SUNLIGHT



FOR HEALTH

Michael F. Holick

Boston University Medical Center

Publics' View of the Sun

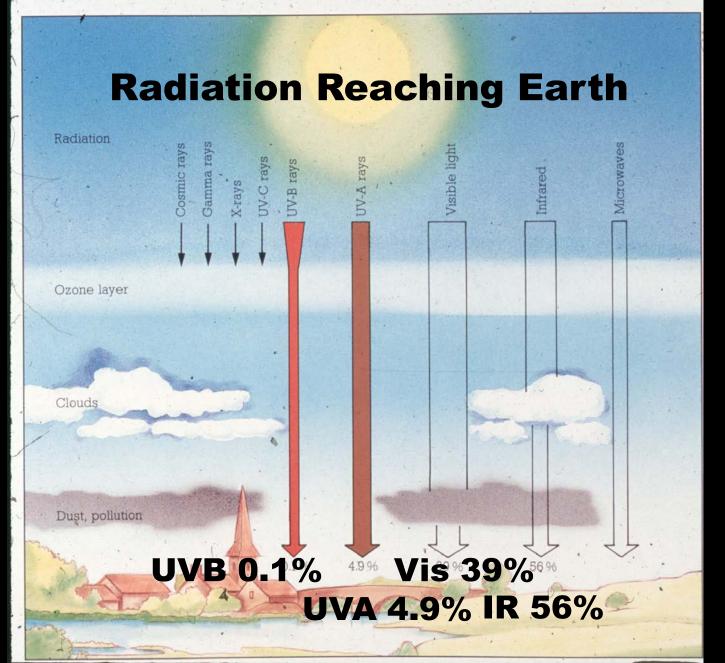


EDUCATION

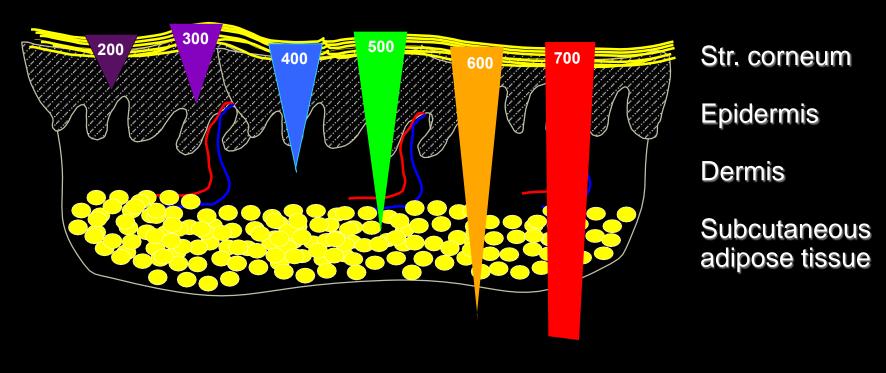
EDUCATION

EDUCATION

Schematic Representation of Electromagnetic Radiations Reaching the Earth Surface



Dependence of wavelength and skin penetration

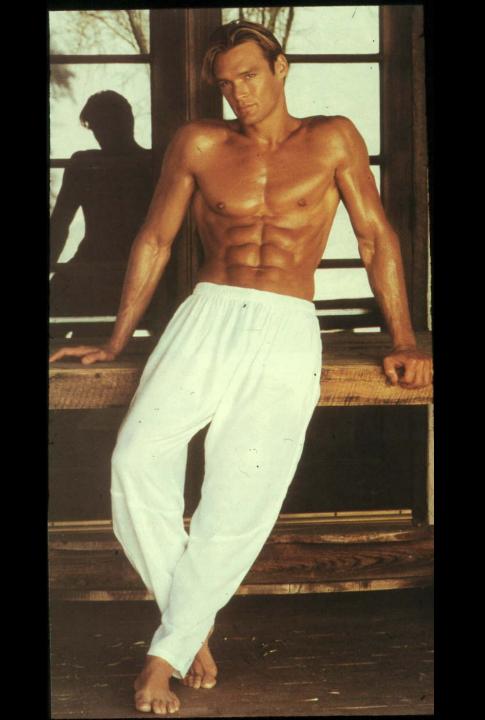


ENERGY NANOMETERS

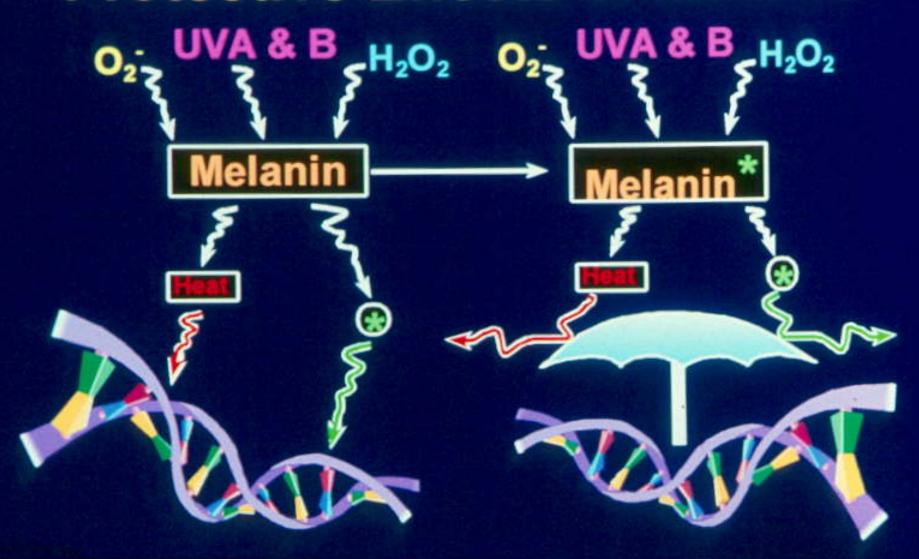
HOW DOESTHESKIN



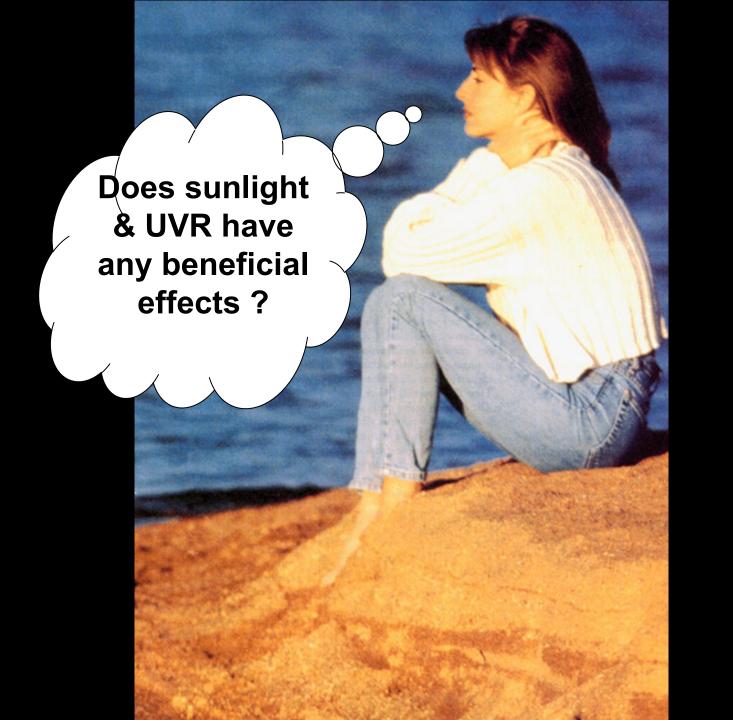
Suntanned hammerheads



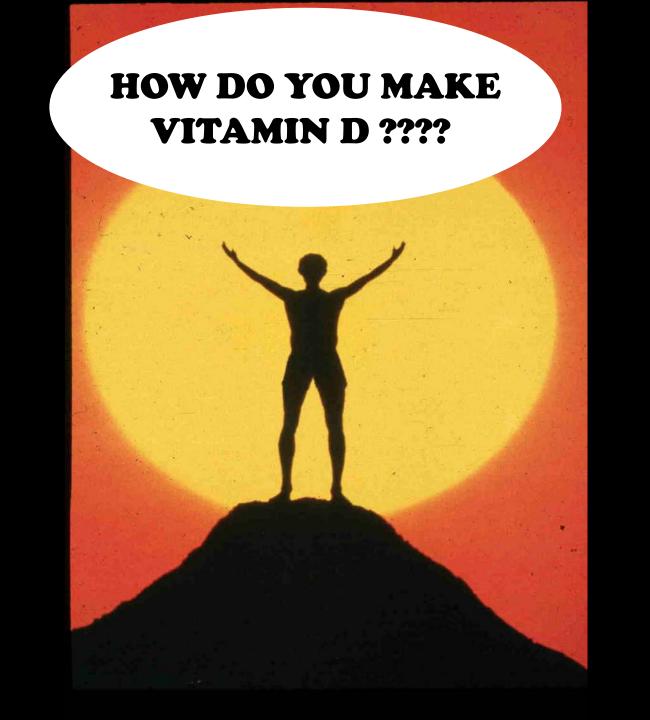
Protective Effects of Melanin



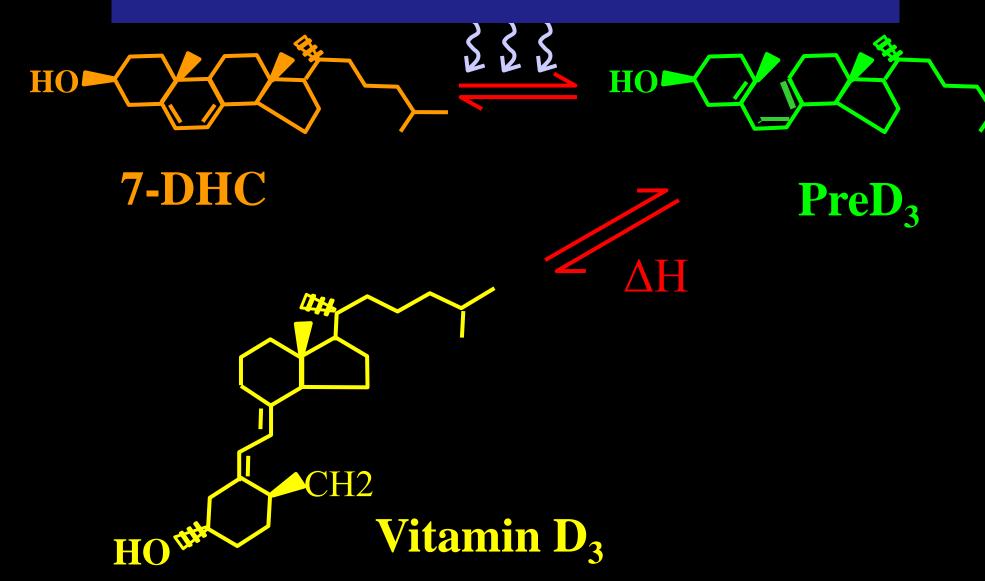


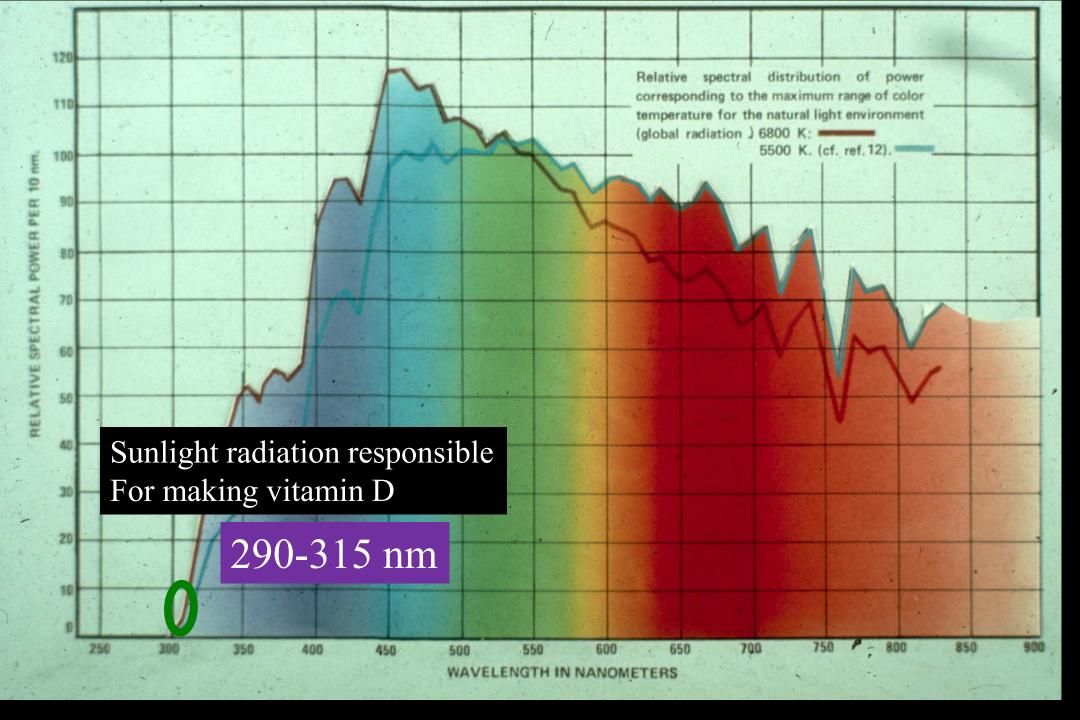






Solar UV Radiation





HOW MUCH

SUN

Total Body





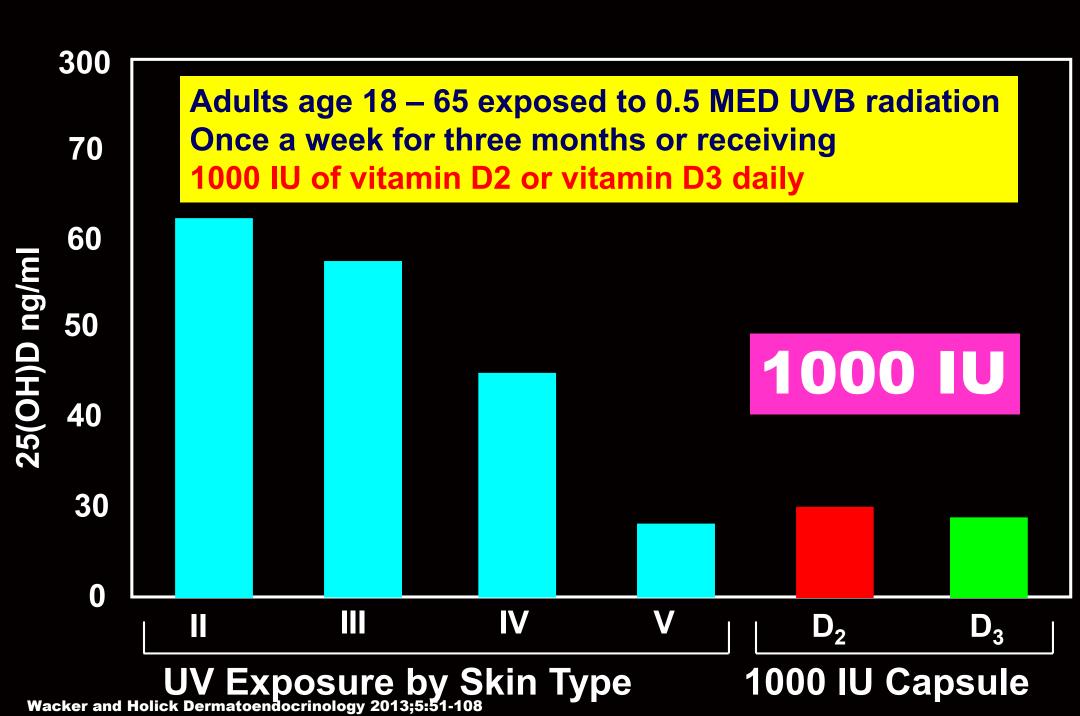
Exposure to

1 Minimum Erythemal

Dose

~20,000 IU Vitamin D₃

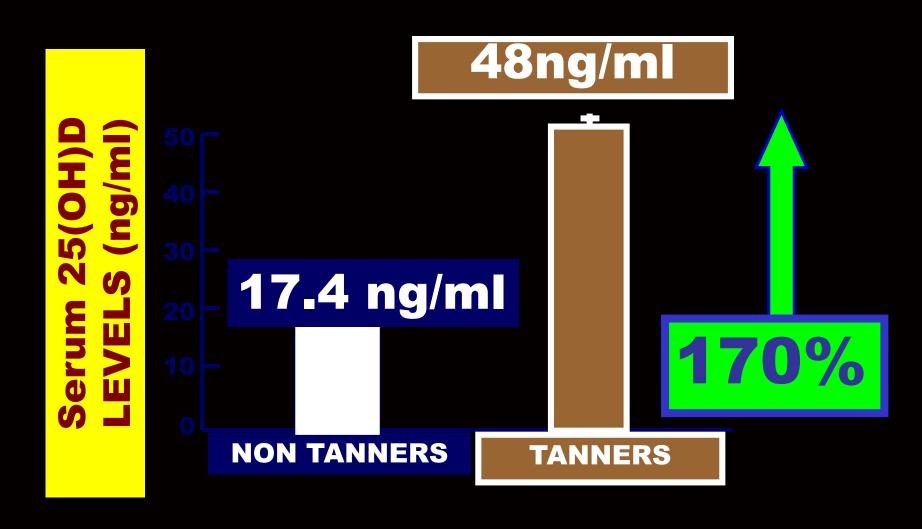
(RDA 600-800 IU)





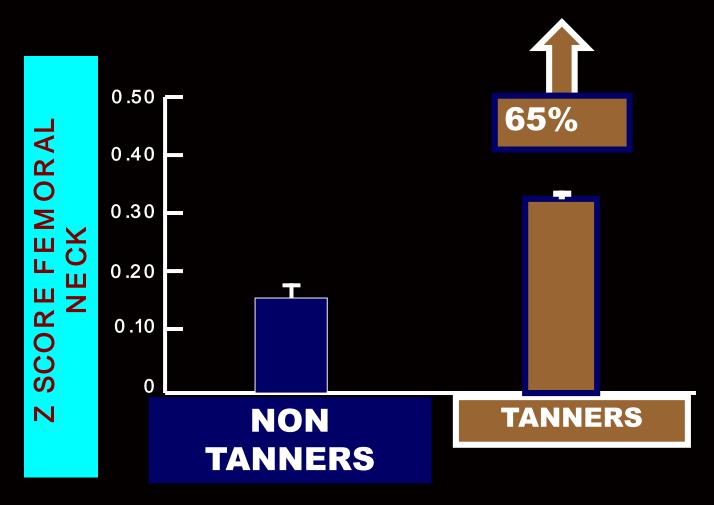


SERUM 25(OH)D LEVELS IN TANNERS AND NONTANNERS END OF WINTER IN BOSTON



Tangpricha, et al Tanning is associated with optimal vitamin D status (serum 25-hydroxyvitamin D concentration) and higher bone mineral density. Am J Clin Nutr. 2004. 80:1645-1649.

Z-SCORE FEMORAL NECK TANNERS VS NONTANNERS



Tangpricha, et al Tanning is associated with optimal vitamin D status (serum 25-hydroxyvitamin D concentration) and higher bone mineral density. Am J Clin Nutr. 2004. 80:1645-1649.



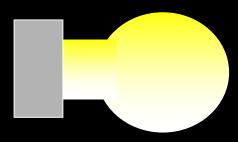
Why do People Feel Good in Sunlight

??????

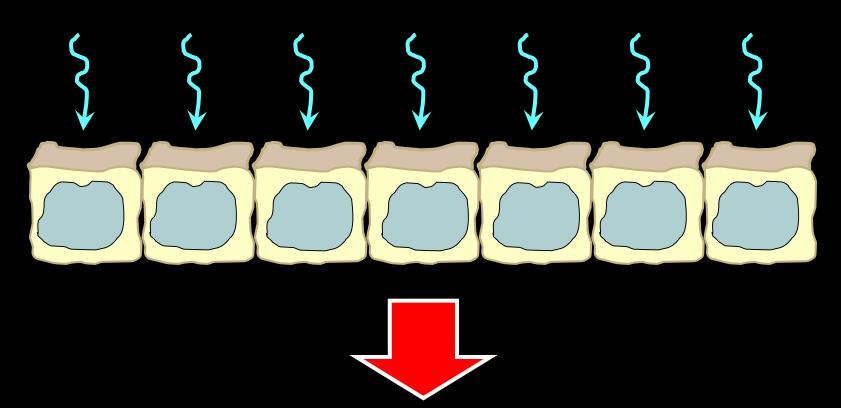
Do Human Skin Cells produce a Happy Substance in response to sunlight

??????

Does Human Skin Produce **B-Endorphin** 77777

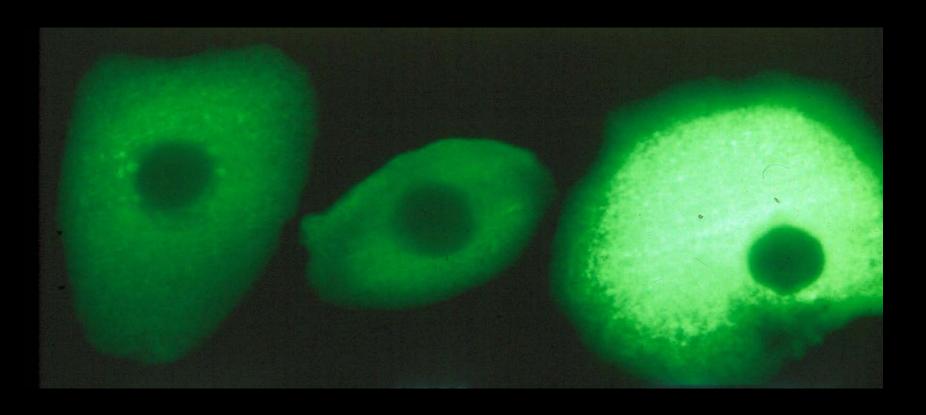


Simulated Sunlight



measure β-endorphin

β-Endorphin Induced by UVB in Cultured Human Keratinocytes



Control

UVA

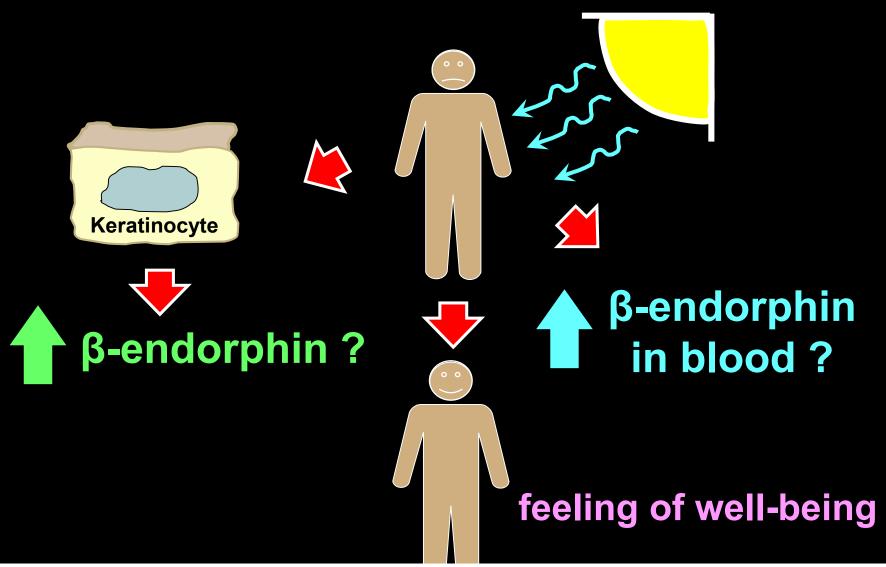
UVA + UVB

. Zanello, S., Jackson, D., and Holick, M.F. An immunocytochemical approach to the study of beta-endorphin production in human keratinocytes using confocal microscopy. *Ann NY Acad Sci.* 885:85-99, 1999.

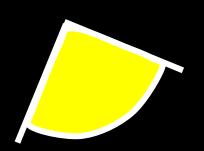
Does

Human Skin Produced β-Endorphin enter the bloodstream to make you Feel Good ??????

Sunlight - β-endorphin – Skin Connection

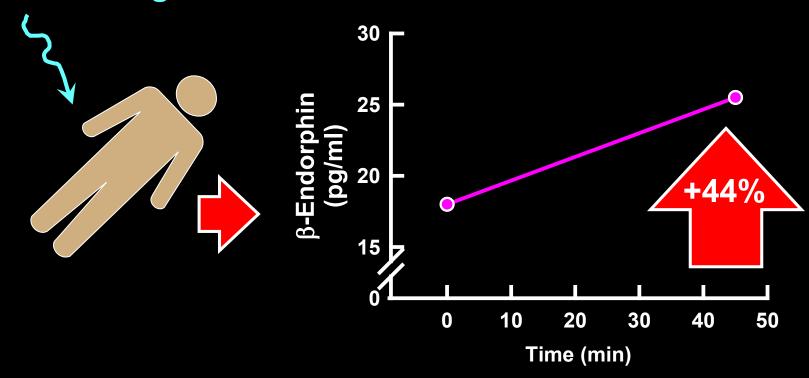


Jussila A, Huotari-Orava R, Ylianttila L, Partonen T, Snellman E. Narrowband ultraviolet radiation induces the expression of β-endorphin in human skin in vivo. *Journal of Photochemistry and Photobiology B:Biology*. 155:104-08, 2016.



Sunlight Exposure May Elevate Serum \(\beta \text{-endorphin} \) Levels in Humans

Simulated Sunlight





β-endorphin: opioid peptide





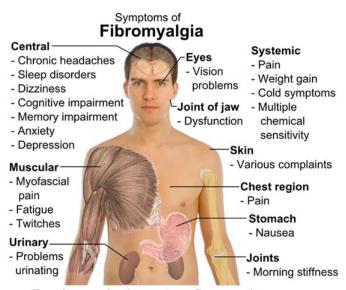
β-endorphins responsible for:

- ✓ Pain relief
- ✓ Feeling of well-being
- ✓ Relaxation

β-endorphin has approximately 33 times the analgesic potency of morphine.

Endorphins: Natural Pain and Stress Fighters





THE JOURNAL OF ALTERNATIVE AND COMPLEMENTARY MEDICINE Volume 15, Number 1, 2009, pp. 15–23
© Mary Ann Liebert, Inc.
DOI: 10.1089/acm.2008.0167

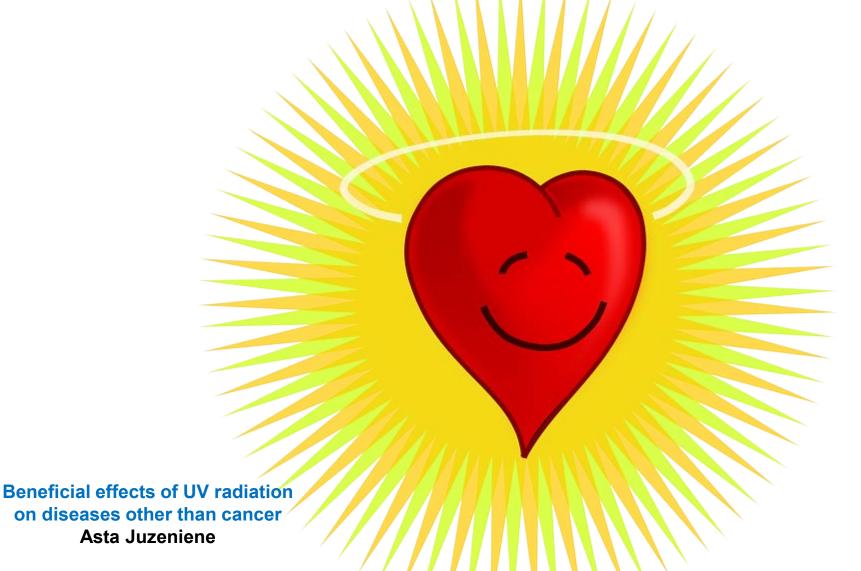
Pilot Study of the Effect of Ultraviolet Light on Pain and Mood in Fibromyalgia Syndrome

Sarah L. Taylor, M.D., M.P.H., Mandeep Kaur, M.B.B.S., M.S., Kristen LoSicco, B.S., Joy Willard, R.N., Fabian Camacho, M.S., Kenneth S. O'Rourke, M.D., and Steven R. Feldman, M.D., Ph.D. 1,2,4

Female reproductive system - Dysmenorrhea

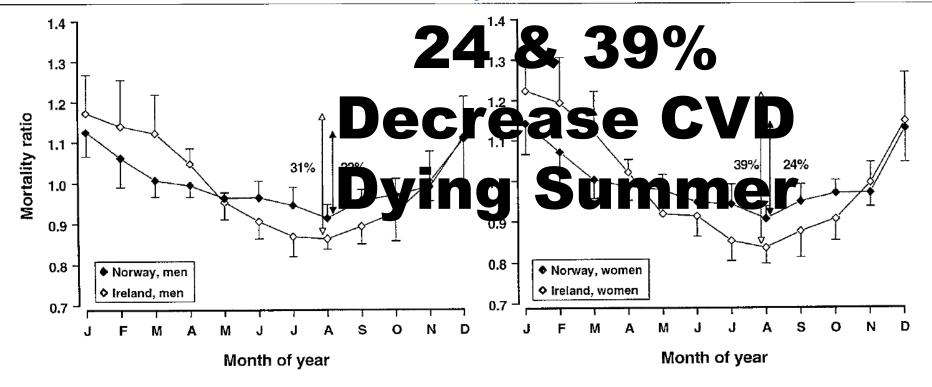
UV may have some potential in reducing chronic pain and improving mood in persons with Fibromyalgia syndrome.

Ultraviolet radiation and cardiovascular disease (CVD)



Seasonal variations in mortality caused by CVDs

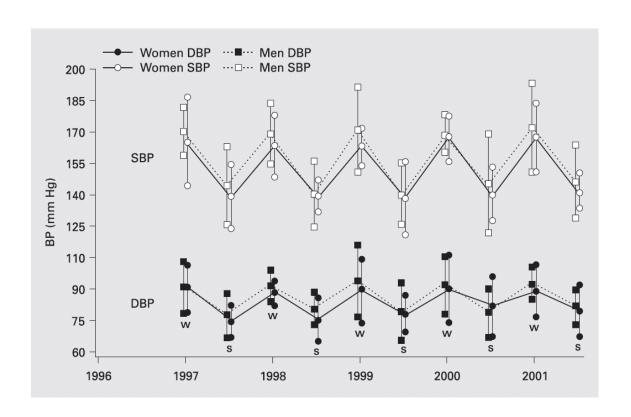




Mean monthly CVD mortality ratio (ICD 9: 390-459) in Norway and Ireland for men (left) and women (right) in the age group 60 years and older for the 10-year period 1985-1994. The figures beside the arrows indicate the percentage difference between the lowest summer mortality and the highest winter mortality.

The seasonal variation in CVDs mortality, with an excess in winter, has been documented in many studies.

Seasonal variations in blood pressure



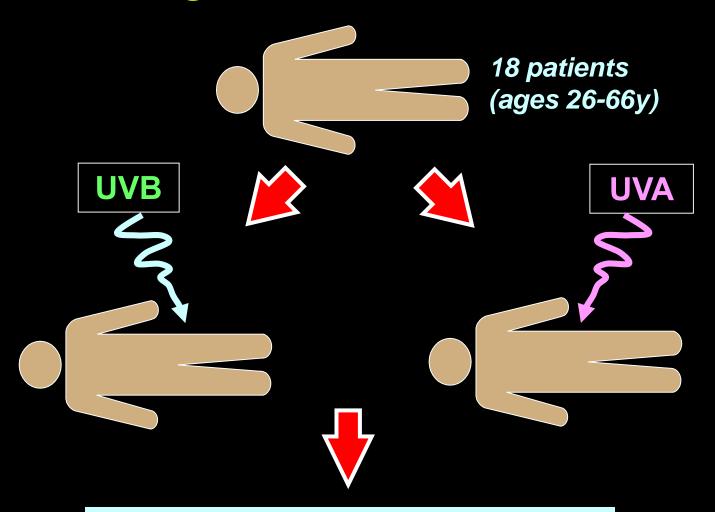
There is a tendency for both systolic (S) and diastolic (D) blood pressure (BP) to rise in winter.

Will UVB irradiation help my hypertension

?

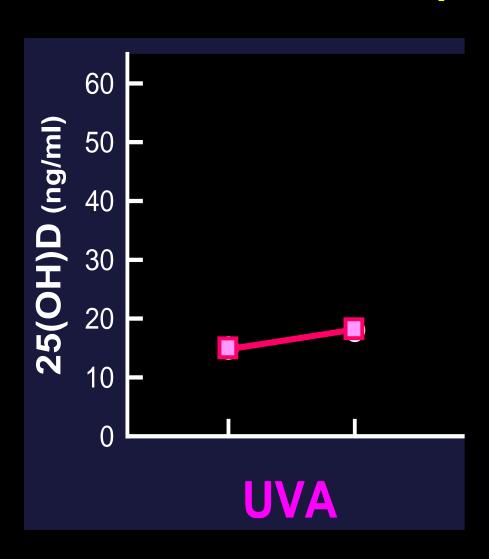


Hypertension

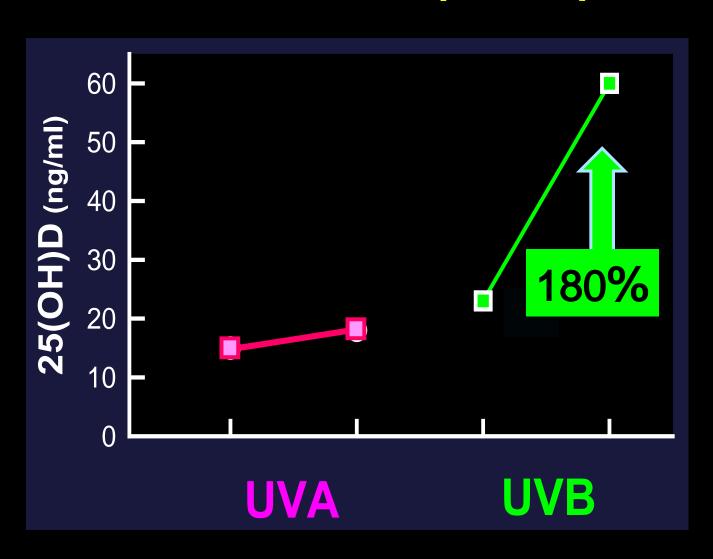


6 Weeks, measure BP

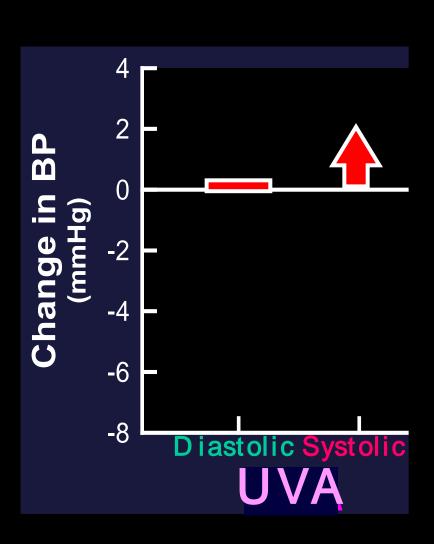
Serum 25(OH)D



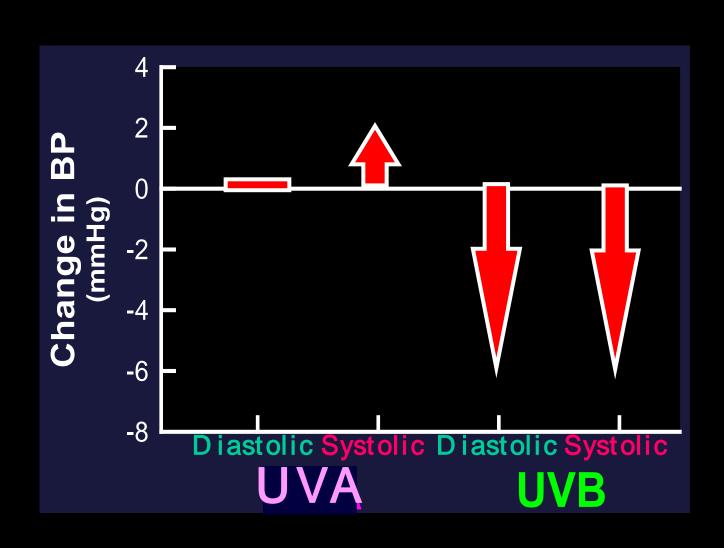
Serum 25(OH)D



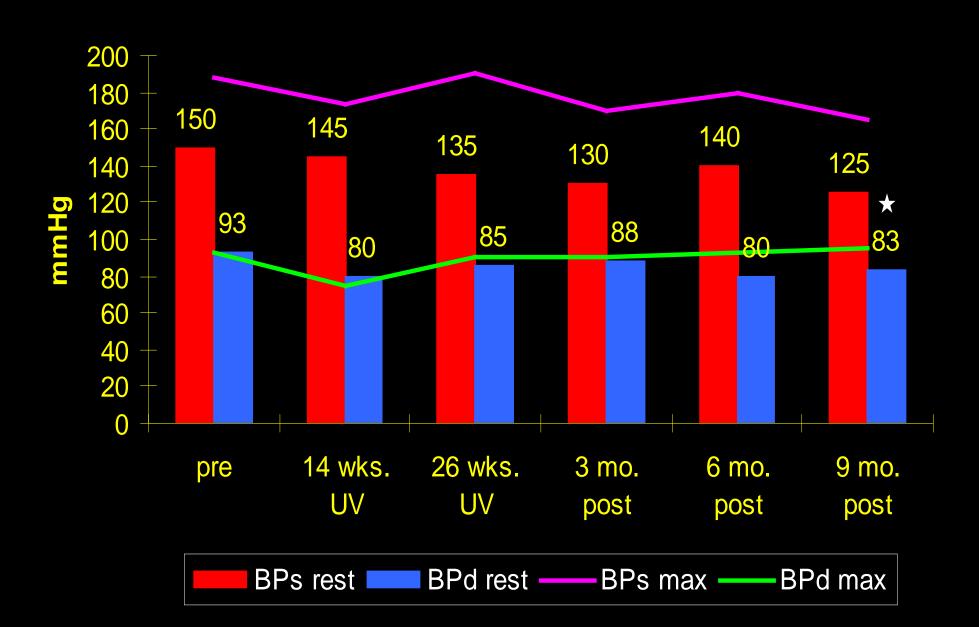
Effect of UVA on Blood Pressure



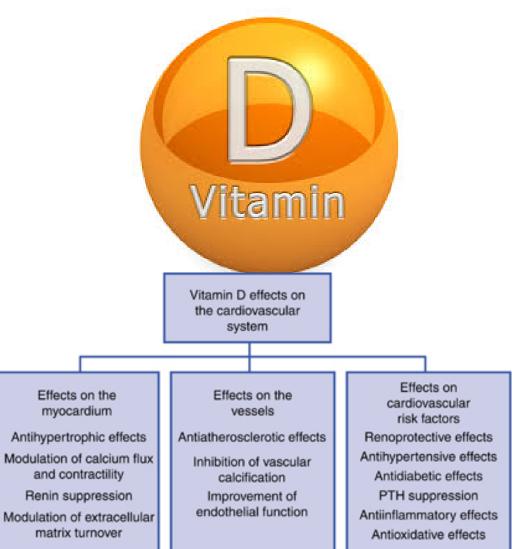
Effect of UVB & UVA on Blood Pressure

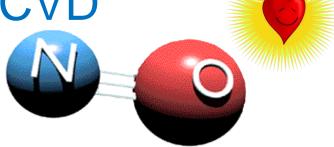


Blood Pressure



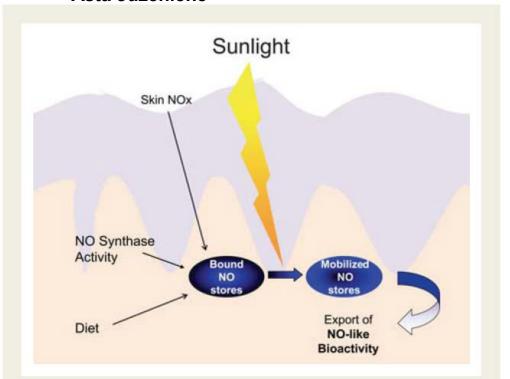
UV radiation and CVD

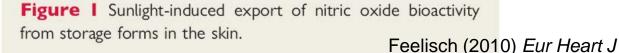


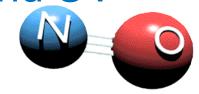


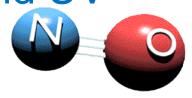
Beneficial effects of UV radiation on diseases other than cancer Nitric oxide (NO) and UV

Asta Juzeniene









$\overline{NO_2}$	\xrightarrow{hv}	NO.	+ RO	S/RNS
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	Species	Nitrite (μM)
Plasma	Human	0.12 ± 0.01
Plasma	Human	0.12
Plasma	Human	0.2 ± 0.02
Erythrocytes	Human	0.29 ± 0.05
Whole blood	Human	$0.18 \pm .02$
Saliva	Human	104
Skin tissue	Human	8.4
Skin tissue	Human	5.1
Sweat	Human	8.7
Sweat	Human	3.4

Suschek (2010) Nitric Oxide

Human skin is capable of releasing nitric oxide (NO) in an enzyme independent manner. This is due to photolysis of nitric oxide stores by UV radiation.

Liu D et al. UVA irradiation of human skin vasodilates arterial vasculature and lowers blood pressure independently of nitric oxide. Synthase. J Int. Derm. 143: 1839-1846, 2014.



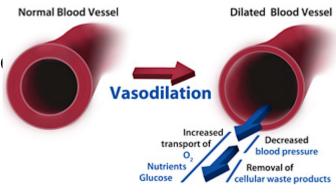
What is the role of nitric oxide (NO)?

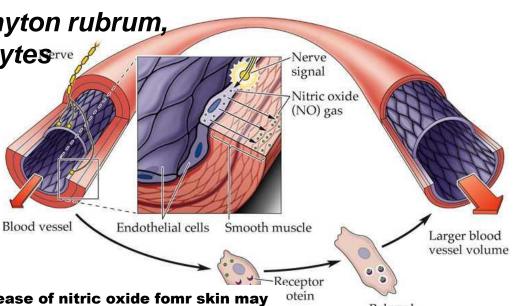


- Vasodilatation
- **✓ UV-induced melanogenesis**
- ✓ UV-induced immunosuppression
- ✓ Inflammation
- ✓ Apoptosis
- Wound healing
- ✓ Antimicrobial effects Bacteria – Staphylococcus aureus

Dermatophytes – Trichophyton rubrum, Trichophyton mentagrophytes –

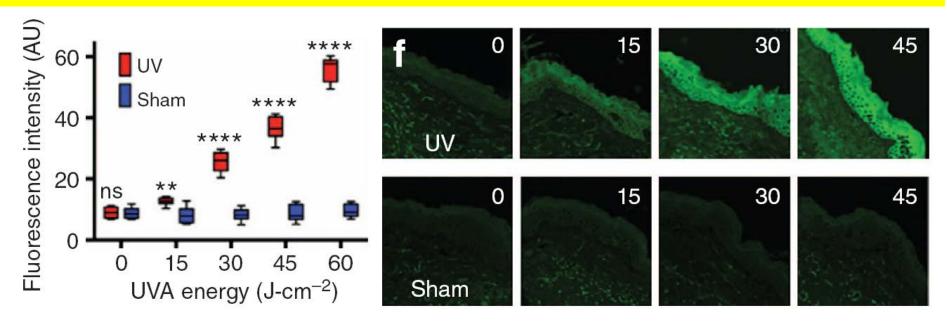
Yeasts - Candida albicans





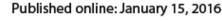
UVA liberates nitric oxide (NO) from the epidermis





Confocal fluorescence microscopy studies of human skin prelabeled with the NO-imaging probe diaminofluorescein 2 diacetate revealed that **UVA-induced NO release** occurs in a NOS-independent, dose-dependent manner, with the majority of the light-sensitive NO pool **in the upper epidermis**.

Liu D et al. UVA irradiation of human skin vasodilates arterial vasculature and lowers blood pressure independently of nitric oxide. Synthase. *J Int. Derm.* 143: 1839-1846, 2014.





Blood Purif 2016;41:130–134 DOI: 10.1159/000441266

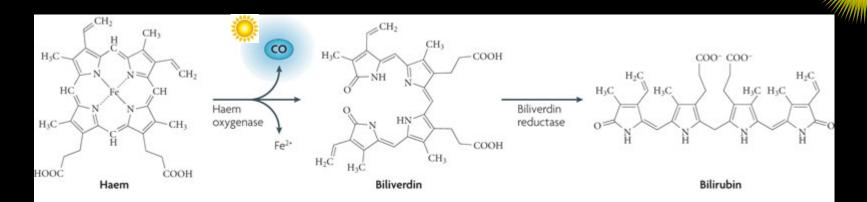
Sunlight Has Cardiovascular Benefits Independently of Vitamin D

Richard B. Weller

Medical Research Council Centre for Inflammation Research, University of Edinburgh, Queen's Medical Research Institute, Edinburgh, UK

risk factors independently of vitamin D. **Key Messages:** All-cause mortality should be the primary determinant of public health messages. Sunlight is a risk factor for skin cancer, but sun avoidance may carry more of a cost than benefit for overall good health.

Carbon monoxide (CO) and UV

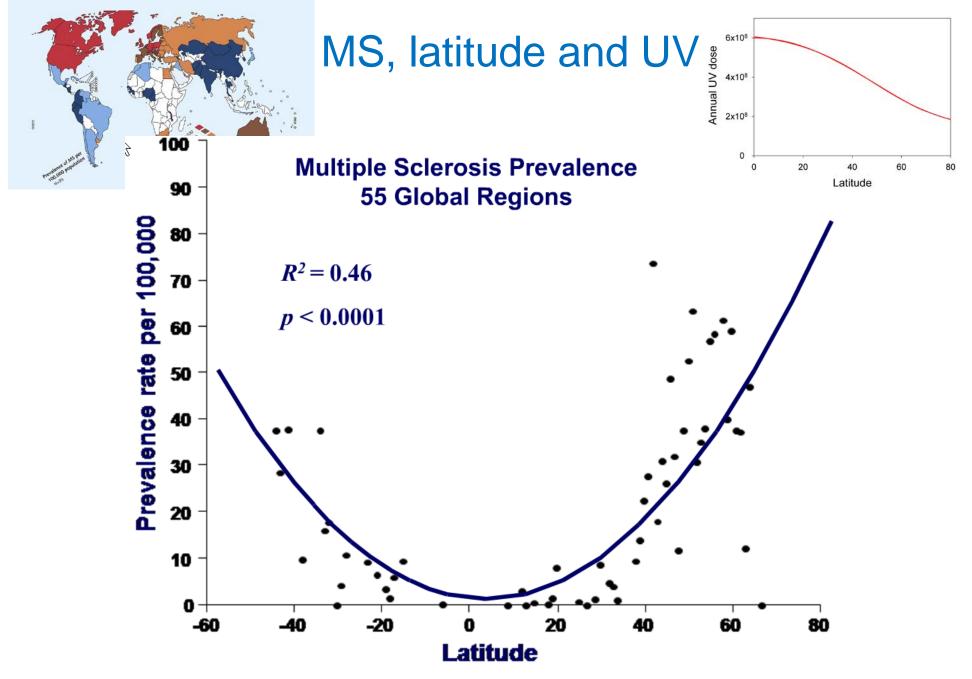


ig Discover

- ✓ Vasodilation
- Neurotransmitter
- Cell signaling
- Relaxation
- Antioxidant
- Anti-inflammation
- Ani-apoptosis

How Does UVR Effect my IMMUNE SYSTEM?

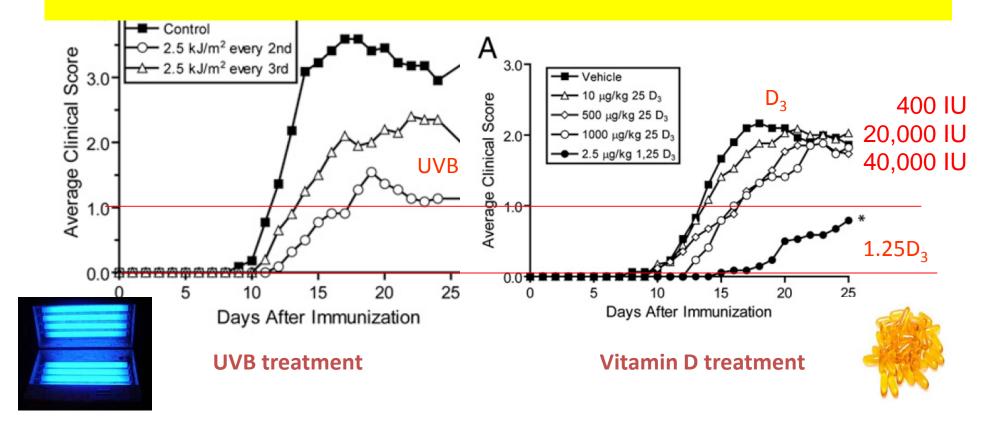




Kurtzke J F. Geographic distribution of multiple sclerosis: an update with special reference to Europe and the Mediterranean region *Acta Neurol. Scand.* 62 65–80, 1980.

MS animal model study:

UV more beneficial than vitamin D

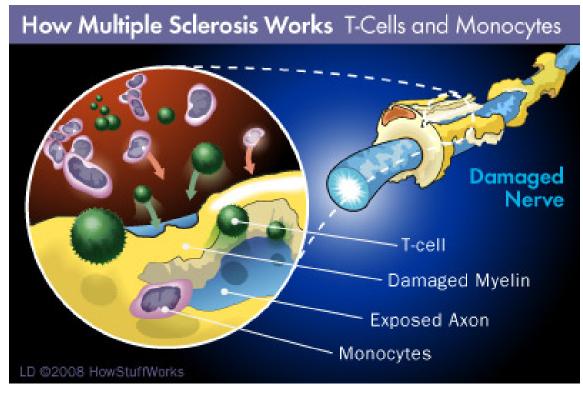


Continuous treatment with UVR dramatically suppresses clinical signs of experimental autoimmune encephalomyelitis (MS).



Multiple sclerosis





Effect of UVR:
Protection against disease

Development and reduces symptoms of MS

Down-regulation of T helper 1 activity involved in demyelination through T regulatory cells and/or B regulatory cells and/or immunosuppressive cytokines

Multiple sclerosis and adrenocorticotropic hormone (ACTH)

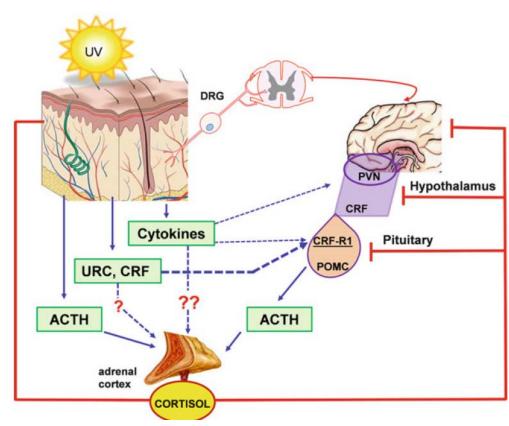


H.P. Acthar® Gel

- Highly purified preparation of adrenocorticotropic hormone (ACTH)
- Key approved indications:
 - Multiple sclerosis (MS) exacerbations
 - Nephrotic syndrome
- Significant off-label usage



A five-day course of intramuscular or subcutaneous ACTH gel improves symptoms associated with acute exacerbations of multiple sclerosis.



Slominski *et al.* (2012) Embryology and Cell Biology

How about UV phototherapy for MS?

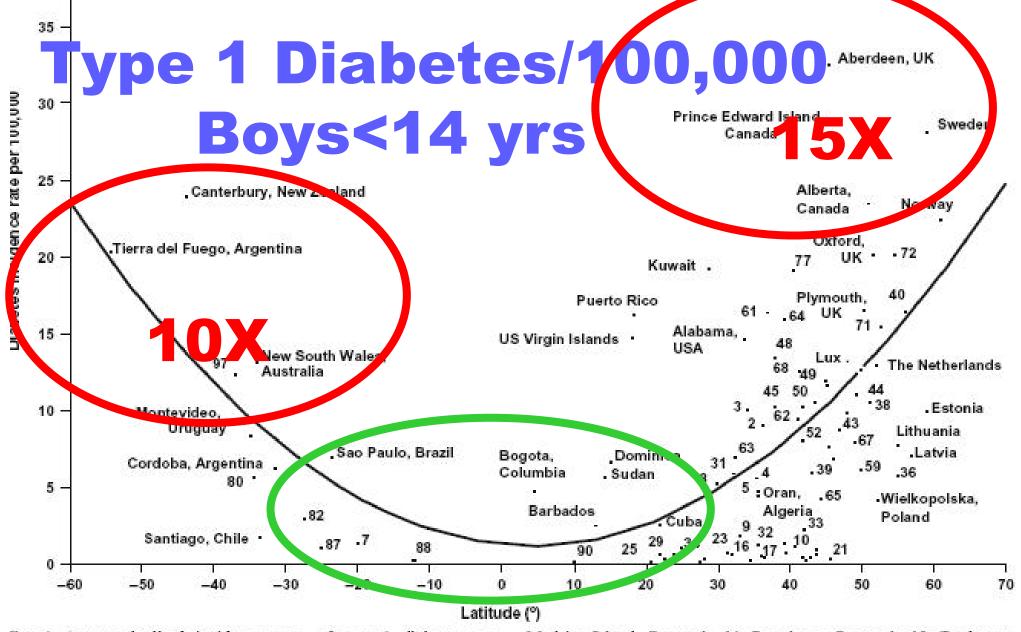


Fig. 1 Age-standardised incidence rates of type 1 diabetes per 100,000 boys <14 years of age, by latitude, in 51 regions worldwide,

Madeira Island, Portugal; 64. Portalegre, Portugal; 65. Bucharest, Romania; 67. Slovakia; 68. Catalonia, Spain; 71. Leicestershire, UK;

Mohr SB, Garland CF, Gorham ED, Garland FC. The association between ultraviolet B irradiance, vitamin D status and incidence rates of type 1 diabetes in 51 regions worldwide. *Diabetologia*. 51:1391-8, 2008.

laneda,

Ramune Jacobsen*, Peder Frederiksen and Berit L. Heitmann

Exposure to sunshine early in life prevented development of type 1 diabetes in Danish boys

Methods: The study population included 331,623 individuals born in Denmark from 1983 to 1988; 886 (0.26%) developed T1D by the age of 15 years. The values of sunshine hours were obtained from the Danish Meteorologi-

40 % Reduction More Sun third trimester

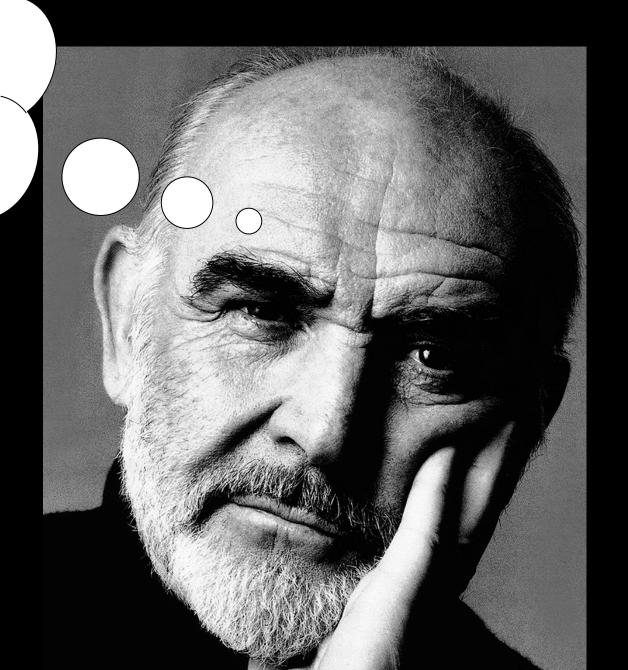
linear variable then was split into two categories separated by the median value.

Results and Conclusions: Cox regression models showed that more sunshine during the third gestational trimester was associated with lower hazards (HR) of T1D at age 5–9 years in males: HR (95% CI): 0.60 (0.43–0.84), p=0.003. Our results should be considered in the context of evidence-based recommendations to the public about skin protection from the sun.

"Der Zauberberg" (Thomas Mann 1924)

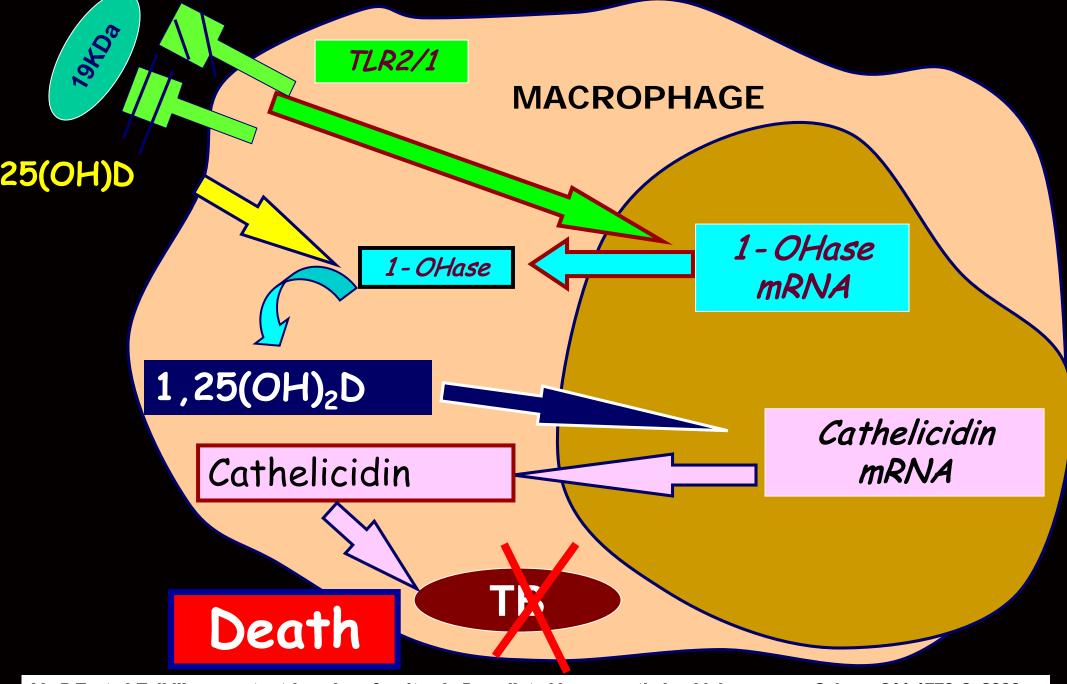


WHAT IS
THE EFFECT OF
VITAMIN D ON
THE
IMMUNE SYSTEM
????????



Vitamin D Protects Against Tuberculosis

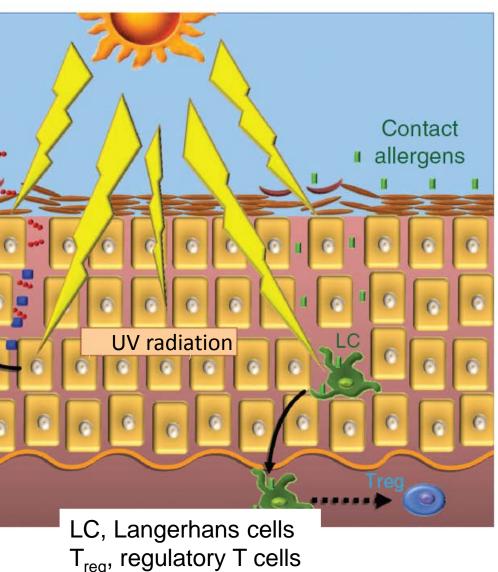
02.23.06, 12:00 AM ET



UV Effects on Other Diseases (T_{req} cells ↑)

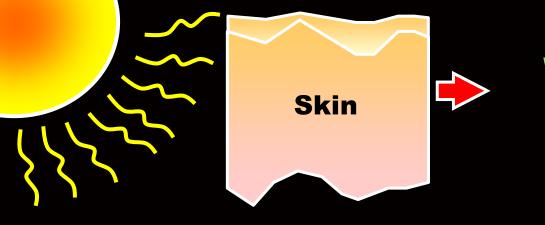
Beneficial effects of UV radiation on diseases other than cancer

Asta Juzeniene



UVR increases the number of regulatory T cells and reduces immune responsiveness

- Suppression of allergy and asthma
- Suppression of chronic inflammatory diseases
- Prevention of autoimmune diseases (Type 1 Diabetes, Multiple sclerosis)

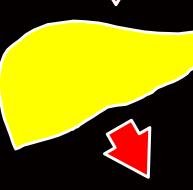


VITAMIN D

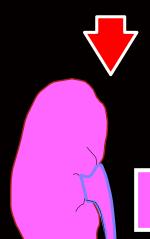


Liver 25-OHase

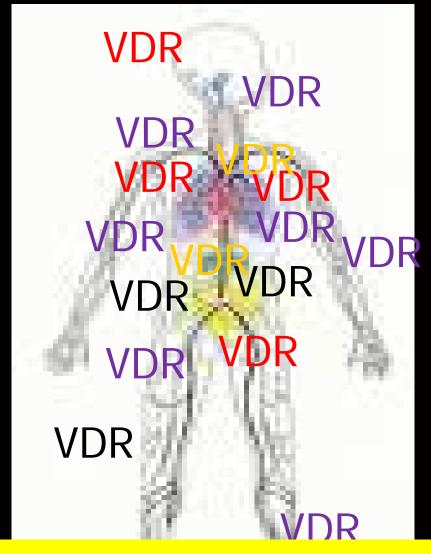
Kidney
1α-OHase



 $25(OH)D_3$



1,25(OH)₂D₃



ESSENTIALLY EVERY TISSUE AND CELL HAS A VDR

Vitamin D and Reduced Risk of Breast Cancer: A Population-Based Case-Control Study

Julia A. Knight,¹ Maia Lesosky,¹ Heidi Barnett,¹ Janet M. Raboud,¹ and Reinhold Vieth²

Table 2. ORs and 95% CIs for an association between vitamin D-related exposure variables at ages 10 to 19 (sun exposure and dietary vitamin D) and breast cancer in cases and controls

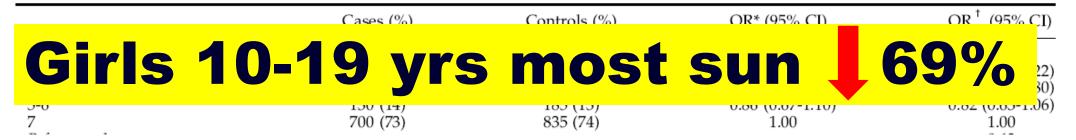


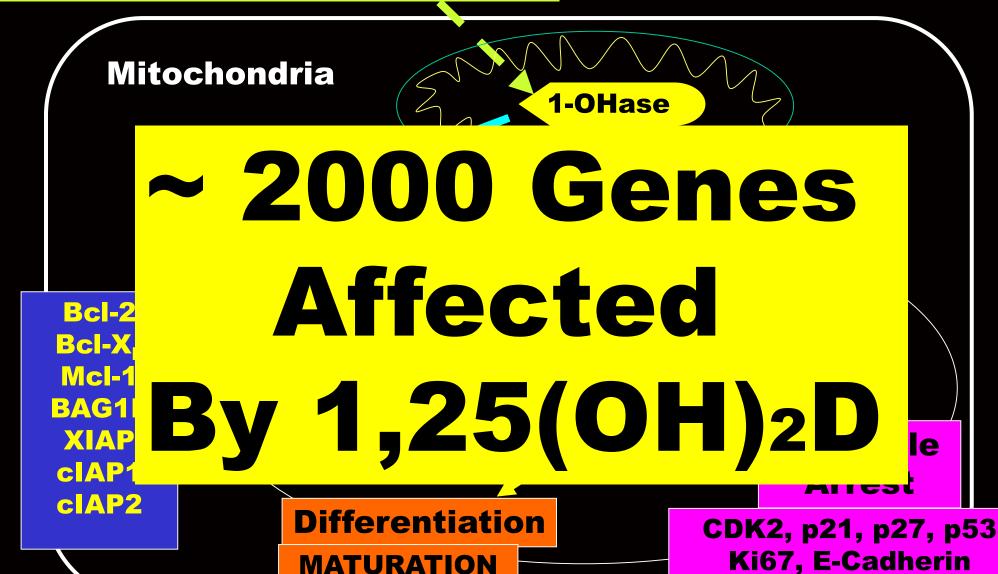
Table 3. ORs and 95% CIs for an association between vitamin D-related exposure variables at ages 20 to 29 (sun exposure and dietary vitamin D) and breast cancer in cases and controls



Women 45-54 yrs most sun 0.0%

25(OH)D>30 ng/ml

Colon Cell

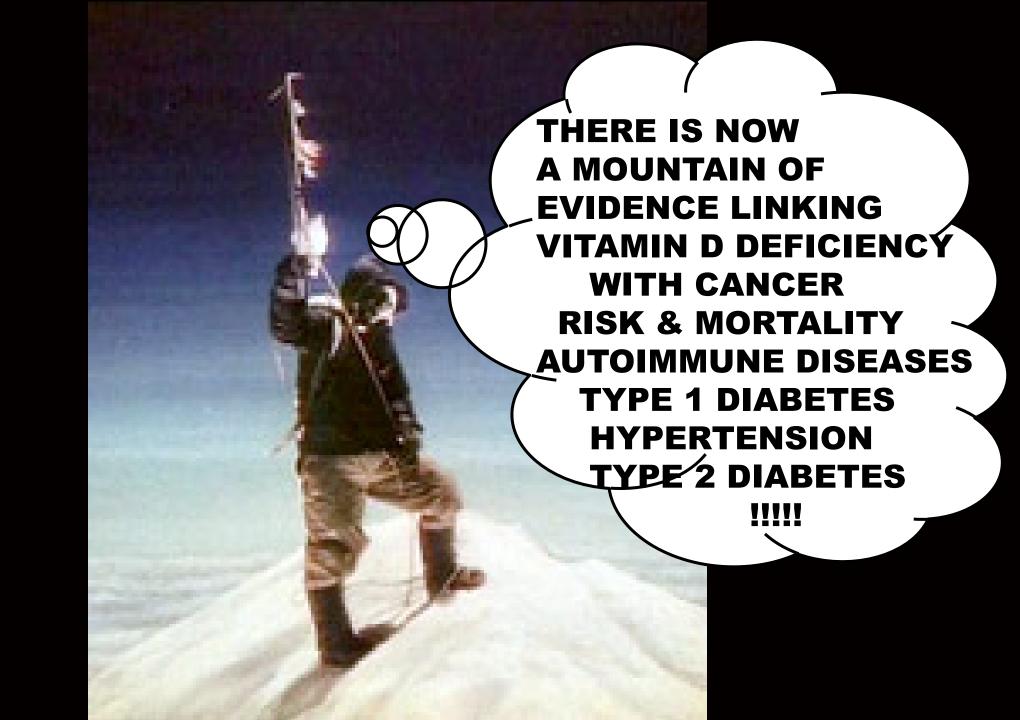


Hossein-nezhad A, Spira A, and Holick MF. Influence of Vitamin D Status and Vitamin D₃ Supplementation on Genome Wide Expression of White Blood Cells: A Randomized Double-Blind Clinical Trial. *PLoS ONE*. 8(3):e58725, 2013.

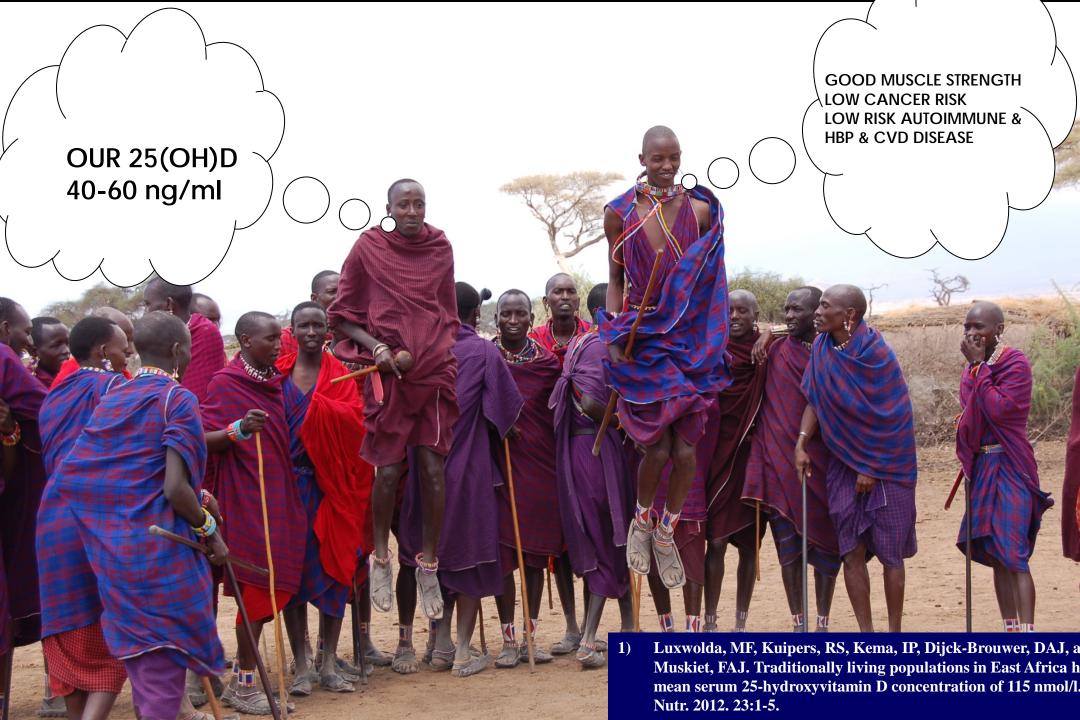
VITAMIN D

DEFICIENCY

IS A DISEASE OF NEGLECT



WHAT IS A ORNA **VERSUS** 25(OH)D LEVEL **7777777**



MICHAEL F. HOLICK, Ph.D., M.D.

Foreword by ANDREW WEIL, M.D.

THE VITAMIN D SOLUTION

A 3-Step Strategy to Cure Our Most Common Health Problems

PREVENT AND TREAT:

Osteoporosis • Heart Disease • Cancer • Autoimmune Diseases • Depression

• Insomnia • Arthritis • Diabetes • Chronic Pain • Psoriasis • Fibromyalgia •

Autism...as well as other diseases, chronic conditions, and mild ailments

The Influence of Painful Sunburns and Lifetime Sun Exposure on the Risk of Actinic Keratoses, Seborrheic Warts, Melanocytic Nevi, Atypical Nevi, and Skin Cancer

Cornelis Kennedy, Chris D. Bajdik,* Rein Willemze, Frank R. de Gruijl, and Jan N. Bouwes Bavinck, for the members of the Leiden Skin Cancer Study

Departments of Dermatology, Leiden University Medical Center, Leiden, The Netherlands; *British Columbia Cancer Agency, Vancouver, British Columbia, Canada

exposure was predominantly associated with an increased risk of squamous cell carcinoma (p-value for trend = 0.03) and actinic keratoses (p-value for trend < 0.0001) and to a lesser degree with the two types of basal cell carcinoma. By contrast, lifetime sun exposure



I should Avoid ALL Sunlight to Prevent Melanoma





The Influence of Painful Sunburns and Lifetime Sun Exposure on the Risk of Actinic Keratoses, Seborrheic Warts, Melanocytic Nevi, Atypical Nevi, and Skin Cancer

Cornelis Kennedy, Chris D. Bajdik,* Rein Willemze, Frank R. de Gruijl, and Jan N. Bouwes Bavinck, for the members of the Leiden Skin Cancer Study

Departments of Dermatology, Leiden University Medical Center, Leiden, The Netherlands; *British Columbia Cancer Agency, Vancouver, British Columbia, Canada

By contrast, lifetime sun exposure appeared to be associated with a lower risk of malignant melanoma, despite the fact that lifetime sun exposure did not diminish the number of melanocytic nevi or atypical nevi. Neither painful sunburns nor lifetime sun exposure were associated with an increased risk of seborrheic warts. Key words: actinic keratoses/atypical nevi/melanocytic nevi/seborrheic warts/skin cancer/ultraviolet light. J Invest Dermatol 120:1087–1093, 2003

WHAT DID THE AUTHORS CONCLUDE?

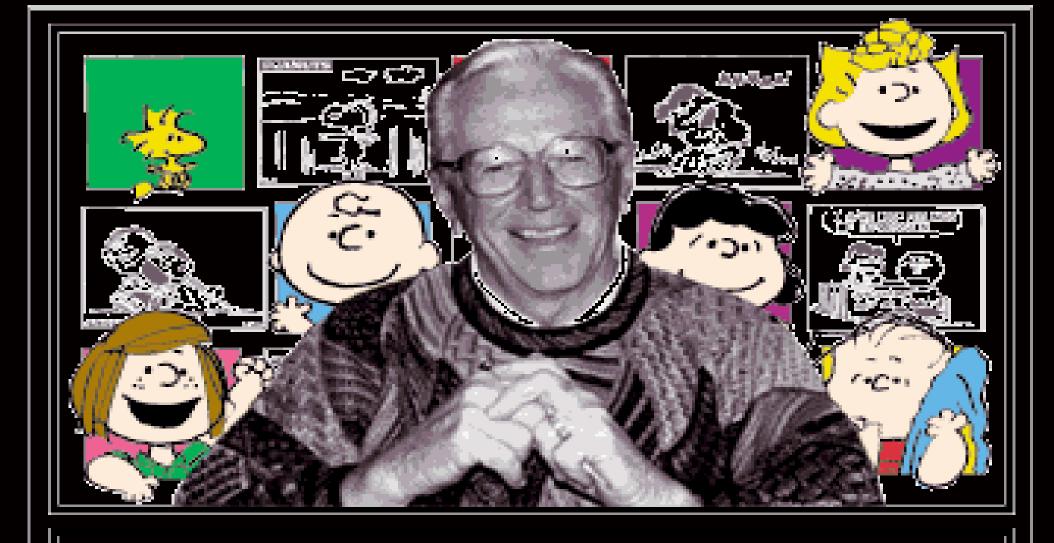
INCREASES RISK MELANOMA

INCREASED # MOLES

BAD GENETICS

RED HAIR COLOR

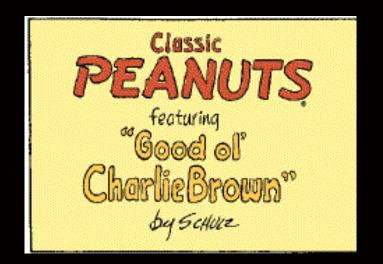
SUNBURNS as CHILD YOUNG ADULT

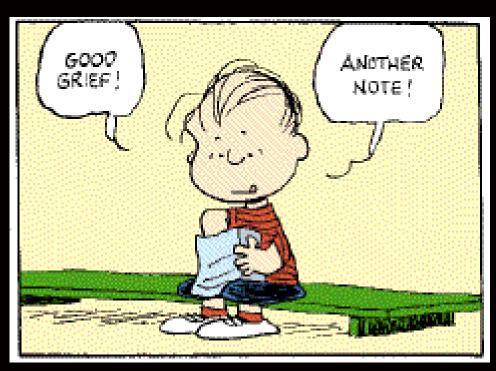


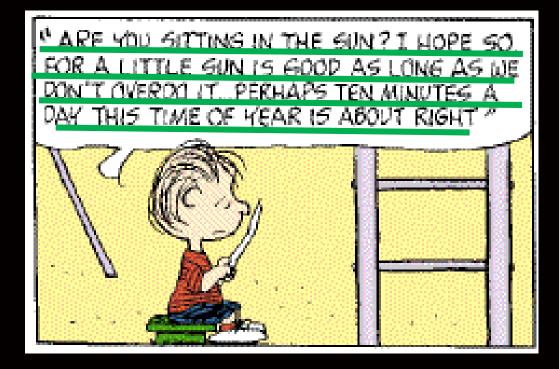
CHARLES M. SCHULZ

1922 - 2860

Them's Yeu For Civing The World Such Heppiness.







What is the Best Source of SENSIBLE

SUNLIGHT

WHEN CAN YOU MAKE VITAMIN D????

How about an App???

http://dminder.info

Lack of
Adequate Sunlight
&
Vitamin D
Deficiency
Has a
Major Impact
On Health



DrHolick.com

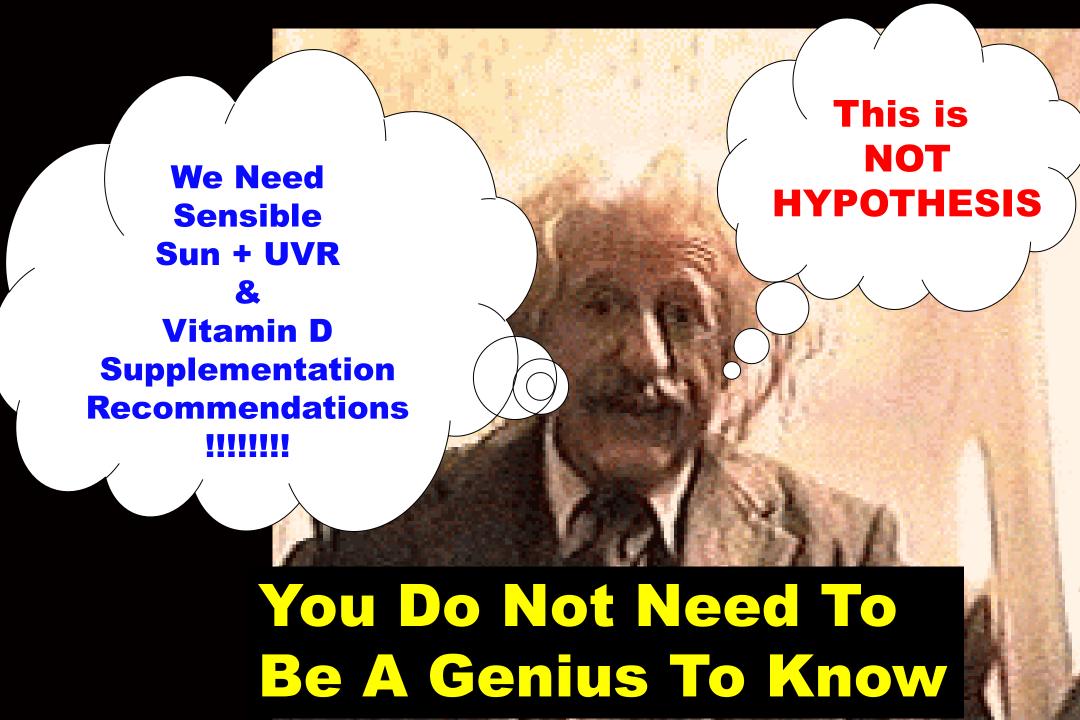
VITAMIN D DEFICIENCY CONSEQUENCES **CAUSES Schizophrenia Depression** SUNSCREEN **MELANIN INFECTIONS LATITUDE** URI WINTER TB **Autoimmune MEDICATIONS Type 1 Diabetes HBP** FEV₁ **Antiseizure** MS CHD Wheezing **Glucocorticoids** Crohn's illnesses Rifampin RA Hepatic Failure AODM **HAART Renal Failure** St John's Wart **Syndrome X CANCER** Nephrotic syndrome Colon **Breast Prostate** etc. **MALABSORPTION Muscle weakness** Crohn's Muscle aches Obesity Whipple's **Cystic Fibrosis** Celiac **Osteoarthritis Liver disease Osteoporosis Osteomalacia Rickets**

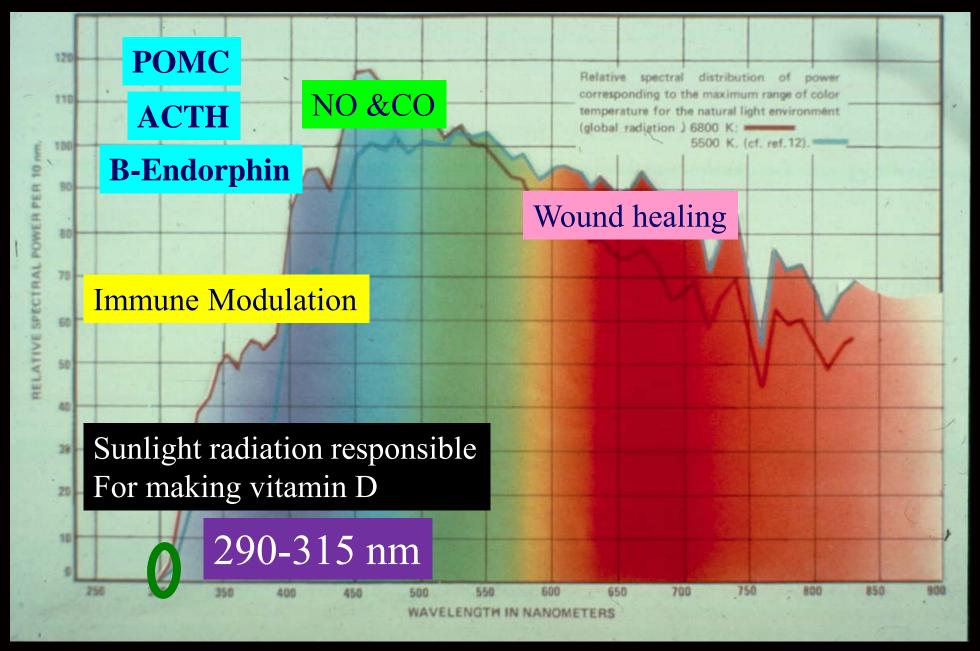
Holick M F. Vitamin D deficiency. New Engl. J. Med. 357 266-8, 2007.

There Is No
Downside
To
Obtaining
Sensible
Sun Exposure
!!!!



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World Health Organization

Knowing how to limit exposure to UV radiation is the key to a healthy and fun sun experience.

- Limit your UV risk when you are working outdoors!
- Enjoy your holiday without getting burnt by the Sun!



WHO INTERSUN the global UV project



Through INTERSUN, WHO provides scientific information and practical advice on the health impact and environmental effects of exposure to UV radiation.

INTERSUN collaborators:

- UNEP: United Nations Environmental Programme
- WMO: World Meteorological Organization
- IARC: International Agency for Research on Cancer
- ICNIRP: International Commission on Nonlonizing Radiation Protection.

Several WHO collaborating centres are also actively involved in INTERSUN.

More information on WHO INTERSUN can be found at: www.who.int/uv/intersunprogramme

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Sunshine and health

How to enjoy the Sun safely



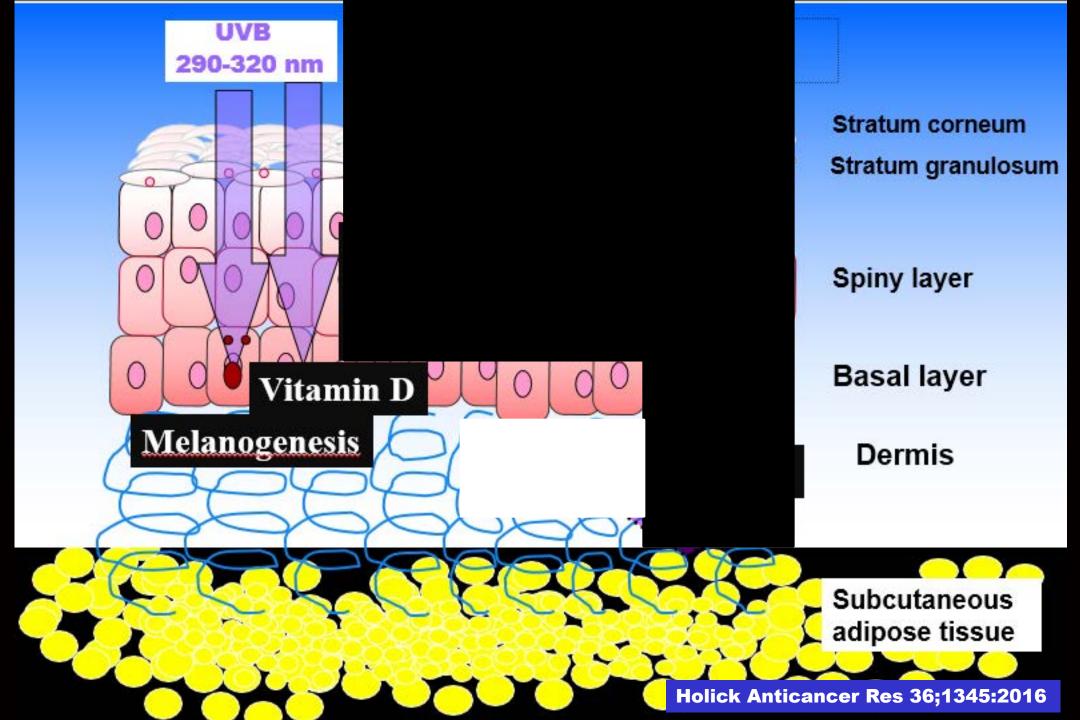


WHO Recomendations

The known health effects of UV

Are there beneficial effects of UV radiation?

The sun's rays provide warmth and light that enhance your general feeling of well-being and stimulate blood circulation. Some UV radiation is essential to the body as it stimulates the production of vitamin D. Vitamin D has an important function in increasing calcium and phosphorus absorption from food and plays a crucial role in skeletal development, immune function and blood cell formation. There is no doubt that a little sunlight is good for you! But 5 to 15 minutes of casual sun exposure of hands, face and arms two to three times a week during the summer months is sufficient to keep your vitamin D levels high. Closer to the equator, where UV levels are higher, even shorter periods of exposure suffice.





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