# If Low Calcium Is Not the Cause of Osteoporosis . . .

## What Is?

#### Rethinking the Nature and Causes of Osteoporosis

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#### Osteoporosis Statistics

54 million American adults over the age of 50 are affected by osteoporosis and osteopenia:

- 10+ million with osteoporosis By 2030: 43.4 million with osteopenia
  - 71.2 million is projected

The real "proof of the pudding" diagnosis of osteoporosis (fragile bones) is the occurrence of a low-trauma fracture.

1 in 2 women and up to 1 in 4 men over age 50 will break a bone due to osteoporosis.

### **Defining Osteoporosis**

Classification	T-score
Normal	-1.00 or greater
Osteopenia	Between -1.00 and -2.50
Osteoporosis	-2.50 or less
Severe/Established Osteoporosis	-2.50 or less and fragility fracture

#### Bone Mineral Density Does Not Predict Fracture Risk

- The US Study of Osteoporotic Fractures (SOF) looked at 8,065 women 65 and older.
- They reported that only 10 to 44% of osteoporotic fractures occurred in those women with an "osteoporotic bone density."
- European trials report that only 18% of all fractures occur in women with "osteoporotic" bone density.

#### So What Is a Better Definition of Osteoporosis?

In 1993, a U.S. Consensus Development Conference defined osteoporosis as:

"A systemic skeletal disease characterized by low bone mass and architectural deterioration of bone tissue with a consequent increase in bone fragility and susceptibility to fractures"

- Consensus Development Conference 1993

# RETHINKING THE NATURE OF OSTEOPOROSIS

#### Osteoporosis is not just . . .

- Thin bone or low bone density
- A disorder of the elderly
- A disorder of women
- Something "gone wrong" in the body





- Peak bone mass is achieved in our late 20s or early 30s.
- Bone loss begins shortly after that.
- There are various reasons for low bone mass in young folks.

## Osteoporosis Is Not Something "Gone Wrong" with Your Bones

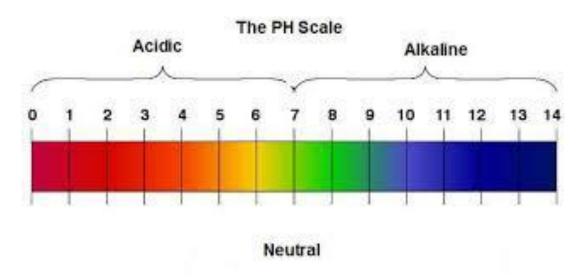




- Osteoporosis is our magical body's intelligent response to long-term imbalances and stressors.
- Bone is a gigantic reservoir of essential nutrients that can be made available to the body instantly.
- Osteoporosis is best seen as a positive coping mechanism by an intelligent biological system that is under stress.

• Osteoporosis is really just the end product "disorder" of our body's lifelong attempt to maintain a crucial internal "order."

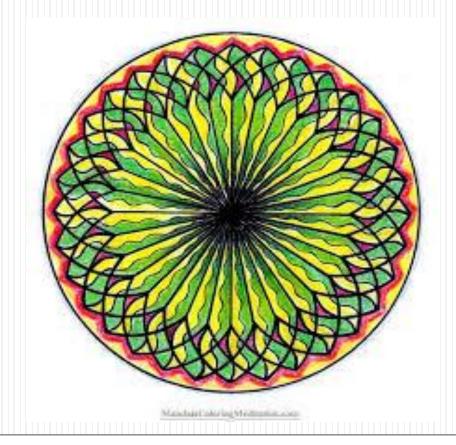




• It is a positive, life-supporting coping mechanism which allows the body to maintain internal balance.

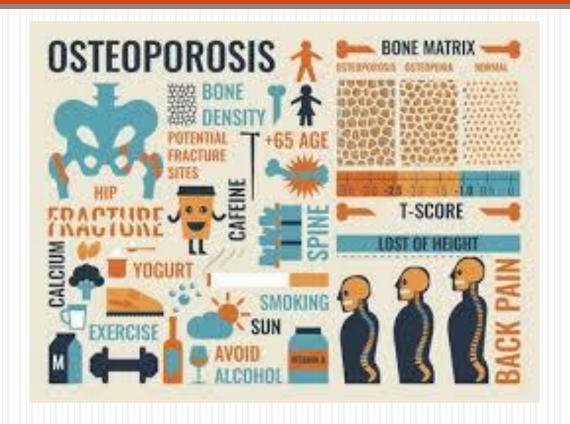
#### Osteoporosis Is a Complicated Disorder . . .

. . . that everyday medicine presents as simple.



#### Osteoporosis Does Not Stand Alone

- The body is one whole, interconnected unit.
- Fragile bones are not an isolated disorder.



#### Muscle and Bone

- As we lose muscle mass over time, we also lose bone mass.
- Back muscle strength has been shown to decrease 50% in women as they age from 50 to 80.
- The average woman loses 47% of her spinal bone mass during her lifetime, while most men lose 30%.

- Riggs et al. 1986; Sinaki 2003; Sinaki et al. 2002

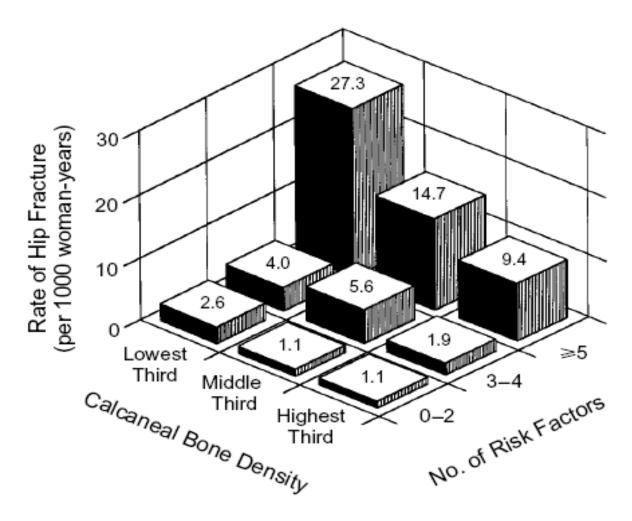
#### Aging and Muscle

- Back muscle strength has been shown to decrease 50% in women and 64% in men as they age from 50 to 80.
- Reduction in height with aging has also been reported.

- Sinaki et al. 2002

# RETHINKING THE CAUSES OF OSTEOPOROSIS





Annual Risk of Hip Fracture According to the Number of Risk Factors and the Age-Specific Calcaneal Bone

#### The Importance of Multiple Risk Factors

Hip fracture risk was 17 times greater among the 15 percent of the women who had five or more risk factors (exclusive of bone density) than the 47 percent of the women with two or fewer risk factors.

- Cummings et al. 1995

#### Muscle and bone wasters include:

- Sugar and sugar substitutes
- Refined carbs
- Processed vegetable oils
- Soda
- Excessive caffeine
- Smoking
- Medications
- Transfats
- Preservatives
- Excessive alcohol



## Stress-Induced Muscle and Bone Loss

Ann N F Acual Ser 2009 August 1172: 34-53, doi:10.1111 j.1749-6632.2009.04414.x.

#### Can meditation slow rate of cellular aging? Cognitive stress, mindfulness, and telomeres

Eiissa Epei, PhD. Jennifer Daubenmier, Ph.D. 1, Judith T. Moskowitz, Ph.D. 2, Susan Folkman, PnD. - and Elizabeth Blackburn, PhD. "LOSF Dect of Payon atry, 3888 California Street, Suite 468, San Francis in Inc. 1 FUCSE Decrit Mezione 2003 Calfornia Street Killin USF Catanment of a new



#### Relaxation Response Induces Temporal Transcriptome Changes in Energy Metabolism, Insulin Secretion and

Inflammatory Pathways

Manoj K. Bhasin<sup>1,4,5</sup>\*, Jeffery A. Dusek<sup>6</sup>\*, Bei-Hung Chang<sup>7,6</sup>\*, Marie G. Joseph<sup>5</sup>, John W. Denninger<sup>1,2</sup>, AFFECTIONS , FECCHIONS , FECTIONS , FUNDS O, LANCETTIONS , LONG STATE OF STATES OF APPRICA 2 Department of Psychiatry, Fertilland of Medicine Manachusetts Coursel Hospital Boston, Massachusetts. Unless States of America 2 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel Hospital States of America 3 Department of Medicine, Manachusetts Coursel States of America 3 Department of Medicine, Manachusetts Coursel States of America 3 Department of Medicine 3 Departm Gregory L. Fricchione 1.2, Herbert Benson 1.31, Towia A. Libermann 1.4.5.1 The state of the state of the state of States of America, & Mistante for Health, and Health, Alabors Hosthwesters Hostinial, Nammorphia, Missonskie, United States of America, a Turk of the state of the state hosticy and Management, Spisson University School. and departs under States of America

#### JBMR°

#### Neuropeptide Y Attenuates Stress-Induced Bone Loss Through Suppression of Noradrenaline Circuits

PA Baldock, 12.3 5 Lin, 1 L Zhang, 1 T Karl, 1.3 Y Shi, 1 F Driessler, 1.2 A Zengin, 1.2 B Hörmer, 2 NJ Lee, 1 IPL Wong, 12 EJD Lin, 1 RF Enriquez, 12 B Stehrer, 1 MJ During, 4 E Yulyaningsih, 1 S Zolotukhin, 3 ST Ruohonen, 6 E Savontaus, A Sainsbury, 1.7 and H Herzog 1.3

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Division of Cell and Molecular Therapy, University of Florida, Gainesville, FL, USA \*Department of Pharmacology, Drug Development and Therapeutics, University of Turku, Turku, Finland

<sup>7</sup>School of Medical Sciences, University of New South Wales, Sydney, Australia

Chronic stress and depression have adverse consequences on many organ systems, including the skeleton, but the mechanisms underlying stress-induced bone loss remain unclear. Here we demonstrate that neuropeptide Y (NPY), centrally and peripherally, 12-25 2 Critical role in protecting against stress-induced bone loss. Mice tacking the anxiolytic factor NPY exhibit more anxious THERE IS SHOULD THE BY DIVICENTING AGAINST SHEASY HIGHER LIGHT 1999, MILE HOLDING SHEADING AGAINST A CAMBRIDE AGAINST HIGH CHINDS THE SHEADING AGAINST A CAMBRIDGE AND A CAMBRIDGE CAMBRID The state of the superstant of the state of 12 TO THE PROPERTY OF THE PROP The second of th THE SECRET OF THE SECRET SECRE

#### How Many Hours Did You Sleep Last Night?



#### Happiness Matters

Older women who are satisfied with their lives have higher bone density and are less likely to develop osteoporosis than their unsatisfied peers, according to a study

in Finland.



– n.a. Nursing Standard 2015

### Highlights of Bone Depleting Factors

- Low Levels of Any of the 20 Key Bone Nutrients
- Chronic Low-Grade Metabolic Acidosis Acid-Base Balance
- Contemporary Life Style:
  - ✓ physical inactivity
  - ✓ emotional over-activity
  - ✓ medication use
  - ✓ medical disorders

## To Protect Your Bones, You Need at Least 20 Key Nutrients, Not Just Calcium



#### Nutrient Repletion: The Key Bone Nutrients

- Chromium
- Vitamin D
- Vitamin C
- Vitamin A
- Vitamin B<sub>6</sub>
- Vitamin B<sub>12</sub>
- Folate
- Vitamins  $K_1$  and  $K_2$
- Essential Fats
- Protein

- Calcium
- Phosphorus
- Magnesium
- Silicon
- Zinc
- Manganese
- Copper
- Boron
- Potassium
- Strontium

#### THE 20 KEY BONE BUILDING NUTRIENTS

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## Sub-Optimal Nutrient Intake Is the Norm, Not the Exception

- Magnesium: all ages, sexes, and classes (except those under 5) fail to consume even the RDA
- Zinc: average intake is 46 to 63% the RDA
- Potassium: average intake is around 50% RDA
- Copper: 75% of diets fail to contain the RDA
- Vitamin D: Low serum levels are common
- <u>Calcium</u>: Average intake is 500 to 600 mg

#### Nutrient Losses in Food (from 1950 to 1999)

Vegetable	Calcium	Iron	Phosphorus
Broccoli	<b>↓</b> 63%	<b>↓32</b> %	↓13%
Carrots	<b>↓</b> 31%	<b>↓</b> 37%	↓19%
Kale	<b>↓</b> 40%	<b>↓23</b> %	↓10%
Onions	<b>↓</b> 37%	<b>↓</b> 56%	<b>↓</b> 25%
Potatoes	<b>↓</b> 36%	<b>↓</b> 8%	↓18%

Fruit	Calcium	Iron	Vit. A		
Apples	None	<b>↓</b> 40%	↓41.1%		
Lemons	<b>↓</b> 57.%	↓14.3%	个3.3%		
Oranges	<b>↓</b> 2.4%	<b>↓</b> 75%	个2.5%		
Strawberries	<b>↓</b> 33.3%	<b>↓62%</b>	<b>↓</b> 67.1%		
Tangerines	<b>↓</b> 65%	<b>↓</b> 75%	个119%		

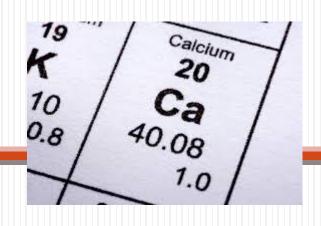
#### Repair Deficit

What keeps you from having perfect health?

Are you in repair deficit?

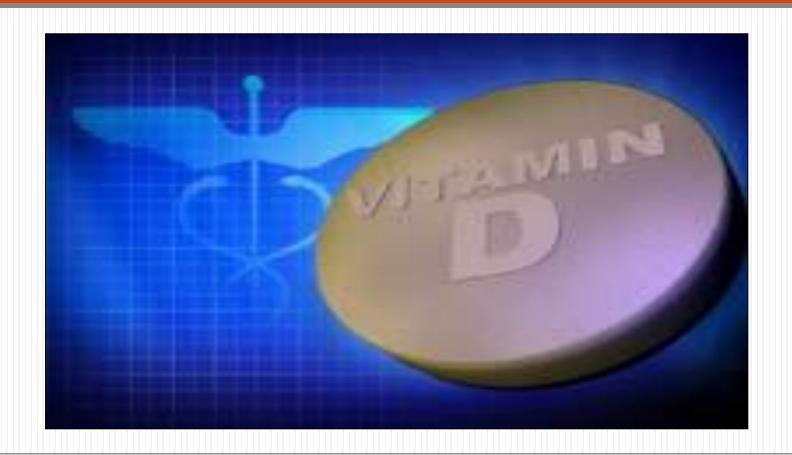
How do we stimulate repair?



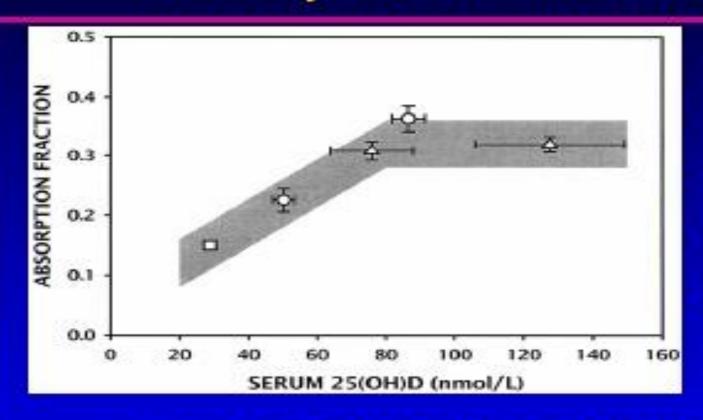


# Are There Nutrients More Important Than Calcium?

#### Vitamin D



#### Relation between Serum 25(OH)D and Calcium Absorption in Vitamin D Insufficiency



Calcium absorption fraction as a function of serum 25(OH)D3 concentrations (combined results of 3 studies)

- ☐ Bischoff et al. J Bone Miner Res 2003; 18: 343-51.
- O Heaney et al. J Am Coll Nutr 2003; 22: 142-6.
- △Barger-Lux et al. J Clin Endocrinol Metab 2002;87:4952–6.

Error bars indicate ± 1 SEM

From Heaney RP Am J Clin Nutr 2005; 80: 1706S-9S.

#### Vitamin D Inadequacy Is Widespread

- 1 billion people worldwide are deficient in vitamin D, especially among the elderly and dark-skinned individuals.
- Half of adults in US, Europe, and up to two-thirds of post-menopausal women are deficient in vitamin D.
- 30 to 40% of children in many cultures are deficient in vitamin D.
- 80% of school children in New Delhi are deficient in vitamin D.
- 97% of hip fracture patients in UK are deficient in vitamin D.
- Half the population of Tasmania is deficient in vitamin D.

## Meta-Analysis Trials Using 700–800 IU Vitamin D<sub>3</sub> and Calcium

In a 1992 18-month study (n=3270), mean age 84, ambulatory elderly, using 1.2g calcium, 800 IU vitamin  $D_3$  . . .

- ✓ 32% fewer non-vertebral fractures
- ✓ 43% fewer hip fractures

#### Vitamin D and Fracture Reduction

The Better Bones Foundation estimates that supplementation with therapeutic levels of vitamin D could result in an overall **50- to 60-percent reduction** in low-trauma osteoporotic fractures.

Other vitamin D researchers who agree with this statement include:

- William Grant, PhD
- Cedric Garland, DrPH
- Michael Holick, MD, PhD

# Vitamin K



#### Vitamin K and Bone

Vitamin K is essential for the carboxylation of osteocalcin, the bone protein which attracts calcium to the crystallization site.

- Hart et al. 1985

#### Vitamin K and Fracture Risk

Work with elderly French patients shows that **ucOC**, **but not conventional calcium metabolism parameters**, predicts the subsequent risk of hip fracture.

- Szulc et al. 1993

European EPIDOS Study found ucOC to be major independent risk factor for hip fracture among healthy elderly women. The women with low BMD and high ucOC had a 5.5x increased risk of hip fracture as compared to those with only low BMD or high ucOC levels.

Vergnaud et al. 1997

# Dietary Vitamin K and Fracture

Data from the Framingham Heart Study revealed that those women and men in the highest quintile of vitamin K intake (250 mcg/d) had one-third the risk of hip fracture as those in the lowest quintile (75 mcg/d).

- Booth et al. 2000

The Nurses Health Study (N=72,732) also reported that the women in the **lowest** quintile of vitamin K intake had an increased risk of hip fracture.

- Feskanich et al. 1999

# Meta-Analysis of Vitamin K and Fracture Reduction

7 RCTs using MK-4 (menatetrenone) (6 using 45 mg and 1 using 15 mg)

- 77% Reduction in Hip Fractures
- 60% Reduction in Vertebral Fractures
- 81% Reduction in all Non-Vertebral Fractures

- Cockayne et al. 2006

#### Ascorbate: The Best Form of Vitamin C



# Vitamin C Is Essential for Healthy Bones

- As an antioxidant to protect bone from free radical damage
- As an electron donor to energize bone cells
- A key nutrient for healthy collagen

According to new research on the effects of vitamin C on bone health, moderate amounts of vitamin C led to a nearly 44% reduction in risk of fracture.

#### **Trace Minerals**

Magnesium, Zinc, Manganese, Copper, Boron, Silica, Iron

Inadequacy Is Common!!

Actual intake levels and therapeutic levels are outlined on the 20 Key Bone Building Nutrients Chart Handout

### Sub-Optimal Nutrient Intake Is the Norm

Magnesium: All ages, sexes, and classes (except those under 5) fail to consume

even the RDA.

**Zinc:** Average intake is 46 to 63% the RDA.

**Potassium:** Average intake is around 50% RDA.

**Copper:** 75% of diets fail to contain the RDA.

Vitamin D: Low serum levels are common.

**Calcium:** Average intake is 500 to 600 mg.

http://www.betterbones.com/bonenutrition/20keybonenutrients.pdf

# Bone Protein Matrix Minerals: Manganese, Zinc , Copper, and Iron

- Copper and iron act as cofactors in the cross-linking of collagen and elastins.
- Manganese participates in the biosynthesis of mucopolysaccharides.
- Zinc deficiency causes a reduction in osteoblastic activity, collagen and chondroitin sulfate synthesis, and alkaline phosphatase activity.

# Magnesium Depletion and Osteoporosis — Experimental Animal Models

- Universal observation is decreased growth of whole body and skeleton
- Reduced osteoblast formation
- Decreased collagen formation
- Impaired mineralization
- Production of brittle and fragile bone

# Magnesium Builds Bone in Girls

RCT of 120 girls (8 to 14 yrs) with magnesium intakes less than 220 mg/day and a 1-year intervention with 300 mg magnesium:

- Significant improvement in hip BMD.
- Slight increase in spinal BMD.

Magnesium Builds Bone in Postmenopausal

Women

Magnesium, Zinc, Copper, Manganese, and Selenium Levels in Postmenopausal Osteonorosis?

Can Magnesium Play a Key Role in Osteonorosis. Magnesium, Zinc, Copper, Manganese, and Selenium Levels in Postmenopausa.

Women with Osteoporosis.

A horner A witness of Turan Law A horner A witness of Tur women with Osteoporosis. Can wiagnesmin Fray a Key Kole in Osteoporosis. Can wiagnesmin Fray a Key Kole in Osteoporosis. Ahmet Aydin, Cemal Akay, Mustafa Kutlu, Mostafa Turan, Ahmet Aydin, Cemal Akay, Mostafa Kutlu, Mosta

Original Article

Introduction: Therehasbeen a resurgence of interest in sta in the develop ment and maintenance of the skeleton. The aim m the develop ment and maintenance of the sketeton. I he at and red blood concentrations of some elements in Posth maternisana meinoas: Seventy-seven postmenopa usan (median interquartile range, 7.5; range, 46 to 74) and 6 menopausal women aged 60 years (median interqual menopausal women aged ou years (median interqual included in the study, Element concentrations in plasma Melanca in incision, element concentrations in para spectrop hotometry in both postmenopausal women wi spectrop no to metry me o the ostmenopausal women we women. Results Only statistically significant difference women. Results Only statistically significant difference women. women. Kesulis: Only statistically againstant amer.

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18 (ml.) and healthy subjects (54.54 [15.42] µg/m magnesium concentration (Z = 2.07, P = 0.039). E

Biol Trace Elem Res (2010) 133:136-143

**Short-Term Oral Magnesium Supplementation** Suppresses Bone Turnover in Postmenopausal Osteoporotic Women

Hasan Aydın · Oğuzhan Deyneli · Dilek Yavuz · Hülya Gözü · Nilgün Mutlu · Işık Kaygusuz · Sema Akalın

Received: 20 April 2009 / Accepted: 20 May 2009 / Published online: 2 June 2009 © Humana Press Inc. 2009

Abstract Magnesium has been shown to increase bone mineral density whe treatment of osteoporosis, yet its mechanism of action is obscure. In this stuof daily oral magnesium supplementation on biochemical markers of bone

#### Magnesium Intake from Food and Supplements Is Associated with Bone Mineral Density in Healthy Older White Subjects

Kathryn M. Ryder, MD, MS,\* Ronald I. Shorr, MD, MS,† Andrew J. Bush, PhD,† Stephen B. Kritchevsky, PhD, † Tamara Harris, MD, MPH, \* Katie Stone, PhD, | Jane Cauley, DrPH, and Frances A. Tylavsky, DrPH<sup>†</sup>

OBJECTIVES: To determine whether magnesium intake from supplemental and dietary sources is associated with bone mineral density (BMD) in older men and women,

DESIGN: Cross-sectional.

Magnesium intake, bone mineral density, and fractures: results from

Tonya S Orchand, Joseph C Larson, Nora Alghothani, Sharon Bour-Tabaku, Jane A Cauley, Zhao Chen, Andrea Z LaCroix,

the Women's Health Initiative Observational Study 1-4

Background: Magnesium is a necessary component of bone, but its

Objective: We examined magnesium intake osteoporotic fractures and altered bone mineral

Design: This prospective cohort study included

tional Study. Total day

SETTING: Memphis, Tennessee, and Pittsburgh, Pennsyl-

PARTICIPANTS: Two thousand thirty-eight older black and white men and women aged 70 to 79 at baseline enrolled in the Health, Aging and Body Composition Study. MEASUREMENTS: Dietary intake of magnesium was assessed using a semiquantitative food frequency questionnaire, and supplement data were collected based on a medication inventory, BMD of the whole body was obtained using a fan-beam densitometer. Additional covariates included age, body mass index (BMI), smoking status, alcohol use, physical activity, estrogen use, and supplemenCONCLUSION: Greater magnesium intake was significantly related to higher BMD in white women and men. The lack of association observed in black women and men may be related to differences in Ca regulation or in nutrient reporting. J Am Geriatr Soc 53:1875-1880, 2005.

Key words: bone mineral density; nutrition; magnesium; osteoporosis: elderly

Osteoporotic fractures are a significant health problem in older adults, and the burden of osteoporosis is expected to increase as the population ages. White woman have a lifetime risk of any clinical fracture approaching 75%2 and a lifetime risk of hip fracture of 16%. White men

#### Zinc Is Essential for Bone Health

- Zinc stimulates osteoblast bone-building formation and mineralization.
- It facilitates bone collagen synthesis.
- It inhibits osteoclastic bone breakdown.
- Studies show women with osteoporosis have lower than normal levels of zinc.

# Multiple Intervention Strategies

2-year multi-nutrient blinded, controlled study, N59 healthy postmenopausal women:

#### <u>Intervention</u>:

- Placebo
- Just Ca citrate malate (1,000 mg)
- Ca citrate malate (1,000 mg), zinc (15 mg), manganese (5 mg), and copper (2.5 mg).

#### **BMD Changes:**

The only significant change from placebo was calcium plus trace minerals. This was the only group to halt loss and gain bone.

#### Multi-Nutrient Intake and Fracture Risk

In a Swedish observational study of 65,000 women aged 48 to 80, when highest quartile of intake was compared to lowest:

- Intakes of iron, magnesium, and vitamin C were found to be independent risk factors for hip fracture (or 3.3; 2.7; 1.9).
- High calcium intake did not protect against hip fracture.

# Potassium: The Unexpected Bone Builder

The RDA for potassium at 4,700 mg is nearly **four times** that of calcium (at 1,200 mg).

**Potassium Protects Bone** 



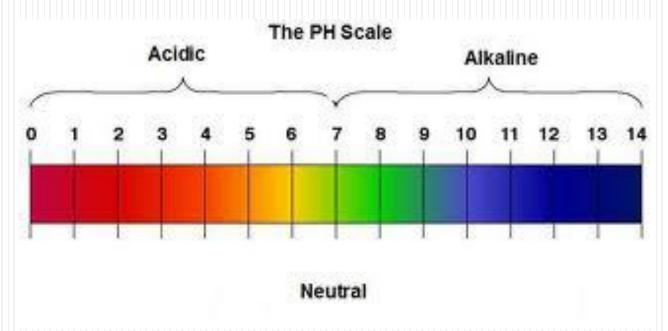
#### Metabolic Functions of Bone

- Serum Calcium Homeostasis
- Systemic Acid-Base Balance

**Bone Sacrifices Structural Functions for Metabolic Functions** 

# Obligatory pH Balance and Chronic Low Grade Metabolic Acidosis

pH homeostatic mechanisms are central to our discussion of toxic metals and bone health.





Our skeleton serves as a gigantic reserve of alkaline mineral compounds ready for transfer into the blood for maintenance of an obligatory, precise pH level.

- Brown and Jaffe 2000

# Bone Is a Gigantic Alkali Buffer Exchange Column Loaded with Alkali Buffer Compounds

Bone and the hydration shell around it contain:

- 80% of body carbonate
- 80% of body citrate
- 35% of body sodium
- 53 to 80% of body magnesium
- 0.1 to 0.2% of body potassium

- Brown and Jaffe 2000; Green and Kleeman 1991a



If we do not maintain adequate alkaline-mineral reserves from our diet, our skeleton willingly sacrifices itself for survival of the whole.

#### Chronic Metabolic Acidosis

- Dietary changes over the last two centuries have resulted in a mismatch between genetically determined nutritional requirements and actual nutrient intake.
- This has resulted in *chronic metabolic acidosis* being the norm rather than the exception:
  - ✓ Deficiency of potassium, magnesium, and other essential minerals from alkaline-forming foods
  - ✓ Excess dietary intake of acid-forming foods
  - ✓ Excess sodium chloride

## Cell Net Acid Excess (NAE)

NAE is a key determinant of bone health because of its role in:

- Regulating the efficiency of protein synthesis
- Controlling bone alkaline mineral reserves
- Regulating the efficiency of our elective protective and anti-toxic mechanisms

# Metabolic Acidosis Is the Norm and It Endangers Bone

- Bone loss can be re-thought of and understood more fundamentally as a "hidden tax of high-tech living."
- People with high-tech diets and lifestyles usually acquire chronic metabolic cellular acidosis and become progressively more deficient in cell mineral reserves that proportionately impair efforts to rebuild bone matrix and detoxify.

- Frassetto et al. 2005; Jehle et al. 2013

# Chronic Low-Grade Metabolic Acidosis Damages Bone

These acid-forming excesses are not mediated by dietary bicarbonates and lead to chronic low-grade metabolic acidosis with ensuing damage to bone. Among other effects you see:

- Urinary loss of calcium
- Increased bone resorption
- Kidney function decline
- Reduction in growth hormone
- Nitrogen and muscle losses

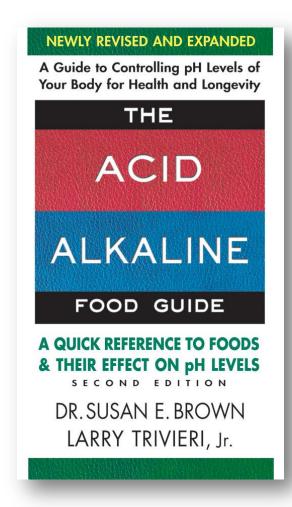
Remember . . .

Chronic Low-Grade Metabolic Acidosis



**Low Mineral Status** 

#### The Acid-Alkaline Food Guide



# Become a Nutrition Detective

