-1.609	-0.377	-0.13	0.002
FSS	-0.102	-0.170	-0.034	-0.13	0.003
MSWS	-1.312	-2.108	-0.516	-0.13	0.001
PDQ	-0.056	-0.091	-0.022	-0.13	0.001
HADS-D	-0.234	-0.362	-0.105	-0.15	< 0.001
HADS-A	-0.190	-0.338	-0.042	-0.11	0.012

### Primary Analysis: MEDAS Score Predicts Cognition

## Multiple Regression Adjusting for All Demographic and Health-Related Covariates

Predictor	В	[95% CI]	β	P
(Constant)	-1.81	[-2.40, -1.22]		< 0.001
AGE	-0.03	[-0.04, -0.02]	-0.37	< 0.001
SEX (F)	0.37	[0.21, 0.52]	0.18	< 0.001
RACE (W)	0.32	[0.13, 0.51]	0.16	0.001
ETHN (H/L)	0.35	[0.12, 0.58]	0.13	0.003
SES	0.02	[0.01, 0.02]	0.29	< 0.001
BMI	0.01	[0.00, 0.02]	0.07	0.077
MVPA	0.00	[0.00, 0.00]	-0.03	0.477
SLEEP	0.04	[-0.10, 0.17]	0.02	0.622
HTN	0.15	[-0.06, 0.35]	0.06	0.157
DM	0.20	[-0.16, 0.55]	0.04	0.284
HLD	-0.08	[-0.27, 0.11]	-0.03	0.416
SMOKE	-0.03	[-0.18, 0.12]	-0.01	0.708
MEDAS	0.08	[0.05, 0.11]	0.20	<0.001



# Current Recommendations: NMSS Wellness Research Working Group



 https://www.nationalmssociety.org/Living-Well-With-MS/Diet-Exercise-Healthy-Behaviors/Diet-Nutrition

- Prepare meals at home as much as possible
- Incorporate colorful fresh fruits and vegetables daily
- If you choose to eat grains, choose whole grains over refined grains
- Avoid/limit processed foods and added sugars as much as possible

## Getting Started...



Aim to make positive changes that will become your lifestyle, rather than looking for a "diet"

#### Think about your goals

- Short-term
- Long-term

#### Consider how your symptoms impact implementation of ideas

- Fatigue: meal planning, energy conservation techniques
- Adaptive cooking tools

#### Include your household in your plans

Children and partners need healthy habits too!

## Getting Started...



#### Practice makes progress

Strive for progress, not perfection

#### Ask for help:

- Talk to your primary care doctor
- Arrange a visit with a dietitian
- Ask if your MS Center has a staff member who is able to counsel about diet

## Be kind to yourself!



## **Tips for Energy Conservation**



#### Plan, plan, plan!

- Set out meal plans at a time interval that is convenient for you in terms of groceries, for example many people use one week at a time
- Plan to do the bulk of preparation at times when you usually have more energy. This may be different for you on different days depending on the other activities you need to do that day.
- Consider dividing up the tasks required to prepare meals so that you don't have to do it all at once
- Include plans for everything you are going to eat, including snacks, so that you won't find yourself hungry and out somewhere where access to healthy food isn't easy or affordable

Consider a grocery delivery service so you don't use time and energy in the store

Utilize recipes that prioritize efficiency: "one pot" or "sheet pan" dinners minimize clean up

## **Tips for Energy Conservation**



Prepare enough to allow for leftovers: tonight's dinner often makes a great lunch for tomorrow and preparing more of the same is less work than preparing another meal

#### Use the freezer and stock your pantry

- Prepare extra/leftovers that can be frozen and used on a day when you don't have the energy to prepare
- Frozen vegetables are often as high quality as fresh (or even higher)
- Cans of beans, nut butters, whole grain pastas are shelf-stable items that can be made into meals

Take family and friends up on offers to help

C. Olsten MS
Patient Wellness
Program





## Key Take-Aways



- 2. There is currently limited evidence for a role for particular dietary factors and patterns on MS outcomes
- 3. General principles of healthy eating are reasonable to recommend now; ask for help to get started!

#### Acknowledgements



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La Toija Striggles

Julisa Cepin

Lady Navia

Konul Azimzade

**Daphne Verter** 

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Jasmin Patel

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Francesco La Rosa

Jordyn Anderson

Sarah Levy

Maggie Lorenzetti

Microbiology

Jose Clemente

Johns Hopkins

Kate Fitzgerald



Our patients and their families!

## **THANK YOU!**





# Q+A

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## How To Sleep Well Despite Your Symptoms





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