

NUTRACEUTICALS FOR VISUAL (SPORTS) PERFORMANCE

COPE 69590-GOI

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Disclosures: ¹Eye Promise Scientific Advisory Board
² Eye Promise Research Grant Recipient

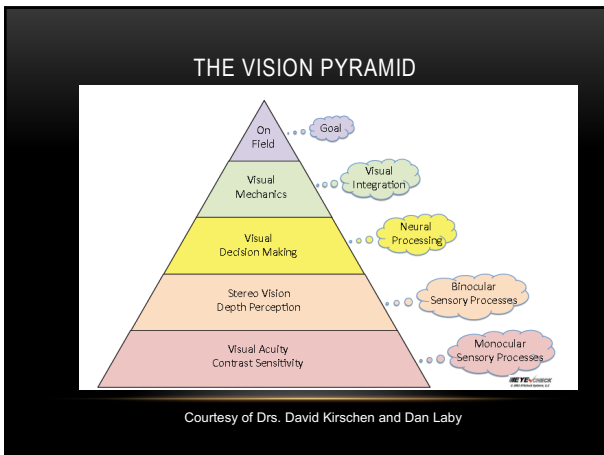
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VISUAL PERFORMANCE ENHANCEMENT AND SPORTS

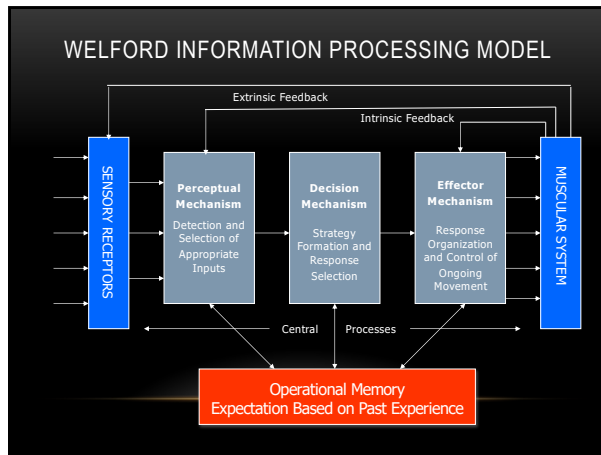


attribution www.nytimes.com

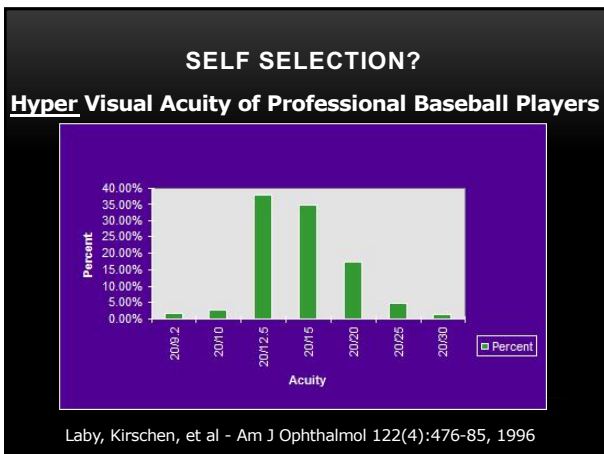
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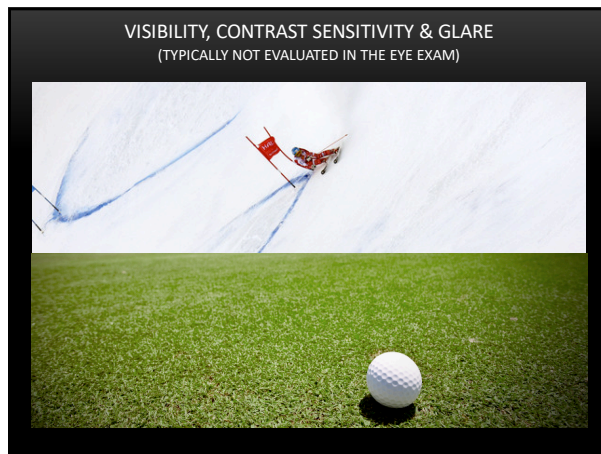
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OPTIONS TO ENHANCE VISUAL ACUITY & CONTRAST SENSITIVITY

- Refractive compensation
 - Methods for refractive compensation
- Filters / tints to enhance visibility of important features
- Visual feedback training to enhance CS function
- **Nutrient intake of carotenoids**

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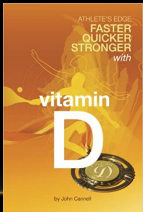
WHICH NUTRIENTS ARE BENEFICIAL FOR EYE HEALTH & FUNCTION?

- Endogenous – 60% of antioxidant capacity
- Exogenous Nutrients – 40% of antioxidant capacity of eye
 - Lutein & Zeaxanthin
 - Vitamin D3 (really a hormone)
 - Omega-3 fatty acids
 - Non-enzymatic antioxidants
 - Vitamins A, C, E, COQ10 and glutathione
 - Minerals
 - Magnesium, Zinc, Selenium
 - OTHER NEWCOMERS:
 - quercetin, resveratrol, pterostilbene, pycnogenol and astaxanthin

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WHAT DO THESE NUTRIENTS DO?

- Protective effects for age-related changes to the crystalline lens & retina (eg, cataracts & AMD)
 - AREDS B carotene, C and E, Zinc, lutein & zeaxanthin
- Epigenetic modulators (longevity)
 - Omega-3's - dry eye / MGD / AMD
 - Vitamin C, D3 & magnesium- arcus, retina CVDz
 - Vitamin A & selenium, night vision, dry eye
- Protect both retina and brain
 - Lutein, Zeaxanthin, DHA



John Cannell, MD

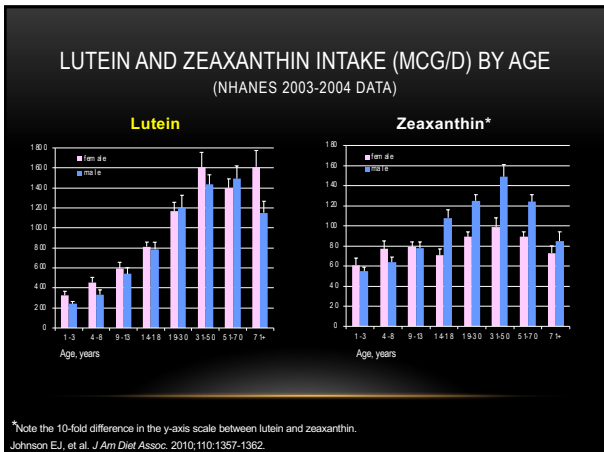
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WHERE ARE THESE NUTRIENTS FOUND?

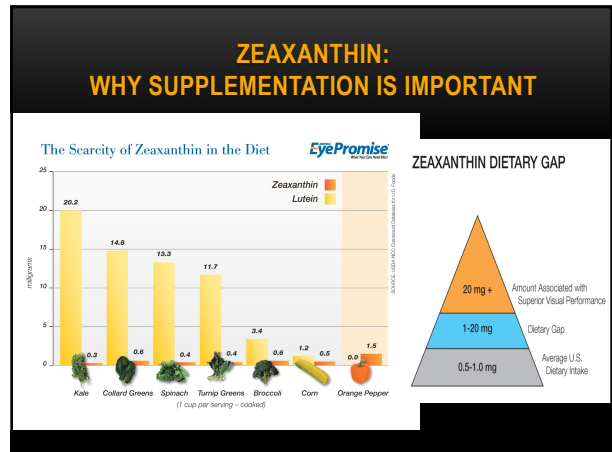
- Green, leafy vegetables such as spinach, kale, collards and orange peppers / paprika / goji berries
- Salmon, tuna, and other oily (cold-water) fish
- Eggs, nuts, beans, and other protein sources
- Oranges and other citrus fruits - C / bioflavonoids
- Colorful fruits and vegetables (bell peppers, berries)- polyphenols
- Whole grains such as quinoa, brown rice, whole oats
- Flaxseed oil (females), EPR, black currant seed oil



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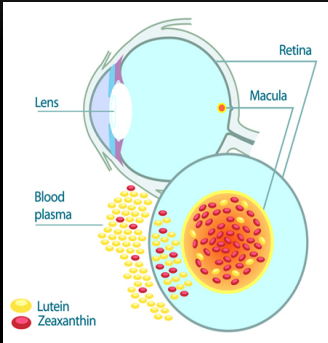
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THINK - ZEAXANTHIN INTAKE

- Zeaxanthin and Lutein accumulate at a 2:1 ratio (Z to L) in the fovea
 - Zeaxanthin isomers (Z & M) are the foveal carotenoids
 - The fovea has 3x the metabolic activity of any other tissue
 - Potently increase foveal (1 degree) MPOD
 - Z publication data supports fewer anti-VEGF injections required
- AVG American diet consists of 5:1 ratio of L to Z and the avg American gets << 1 mg of Z per day
 - Why Supplementation is Important

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COMPOSITION OF MACULAR PIGMENT



Zeaxanthin and Lutein – 2:1 ratio in the macular pigment.

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NUTRITION FOR BETTER PERFORMANCE: ZEAXANTHIN

- Nutrient found in bright colored fruits and vegetables
- 20+ years of research, 150+ published studies to support zeaxanthin's role in healthy vision
 - Masked, placebo-controlled studies
- Higher daily intake of zeaxanthin relates to improved vision, faster response time



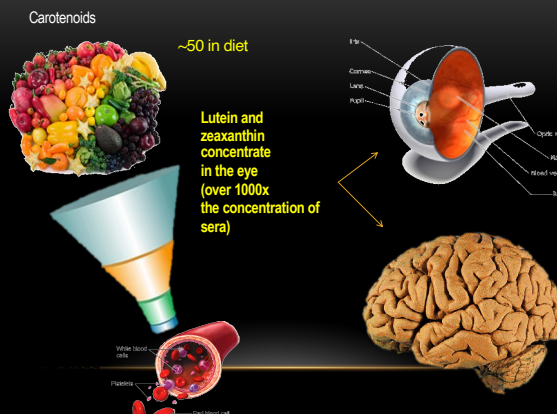
Richer et al, The Zeaxanthin and Visual Function Study (ZVF), 2011, ClinicalTrials.gov Identifier: NCT00564902

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Carotenoids

~50 in diet


Lutein and zeaxanthin concentrate in the eye (over 1000x the concentration of sera)



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LUTEIN AND ZEAXANTHIN MISSING FROM THE DIET CAN INFLUENCE VISUAL FUNCTION THROUGH OPTICAL AS WELL AS BIOLOGICAL MECHANISMS

- **Optical mechanisms**
 - GD, GR, CSF, CC, VISUAL RANGE
- **Biological mechanisms – eyes + brain**

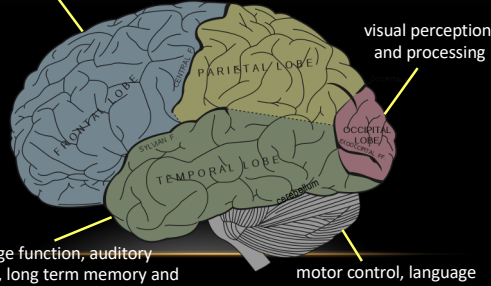


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REGIONS OF THE BRAIN ANALYZED FOR CAROTENOIDS

(CRAFT ET AL., 2004; VISHWANATHAN ET AL., 2011, JOHNSON ET AL., 2012)

thinking, planning, central executive function, attention and motor execution

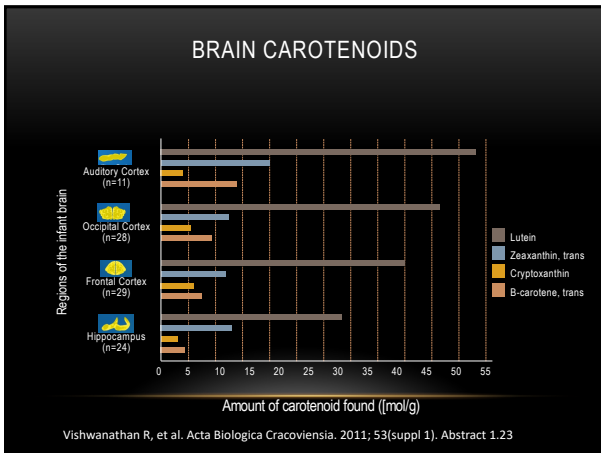


visual perception and processing

Language function, auditory perception, long term memory and emotion

motor control, language and attention

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VISUAL PERFORMANCE & MACULAR PIGMENT

- SPORTS
- MILITARY
- DRIVING

- CFF
- CSF
- DG
- Visual Adaptation
 - Cones
 - Rods
- UFOV
- Peripheral Motion Sensitivity

CNS Benefits
 "MPOD & Steady State VEPs"
S Anna Ceravalo et al 2019
 1) Better Visual Memory
 2) Faster Visual Processing
 3) Superior power of EEG activation
 4) Better signal to noise ratio

fMRI Lidbergh et al 2019
 * Neuroplasticity compensation in older adults

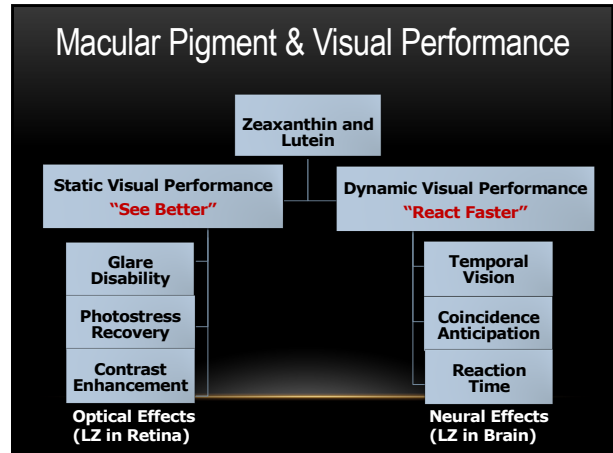
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INDIVIDUAL DIFFERENCES IN VISUAL FUNCTION

HAMMOND & BUCH, EXP EYE RESEARCH 2020

- There are dramatic individual differences in visual function even in college students
 - Vernier acuity
 - Resolution Acuity
 - Lens density
 - Intra-ocular light scatter
 - MPOD by retinal eccentricity
 - Photoreceptor loss
- Performance declines before midlife aging and obvious disease, and may indicate poor aging and susceptibility to disease
- Concierge optometry ?

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EFFECTS OF SUPPLEMENTATION

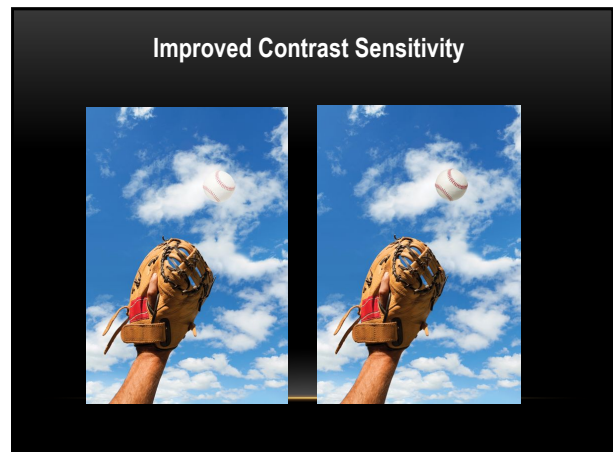
TABLE 3. Changes in Macular Pigment and Visual Function Compared to Placebo

Variable	Slope, Change per Day*	SE of Slope	P Value
MPOD 10'	0.00025	0.00006	<0.0001†
MPOD 30'	0.00025	0.00005	<0.0001†
MPOD 60'	0.00013	0.00005	0.006†
MPOD 105'	0.00016	0.00004	0.0004†
Photostress recovery	-0.019	0.008	0.013†
Glare disability	0.00018	0.00014	0.21
Chromatic contrast	0.00037	0.00017	0.030†

* Daily change in treatment group versus daily change in placebo group.
 † P < 0.05.

Hammond BR, et al. A double-blind, placebo-controlled study on the effects of lutein and zeaxanthin on photostress recovery, glare disability, and chromatic contrast. Invest Ophthalmol Vis Sci. 2014;55:8583-8589. DOI:10.1167/iov.14-15573

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CONTRAST SENSITIVITY IN SPORT

- Judgment of subtle differences in contrast:
 - Between the “target” and its background
 - Rotation of the “target”
- Helps to better judge speed & trajectory
 - **Research: Athletes > Non-athletes**
- Reduced sensitivity may contribute to performance inconsistency due to variable lighting conditions, figure-ground characteristics

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YELLOW RANGE FILTERS

- Selectively filters shorter wavelength light
- Ocular media scatters short wavelength light more - improves contrast by eliminating some of this “internal glare”
- May enhance contrast differences (contours) by reducing chromatic aberration




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CHROMATIC ABERRATION

SYMPOSIUM PAPER

Effect of Ocular Chromatic Aberration on Monocular Visual Performance

LARRY N. THIBOS*, ARTHUR BRADLEY*, and XIAOXIAO ZHANG†
Department of Visual Sciences, School of Optometry, Indiana University, Bloomington, Indiana

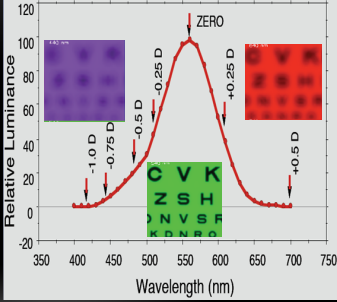
“In the current view, chromatic aberration is the most important optical imperfection of the well corrected eye.”

Thibos, L., et al., Optom. & Vis. Sc., 1991

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CHROMATIC ABERRATION

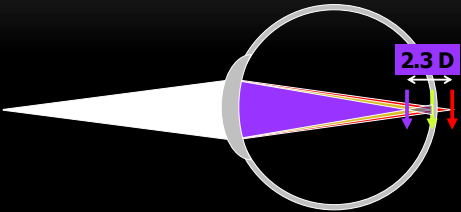
The inability of the ocular system to bring the various colors of light to focus at a single point.



Thibos, 5th Wavefront Congress, Feb 2004

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CHROMATIC ABERRATION

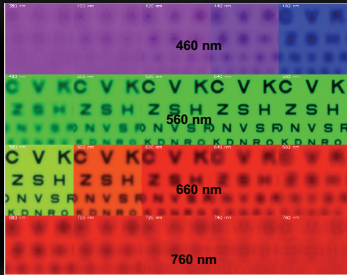


Chromatic Aberration of the Eye

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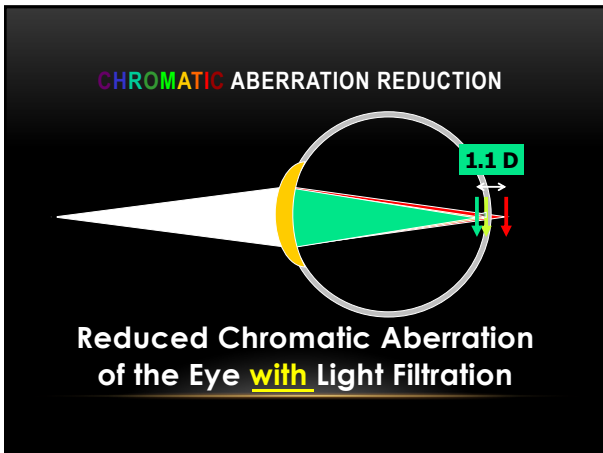
CHROMATIC ABERRATION

Decreases the visual system's potential speed of reaction to individual object's color, shape and speed differences across the range of wavelengths

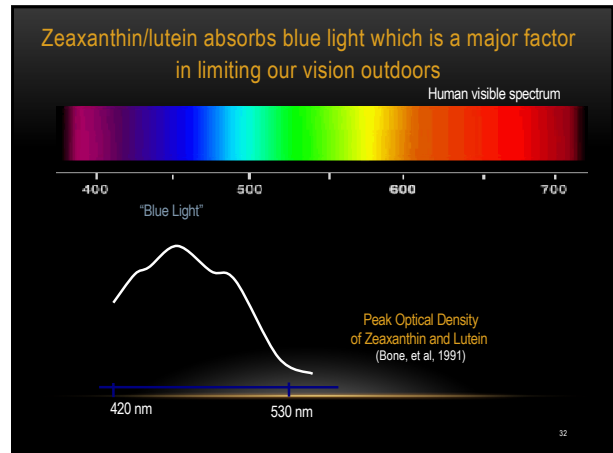


Thibos, 5th Wavefront Congress, Feb 2004

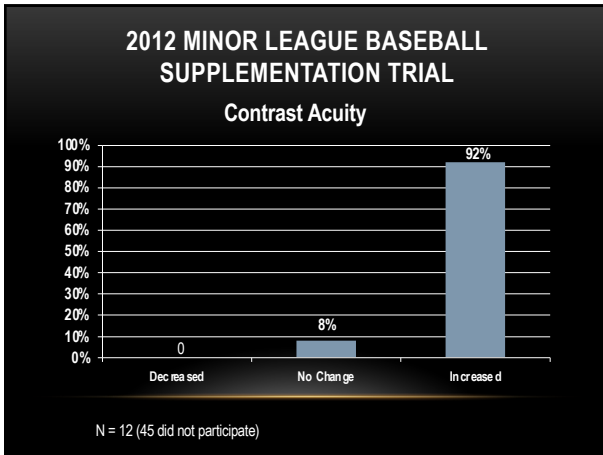
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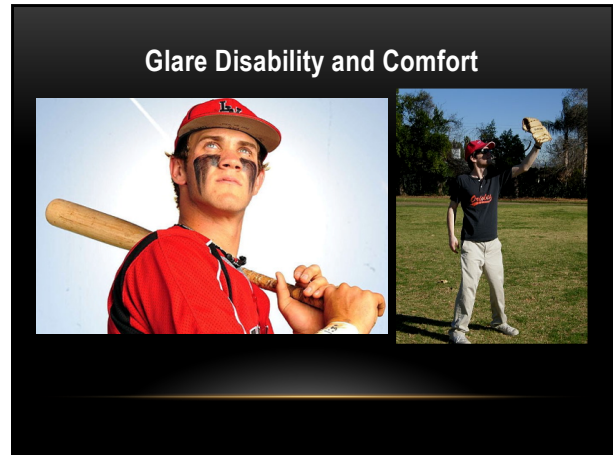
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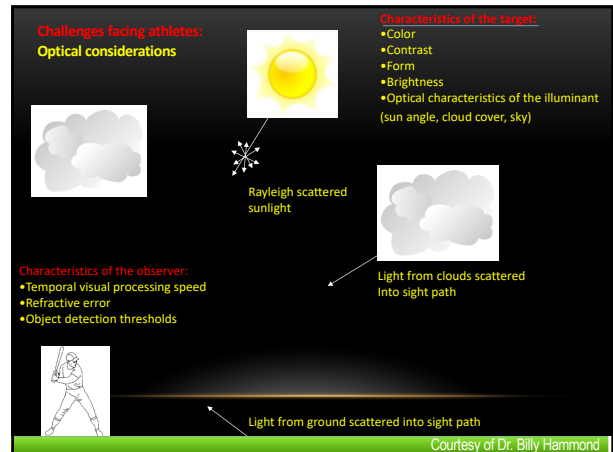
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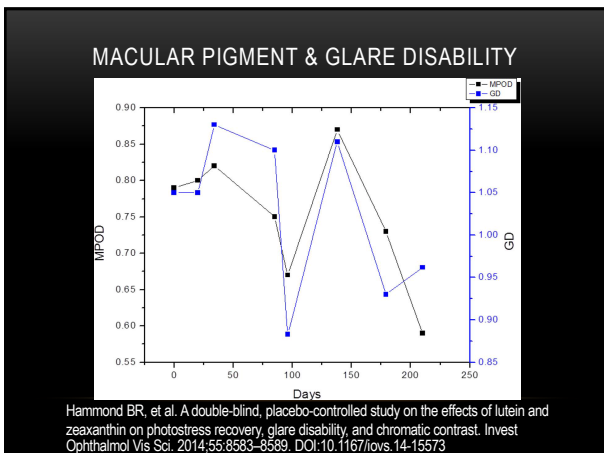
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- ### GLARE EFFECTS IN SPORT
- Judgment of "target" speed and trajectory can be affected by
 - Sun angle and intensity
 - Stadium lighting
 - Color contrast between "target" and background
 - Glare sensitivity and slow glare recovery may contribute to errors in certain game conditions

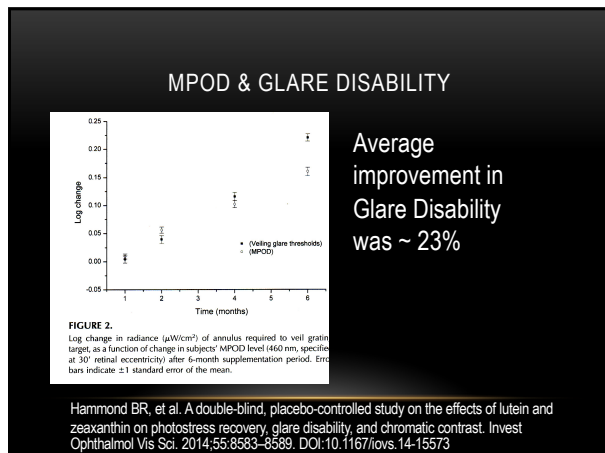
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PHOTOSTRESS: VISUAL RECOVERY AFTER A BLINDING LIGHT EXPOSURE

- Improvement in photostress recovery has been found to be proportional to the level of macular pigment

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AVERAGE GLARE DISABILITY IMPROVEMENT FROM LOW TO HIGH MACULAR PIGMENT IS ABOUT 40% (AVERAGE IS ~23%)

Increased straylight 40%

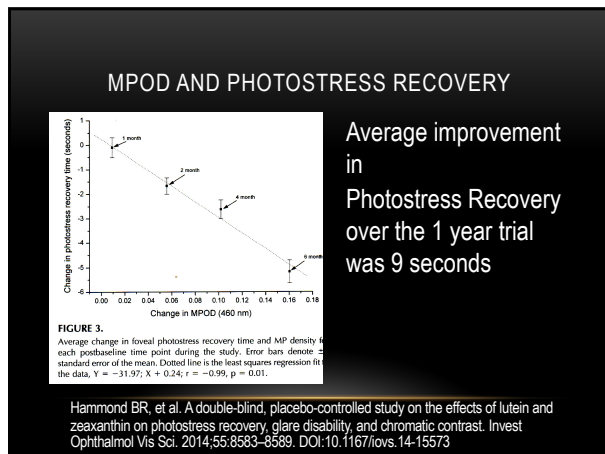
Image: TJTP Van Den Berg

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PHOTOSTRESS RECOVERY EFFECTS IN SPORT

- Some sport situations require the athlete to move quickly between areas of bright sunlight and shadow
- Slow visual recovery following photoreceptor supersaturation can lead to performance errors
 - These errors may put the athlete at risk for injury, such as in cycling sports or downhill skiing

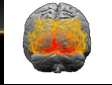
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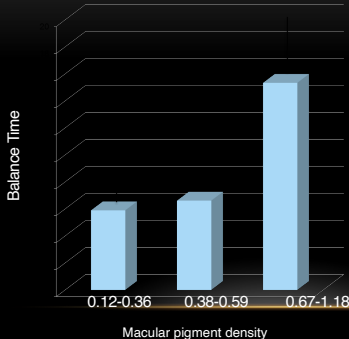
NUTRITION EFFECTS ON NEURAL PERFORMANCE

- MPOD is linked to L and Z levels in the brain
- Neuroimaging of brain structure in vivo confirms L&Z influences white matter integrity, particularly in regions vulnerable to age-related decline
- L and Z are incorporated in cell membranes and axonal projections, which serve to enhance inter-neuronal and neural-glia communication
- 66-76% of total carotenoid concentration in occipital cortex, but highly variable

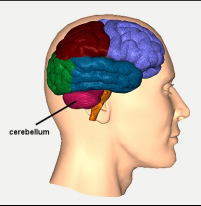


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CNS EFFECTS OF L AND Z



Macular pigment density	Balance Time
0.12-0.36	~1.5
0.38-0.59	~2.0
0.67-1.18	~3.5



Renzi, LM, Bovier ER, Hammond BR. A role for the macular carotenoids in visual motor response. *Nutritional Neurosci* 2013; 16:262-8.

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VISUAL MOTOR EFFECTS OF A CAROTENOID INTERVENTION (1 YEAR STUDY)

Study design: Randomized, double-blinded, placebo-controlled, intervention trial

Subjects: Healthy young healthy adults (18-32 yrs) N = 92 in 1 year study **4 month data**

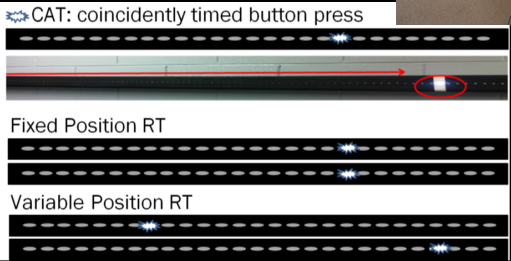
3 Interventions:

- 1) Placebo (n=10)
- 2) Zeaxanthin, 20 mg/d (n=29)
- 3) Multi-condition, 26 mg Z, 8 mg L, 1480 mg Omega-3 fatty acids (n =25) (Eye Promise Vizual Edge Pro)

Bovier ER, Renzi LM, Hammond BR. A double-blind, placebo-controlled study on the effects of lutein and zeaxanthin on neural processing speed and efficiency. *PLoS ONE* 9 2014; 9:e108178.

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Cross-sectional arm (n = 92): LZ measured in the retina strongly related ($p < 0.01$) to fixed and variable reaction time, temporal contrast sensitivity, Critical flicker fusion thresholds, Coincidence anticipation timing (15 mph).

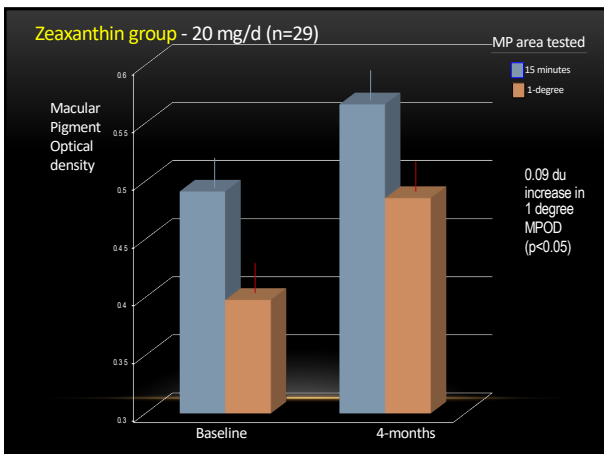


CAT: coincidentally timed button press

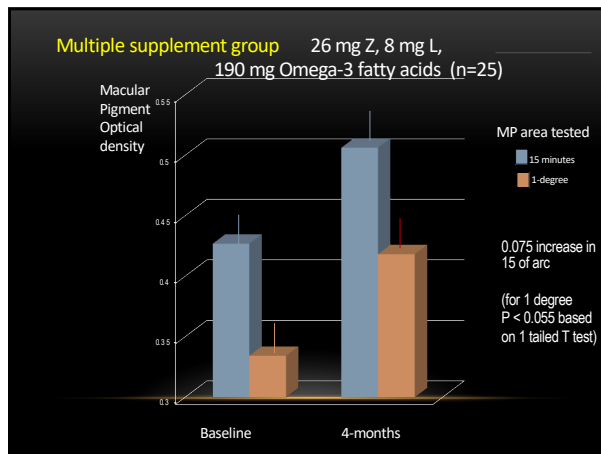
Fixed Position RT

Variable Position RT

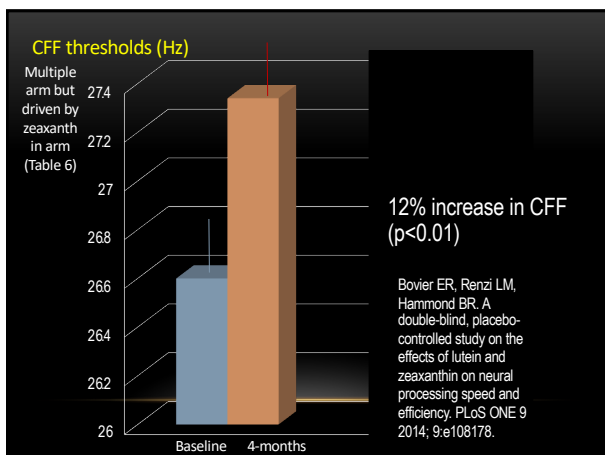
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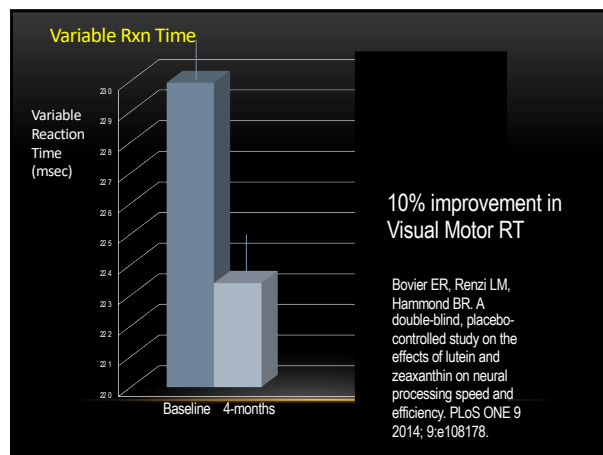
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MACULAR PIGMENT AND TEMPORAL CONTRAST SENSITIVITY

PRESENT A GRATING IN TIME RATHER THAN SPACE

- Light & dark adaptation
- Age related vision loss
- Disease & Performance

Temporal Frequency	Group	Baseline	Final	Change	z-value	p-value*
Foveal 1.0 deg/beat	Treatment	0.32 ± 0.13	0.29 ± 0.14	-0.06	2.53	<0.05
	Placebo	0.22 ± 0.18	0.25 ± 0.19	-0.09	0.34	2.37
Foveal 1.0 deg/beat	Treatment	1.45 ± 0.51	1.37 ± 0.50	-0.15	0.64	0.52
	Placebo	1.34 ± 0.34	1.37 ± 0.10	-0.03	0.90	<0.05
Foveal 0.4 deg/beat	Treatment	1.85 ± 0.51	1.77 ± 0.53	-0.09	0.75	0.45
	Placebo	1.23 ± 0.36	1.27 ± 0.10	-0.02	0.37	2.16
Peripheral 1.8 deg/beat	Treatment	0.26 ± 0.16	0.44 ± 0.20	+0.18	2.80	0.01
	Placebo	0.47 ± 0.23	0.65 ± 0.21	+0.18	1.45	0.15
Peripheral 1.8 deg/beat	Treatment	1.10 ± 0.51	1.39 ± 0.50	+0.30	0.48	<0.05
	Placebo	1.14 ± 0.15	1.20 ± 0.11	+0.06	1.88	0.04
Peripheral 0.4 deg/beat	Treatment	0.95 ± 0.14	1.00 ± 0.11	+0.10	1.41	<0.05
	Placebo	1.05 ± 0.15	1.11 ± 0.11	+0.05	1.35	0.08

* z-values without Bonferroni correction for multiple comparisons

Average improvement in t CSF was 20%

Bovier ER, Hammond BR. A randomized placebo-controlled study on the effects of lutein and zeaxanthin on visual processing speed in young healthy subjects. Arch Biochem Biophys. 2015;15(572):54-57.

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WAYS OF MAKING THE BRAIN FASTER AND MORE EFFICIENT

Caffeine blocks adenosine receptors; makes the brain faster

LZ connects cells makes them interact more efficiently; also makes the brain faster through structural change

ions and small molecules

Cell 1, Connexion, Cell 2

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SUMMARY OF EFFECTS

Sources of influence

- Filtering scatter
- Filtering chromatic borders
- Optics
- Synaptic Efficiency
- Antioxidant-inflammatory
- Biology

Levels of effects

Outcome

- Improved Ocular Optics
- Lower oxidative stress
- Lower inflammatory stress
- Reduced light damage
- Increased Neural efficiency

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INCREASED VISUAL PROCESSING SPEED

THE SCIENCE OF THE SWING

LOOKING: It takes 100 milliseconds for the eye of the batter to see the ball and send an image to the brain.

THINKING: It takes 75 milliseconds for the brain to process the information and decide the speed and location of the pitch.

BEINGING: The batter then must decide, in just 25 milliseconds, whether to swing or let the ball go by. It takes another 25 milliseconds to pick a swing pattern - high, low, inside, outside.

ACTING: The swing starts when the brain sends signals to the legs to start the batter's stride. This takes at least 15 milliseconds.

CONTACT: The bat must meet the 3 inch spinning sphere within an eighth of an inch of disc center and at precisely the right millisecond.

SWINGING: The swing itself takes 150 milliseconds. During the first 50 milliseconds, the batter can stop. But after 100 milliseconds, the bat is moving too fast, and the swing cannot be checked.

MARGIN OF ERROR: If the batter is slower than 100 milliseconds, the ball will pass the first or third base lines.

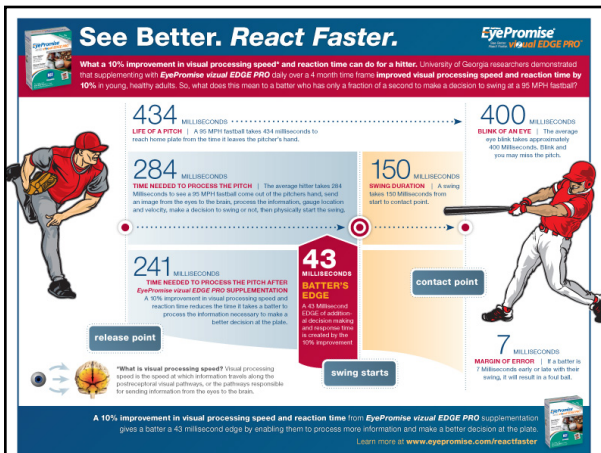
Batter must start to swing at this point

Ball: 2 1/4" dia, diameter, 42" max. length, average 52" inches weight

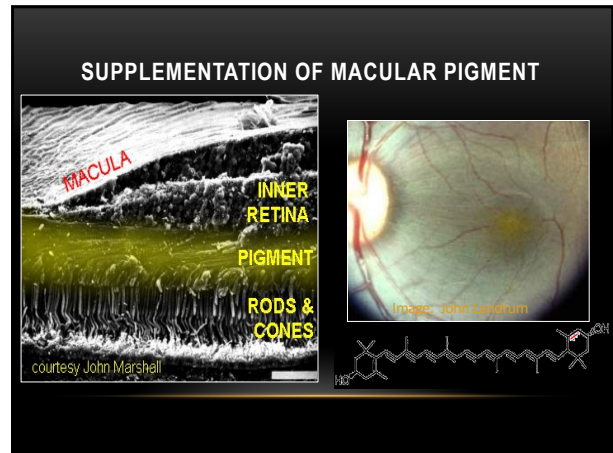
Bat: 34" dia, 3" diameter, 3' in 5/8" inches weight

* SF Chronicle

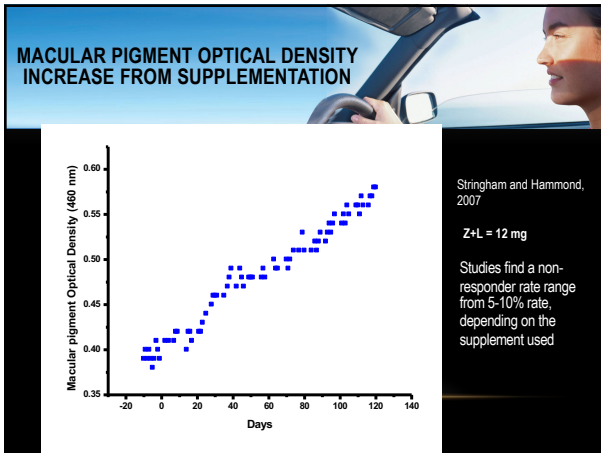
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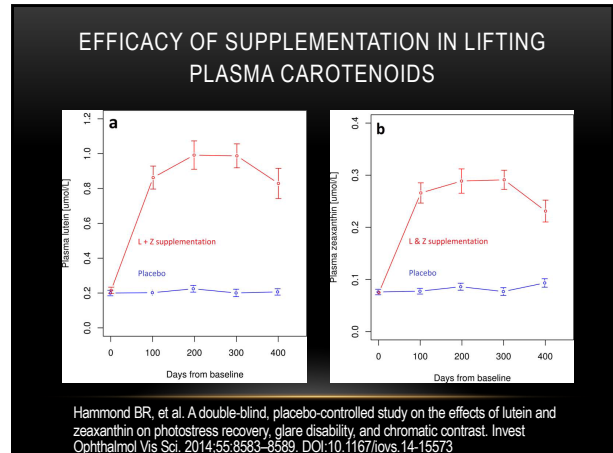
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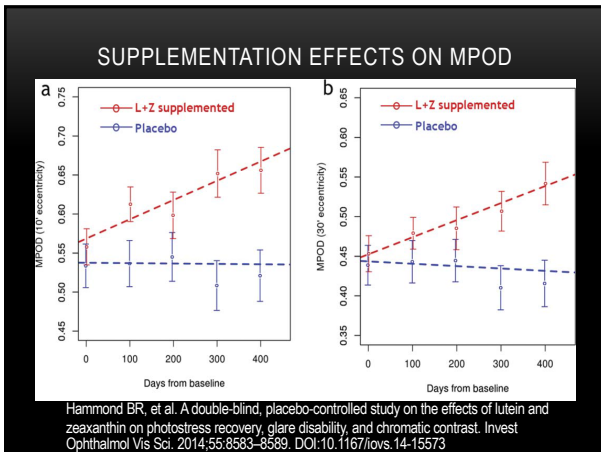
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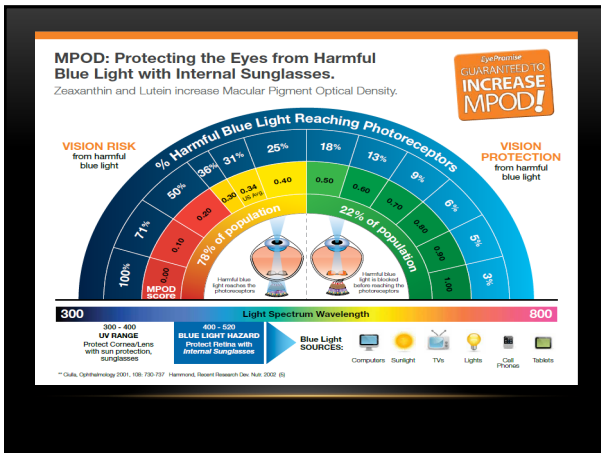
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Simple & Efficient 5 minute clinical test of foveal macular pigment for 1 or 2 eyes

QuantEYE MPS II

- Validated
- Identify low macular pigment
- Assess change with supplementation

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2012 MLB Season Trial

- **100%** of the players did not need their sunglasses as often outside.
- **89%** of players were less sensitive to light.
- **63%** of players had less issue with losing the ball in the sky.
- **63%** of players were able to find the ball easier in the night sky and/or stadium lights.
- **71%** of players were able to pick up on the ball's rotation and/or see the threads on the ball better.

N = 43

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EXTRA PROTECTION BY INCREASED MP

- Reduced "blue-light" damage
- ↑ antioxidant and anti-inflammatory effects
- Possible protection from screen time (esp. in e-sports)

Barker et al., 2011, IOVS

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SUPPLEMENTS AND COMPETITIVE ATHLETES

- Advise athletes to verify any supplements to assure that the ingredients are certified as acceptable for sports competition regulations.
- I reference the NSF International website:
 - <http://info.nsf.org/Certified/BannedSub/listings.asp>
 - Products will say: *NSF Certified for Sport®*

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TIMING OF SUPPLEMENTATION BENEFITS

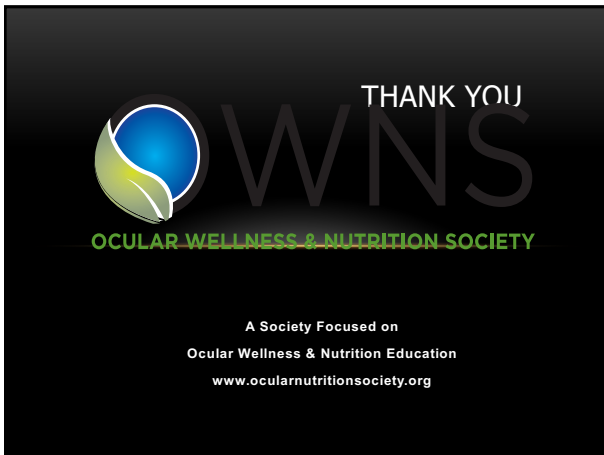
- **Protects Vision: Immediately**
- **Vision Quality Benefits: 1-2 months**
- **Visual Processing Speed Benefits: 3-4 months**

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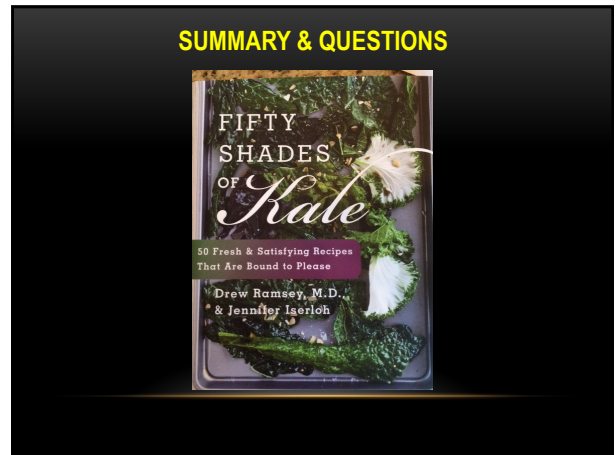
CHALLENGES OF NUTRIENT SUPPLEMENT STUDIES

- Nutrient trials are not pharmaceutical trials because all subjects have been exposed to the nutrients
 - No real "placebo" group since everyone ingests nutrients
 - A true RCT would require subjects that are nutritionally deficient
 - Cannot adequately control for daily nutrient intake
 - Nutrients are pan-systemic vs. system targeted
 - As a disease/condition prevention, the sample size to show effect would have to be extremely large

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