

THE WEBINAR BEGINS AT 19.30 GMT

Perimenopause and menopause transition: optimising hormones for long-term health



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PhD in Medical Sciences (University of Groningen)

MSc in clinical PNI and Evolutionary Medicine

Degree in Physiotherapy and Osteopathy

Health Coach and Therapist according to cPNI

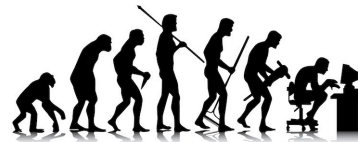
President of the AEPNIC

Co-CEO of Healthy Institute

Scientific advisor



@drbegoruiznunez
@healthyinstitute



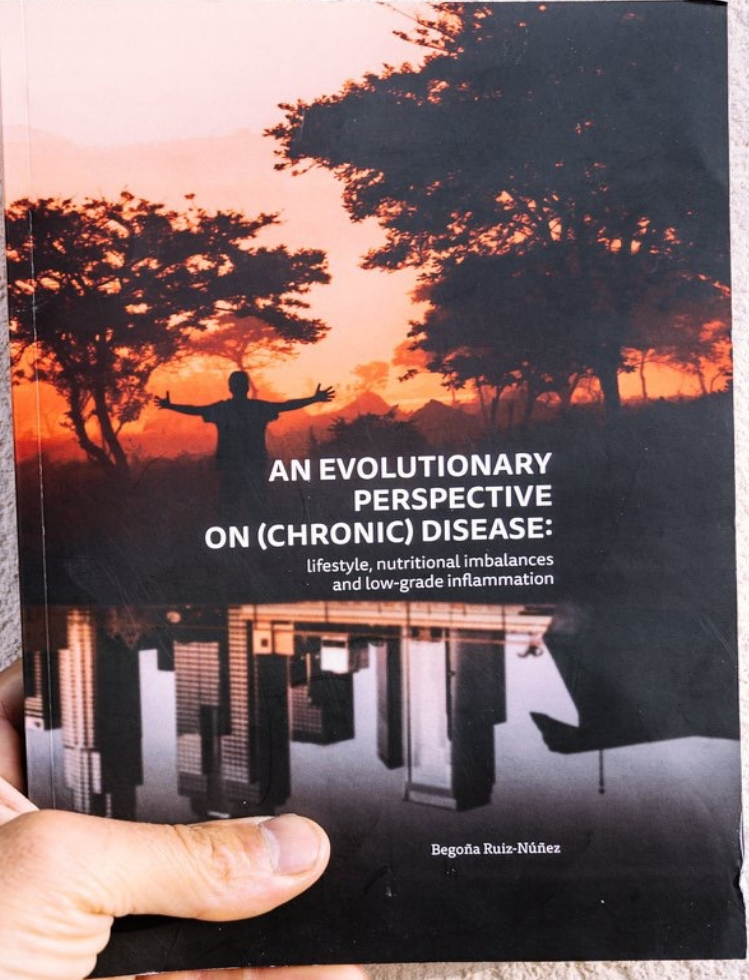
AEPNIC

Asociación Española de
Psico-Neuro-Inmunología Clínica



www.healthyinstitute.es





**AN EVOLUTIONARY
PERSPECTIVE
ON (CHRONIC) DISEASE:**

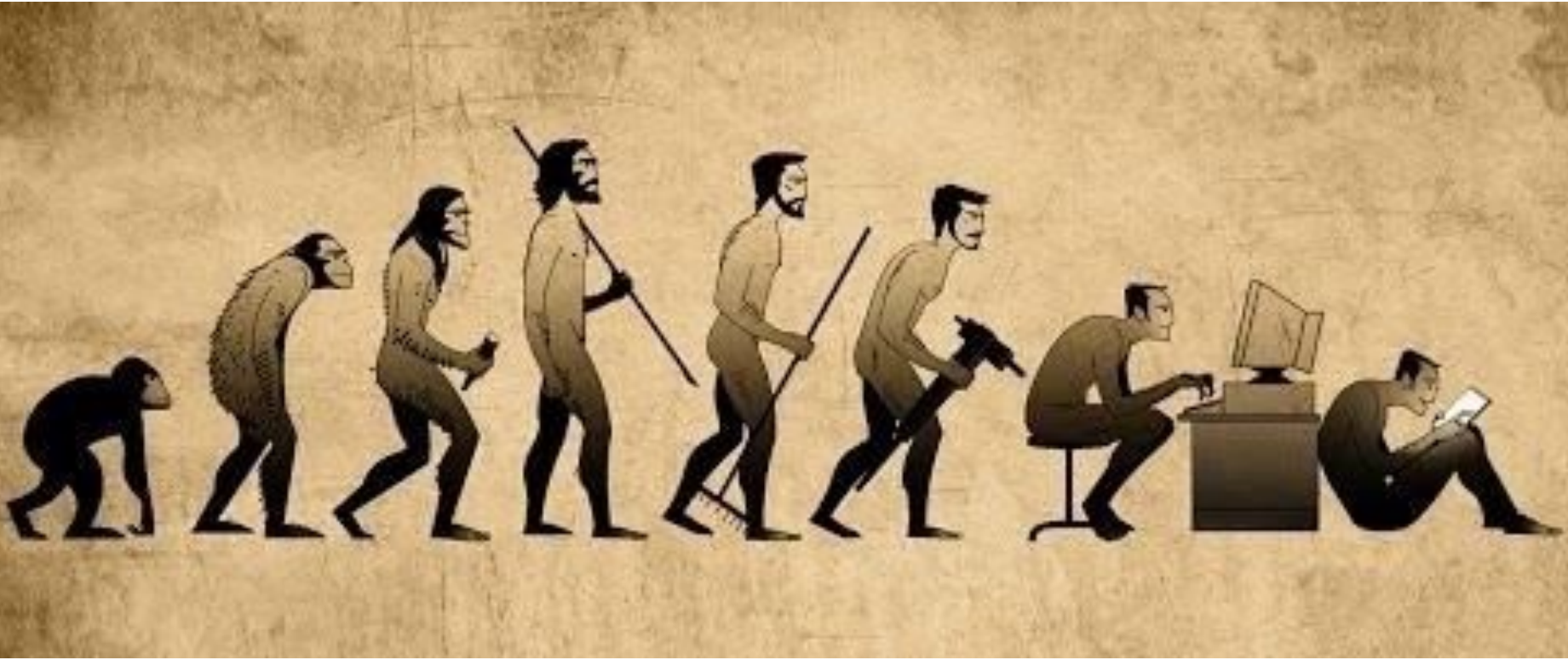
lifestyle, nutritional imbalances
and low-grade inflammation

Begoña Ruiz-Núñez

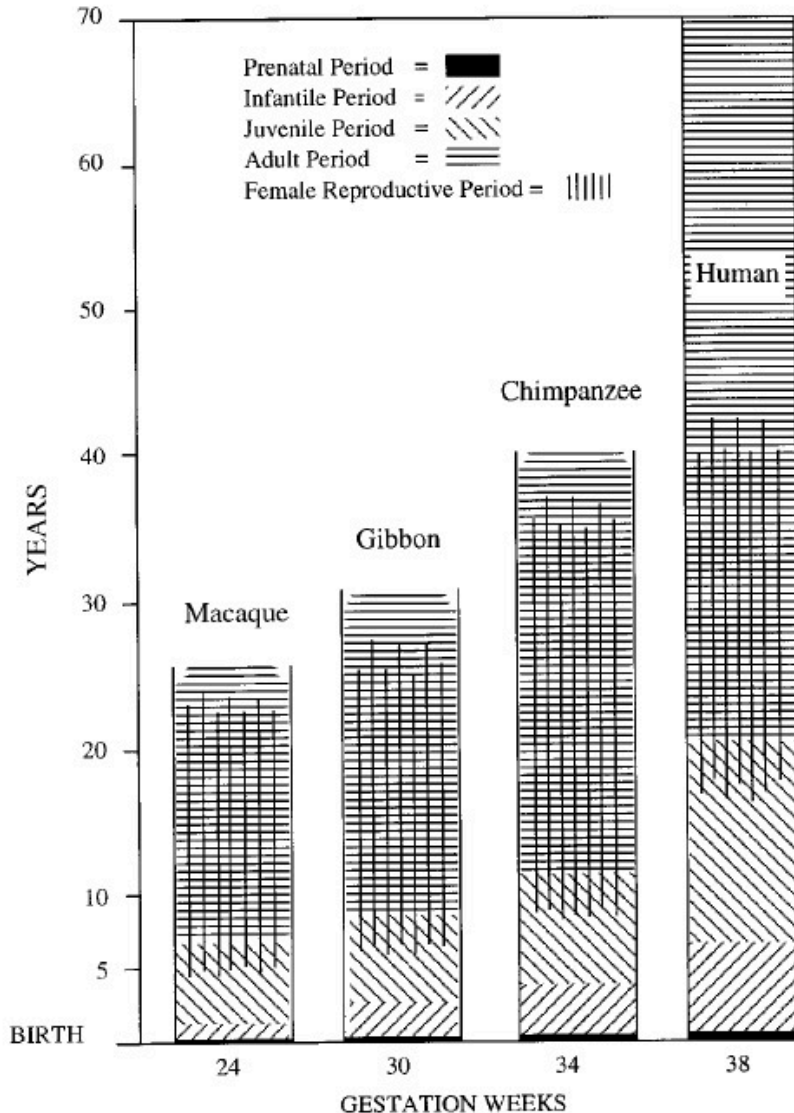
And... here we are!



Evolution?



Menopause and evolution



Compared to other primates:

- ✓ Longer gestational period
- ✓ Longer prenatal, infantile and juvenile periods
- ✓ Much longer adult period
- ✓ Shorter female reproductive period
- ✓ Longer post-menopausal lifespan

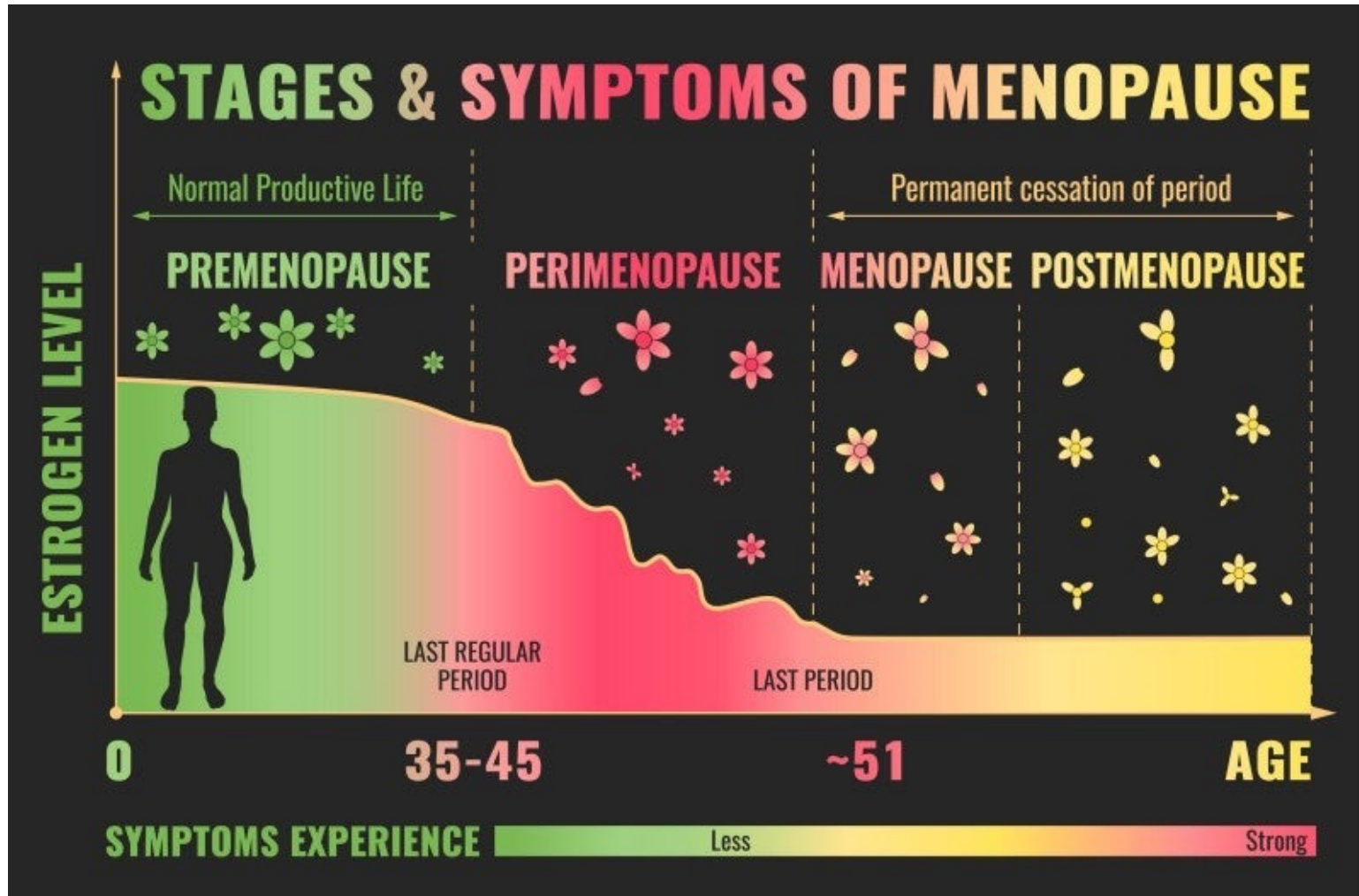
Menopause and evolution

- ✓ Avoids mutations
- ✓ Healthier offspring
- ✓ Normal coresidence between older mothers and fertile daughters (not sons)
- ✓ Help daughter(s) with food supply and production
- ✓ Extended childhood learning



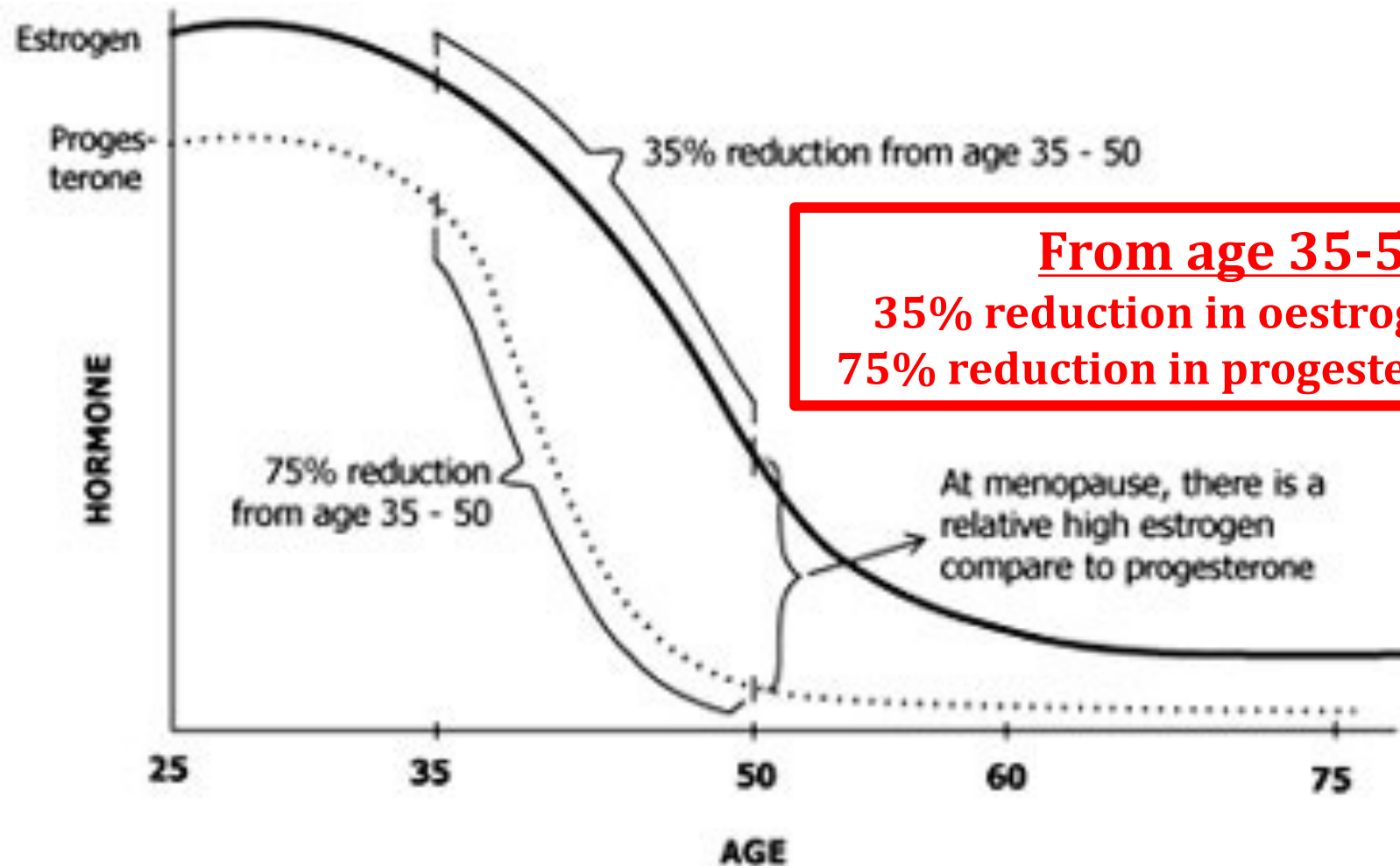
Hawkes, K., O'Connell, J. F., Jones, N. B., Alvarez, H., & Charnov, E. L. (1998). Grandmothering, menopause, and the evolution of human life histories. *Proceedings of the National Academy of Sciences*, 95(3), 1336-1339.

Menopause



FACTS

Perimenopause Ovarian hormones



Menopause

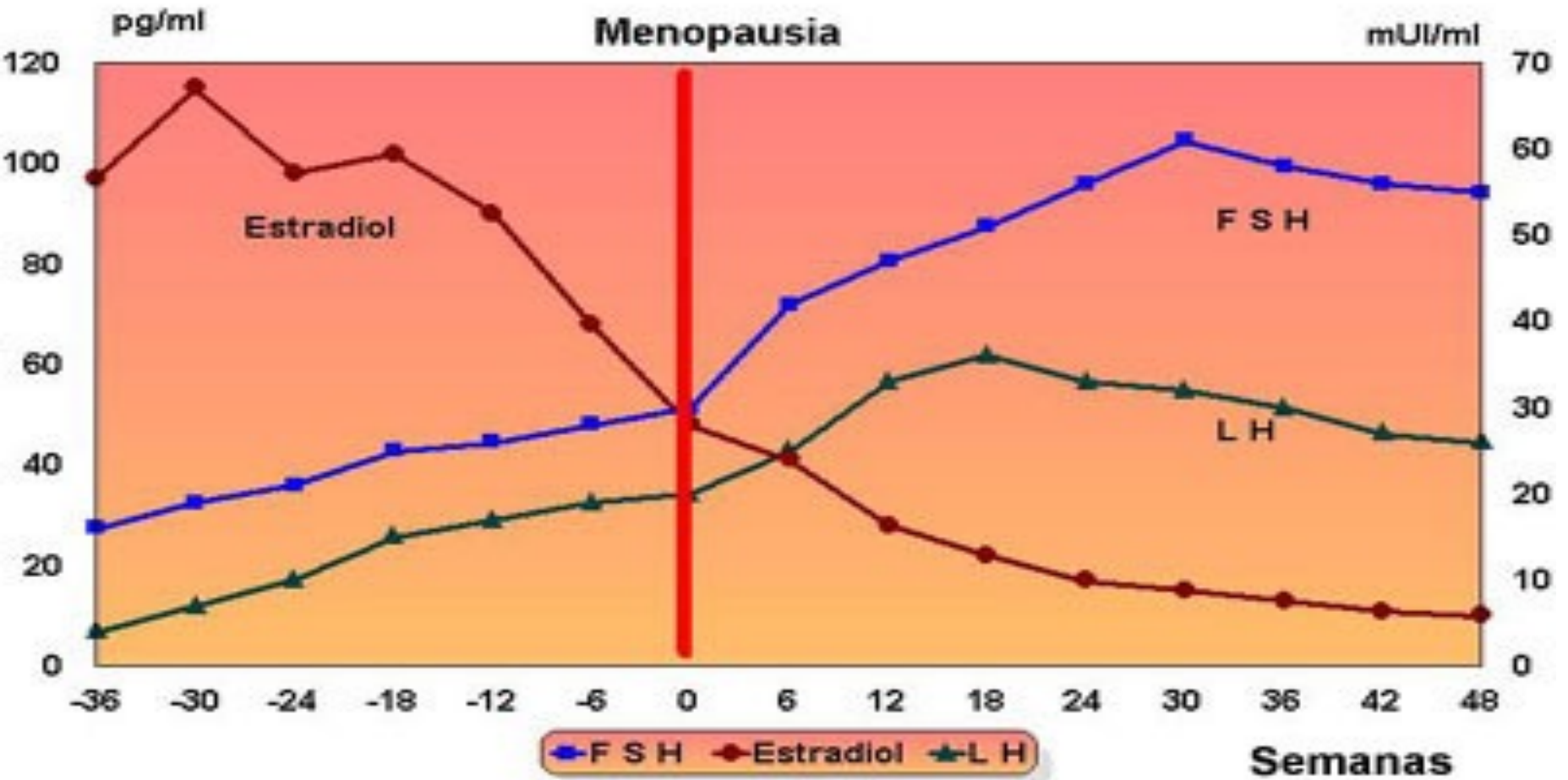


Figura 3. Cambios en los niveles de FSH, Estradiol y LH durante el climaterio

FACTS

Perimenopause--->heading menopause

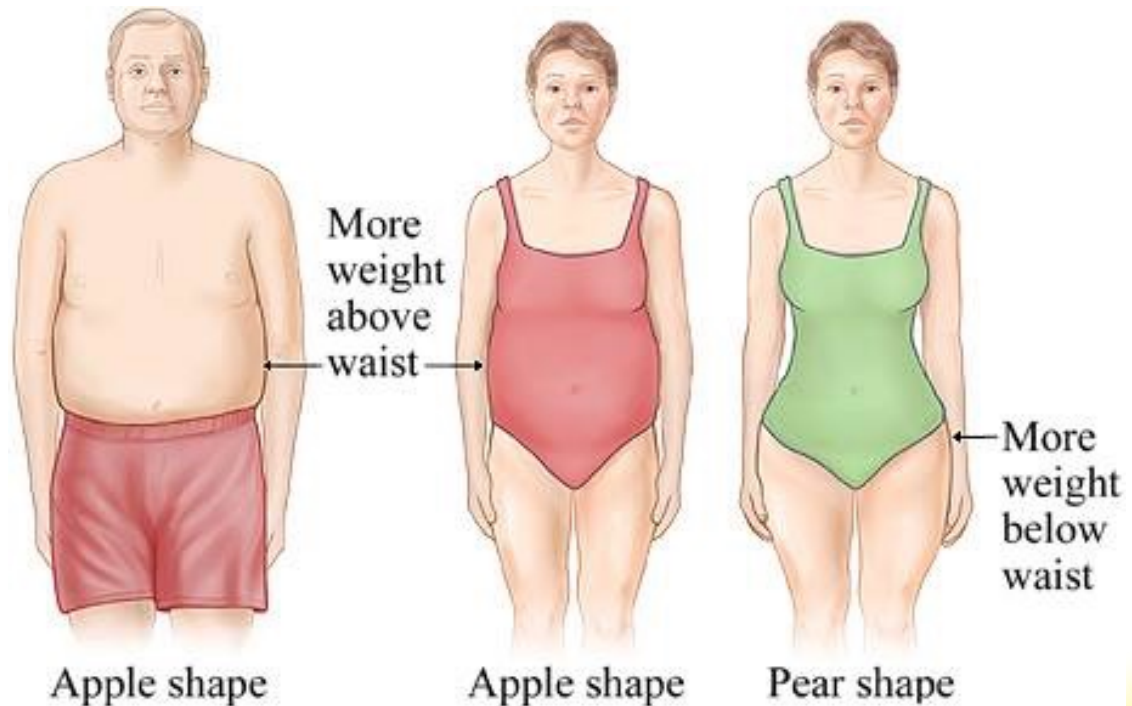
- ✓ Lower oestrogen levels
- ✓ Much lower progesterone levels
- ✓ Higher levels of pituitary hormones (LH & FSH)



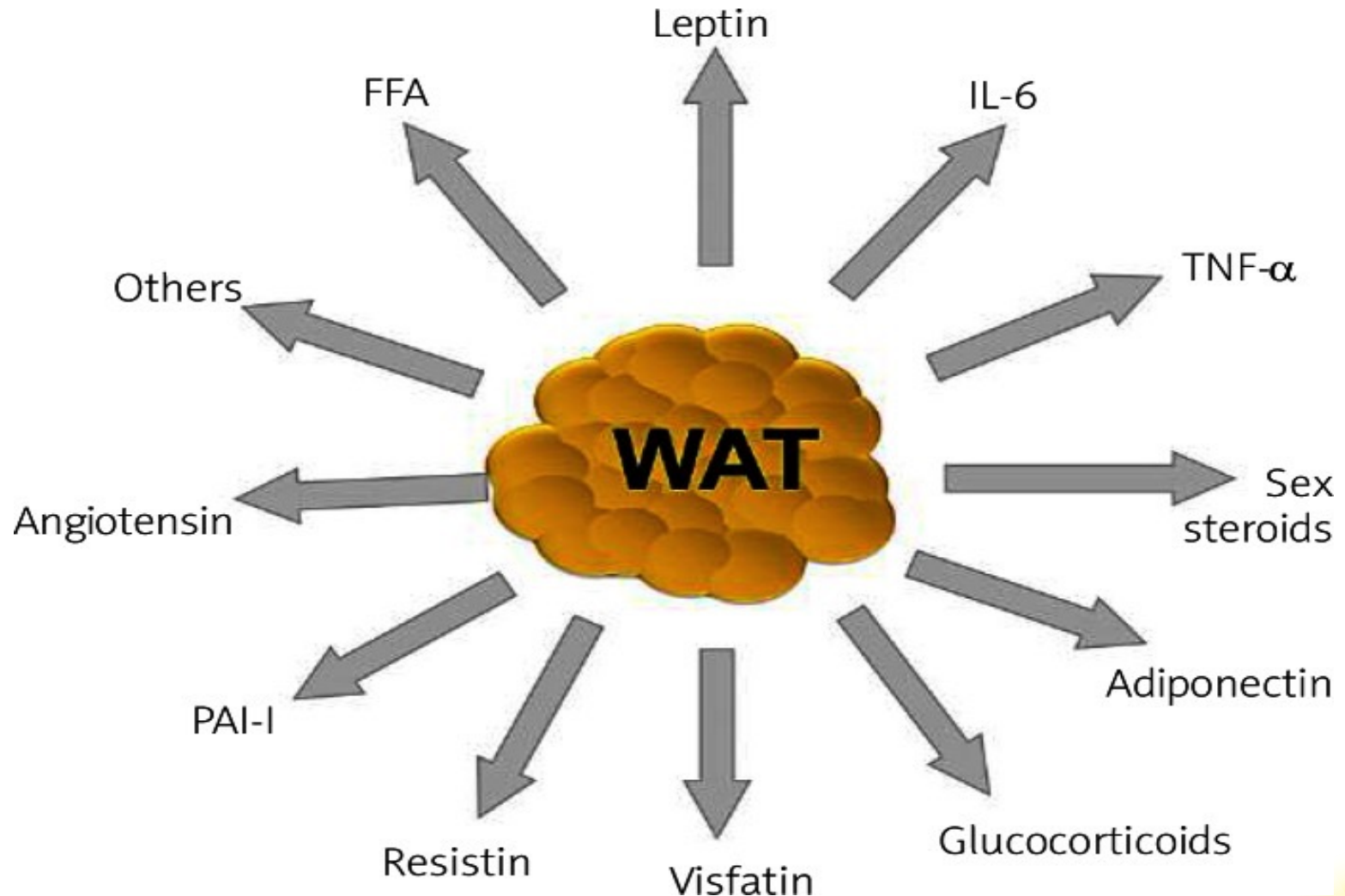
FACTS

However...

The perimenopause is associated with a more rapid increase in fat mass and redistribution of fat to the ABDOMEN, resulting in a transition from a gynoid to an ANDROID pattern of fat distribution and an INCREASE IN TOTAL BODY FAT



Abdominal fat as an endocrine organ



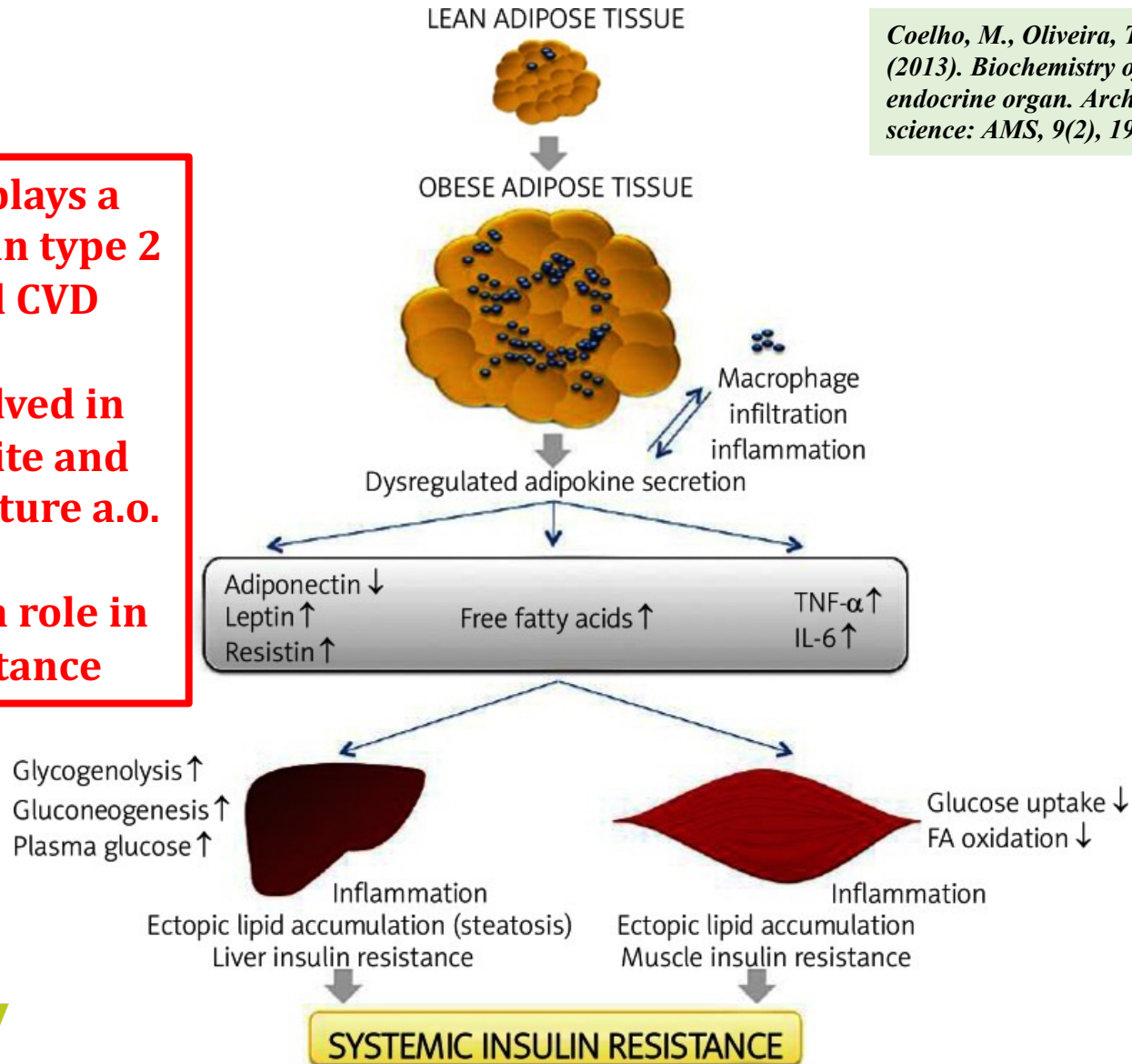
Abdominal fat as an endocrine organ

Coelho, M., Oliveira, T., & Fernandes, R. (2013). *Biochemistry of adipose tissue: an endocrine organ. Archives of medical science: AMS, 9(2), 191.*

Adiponectin plays a protective role in type 2 diabetes and CVD

Leptin is involved in satiety/appetite and energy expenditure a.o.

Resistin plays a role in insulin resistance



Abdominal fat as an endocrine organ

www.nature.com/scientificreports

scientific reports

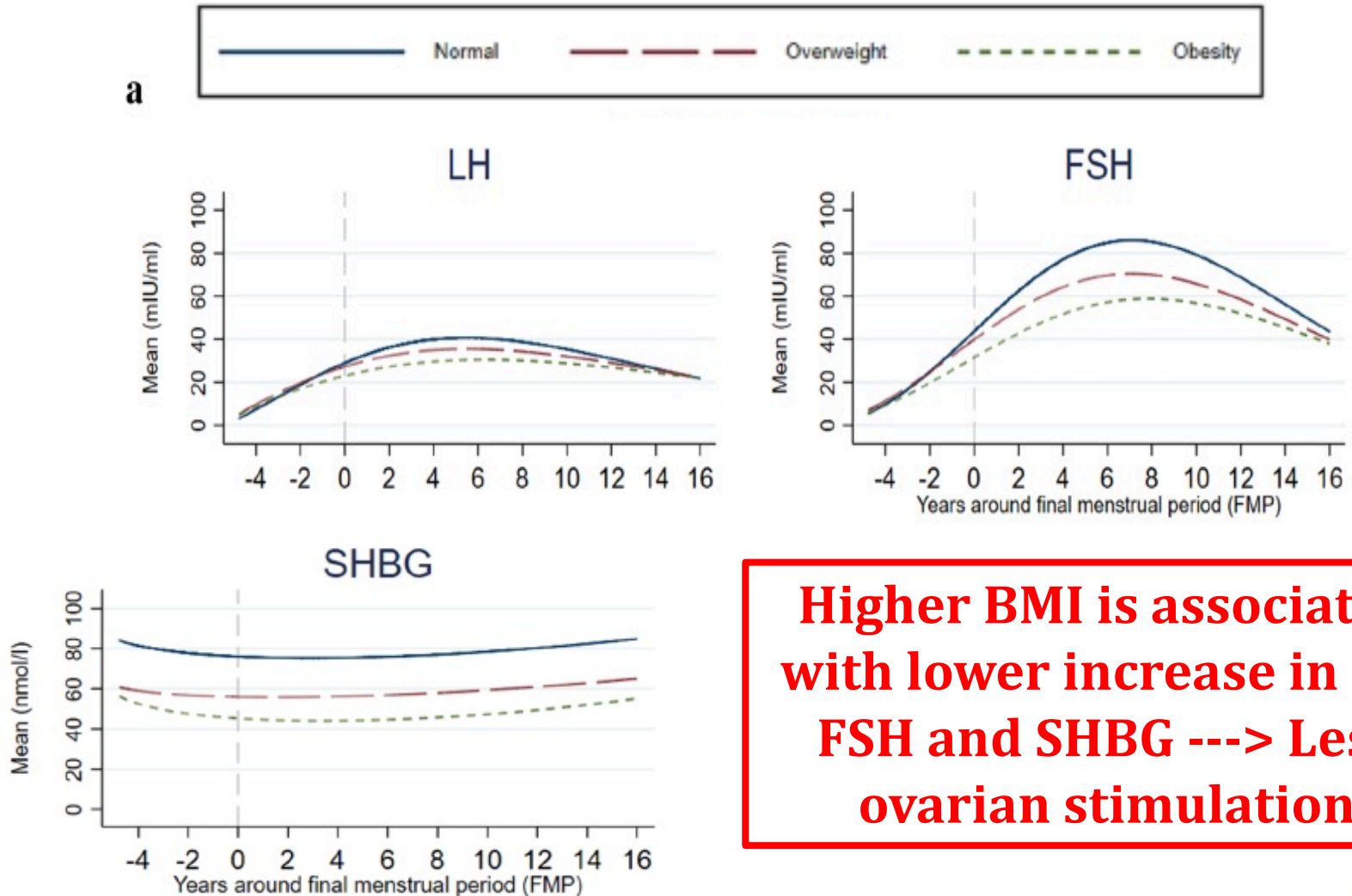
 Check for updates

OPEN

Longitudinal changes in reproductive hormones through the menopause transition in the Avon Longitudinal Study of Parents and Children (ALSPAC)

Ana Goncalves Soares ^{1,2,✉}, Fanny Kilpi ^{1,2}, Abigail Fraser ^{1,2,3}, Scott M. Nelson ^{3,4}, Naveed Sattar ⁵, Paul I. Welsh ⁵, Kate Tilling ^{1,2,3,6} & Deborah A. Lawlor ^{1,2,3,6}

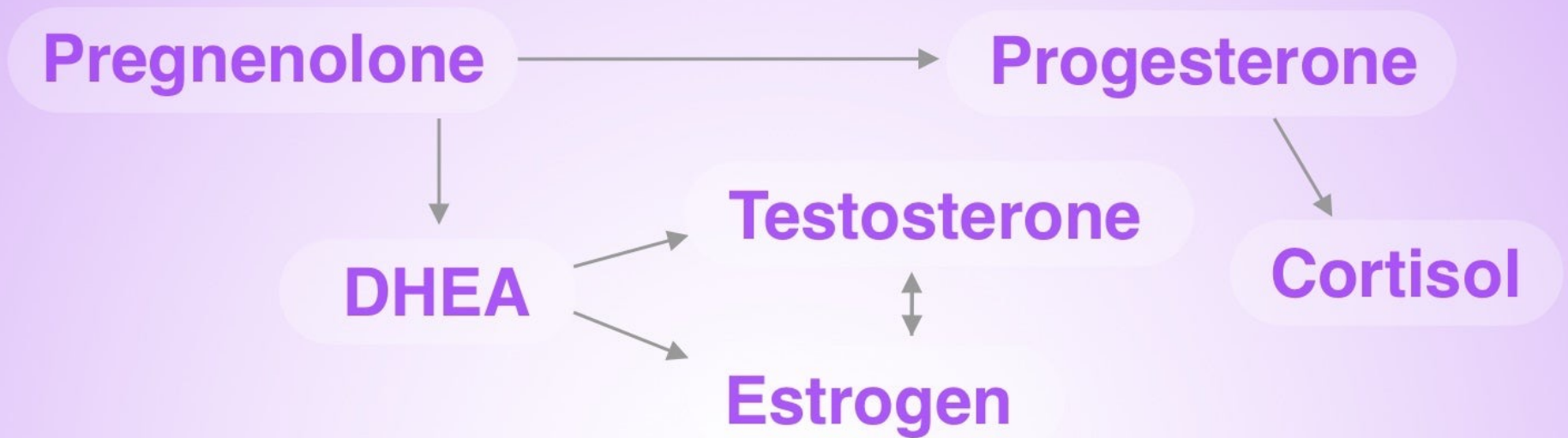
Abdominal fat as an endocrine organ



Higher BMI is associated with lower increase in LH, FSH and SHBG ---> Less ovarian stimulation

FACTS

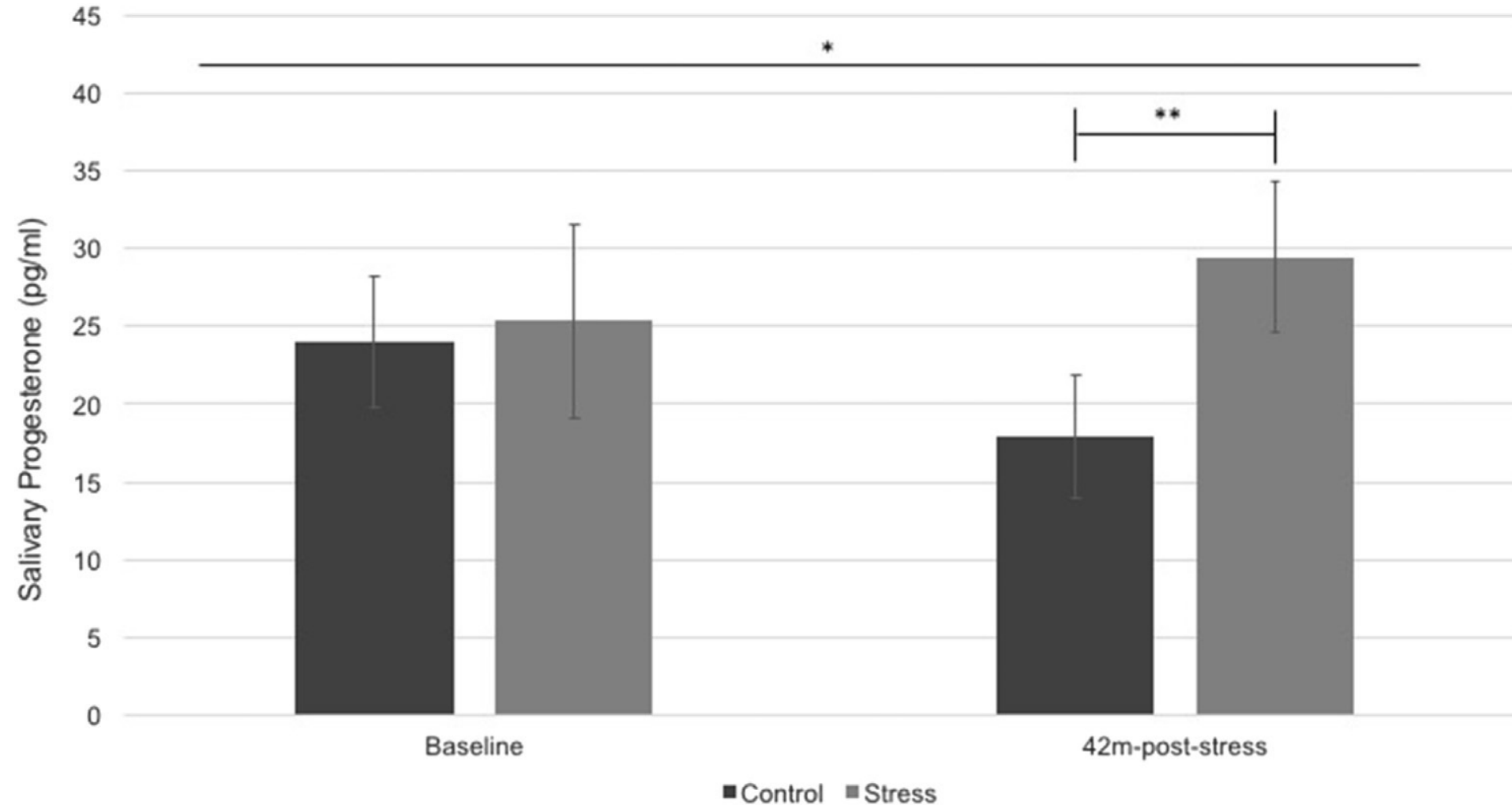
Stress, glucose & sex steroids



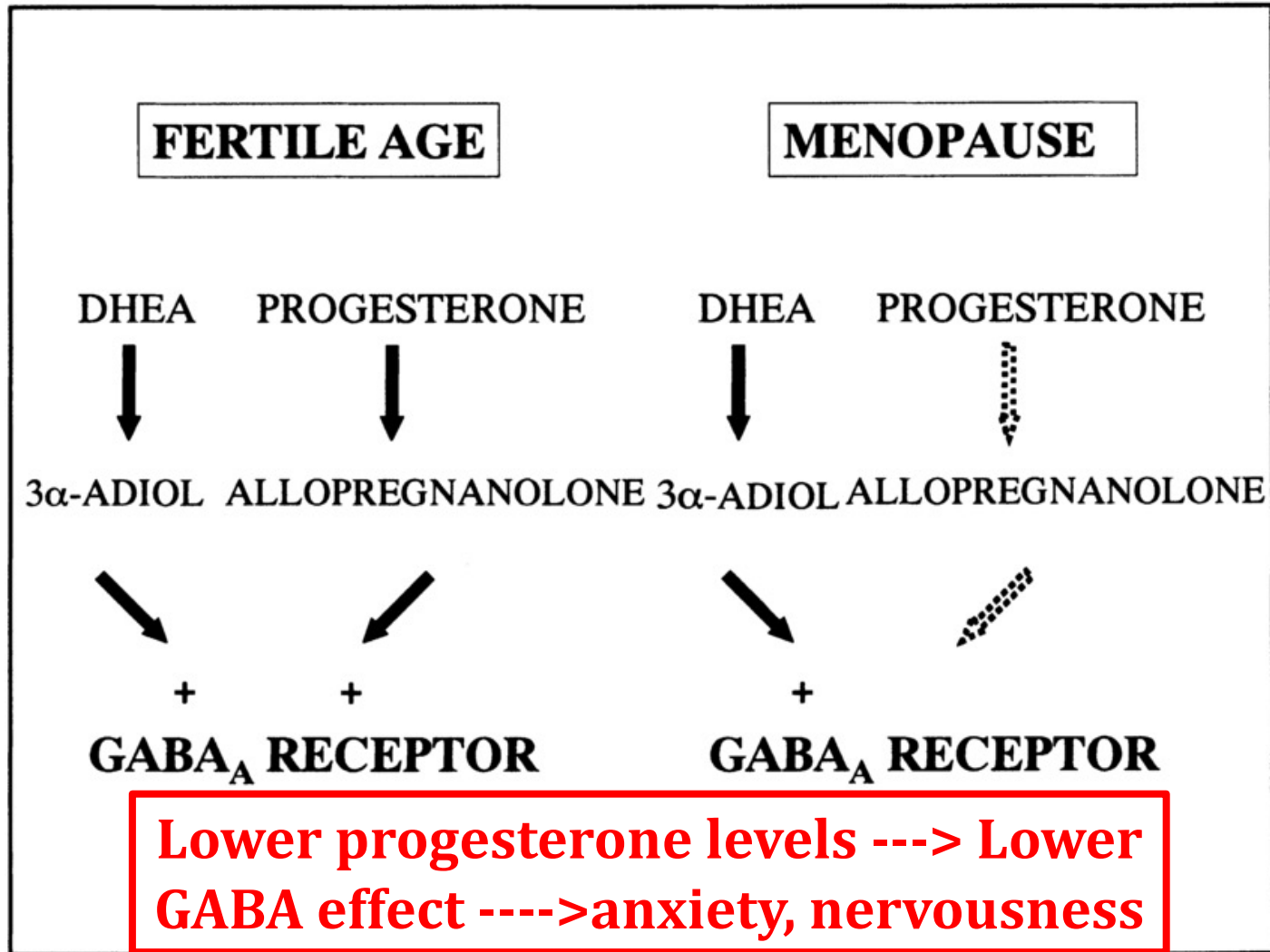
High cortisol/ACTH (stress) decreases DHEA levels
High glucose levels decrease DHEA levels
High stress levels may decrease progesterone, testosterone and oestrogen

FACTS

Progesterone is a “stress-buffering hormone”



Menopause Progesterone & GABA



BARBACCIA, M. L., et al. Plasma 5α-androstane-3α, 17βdiol, an endogenous steroid that positively modulates GABA_A receptor function, and anxiety: a study in menopausal women. Psychoneuroendocrinology, 2000, vol. 25, no 7, p. 659-675.

FACTS

Menopause Sleep

Menopause: The Journal of The North American Menopause Society
Vol. 27, No. 3, pp. 000-000
DOI: 10.1097/GME.0000000000001462
© 2019 by The North American Menopause Society

Effects of menopause on sleep quality and sleep disorders: Canadian Longitudinal Study on Aging

*Sheida Zolfaghari, MD,^{1,2} Chun Yao, MSc,^{1,2} Cynthia Thompson, PhD,³ Nadia Gosselin, PhD,^{3,4,8}
Alex Desautels, MD, PhD,^{3,5,8} Thien Thanh Dang-Vu, MD, PhD,^{6,7,8}
Ronald B. Postuma, MD, MSc,^{2,3,8} and Julie Carrier, PhD^{3,4,7,8}*

Conclusions: Menopause is associated with increased sleep-onset insomnia. Postmenopausal women also are more likely to screen positive for OSA. However, menopausal status is not associated with sleep maintenance, somnolence, or RLS, and RBD.

**Menopausal women require >30 extra minutes to fall sleep
and are more likely to suffer from obstructive sleep apnea**

FACTS

Menopause

The role of magnesium

Decreased levels of oestrogen are associated to decreased levels of magnesium

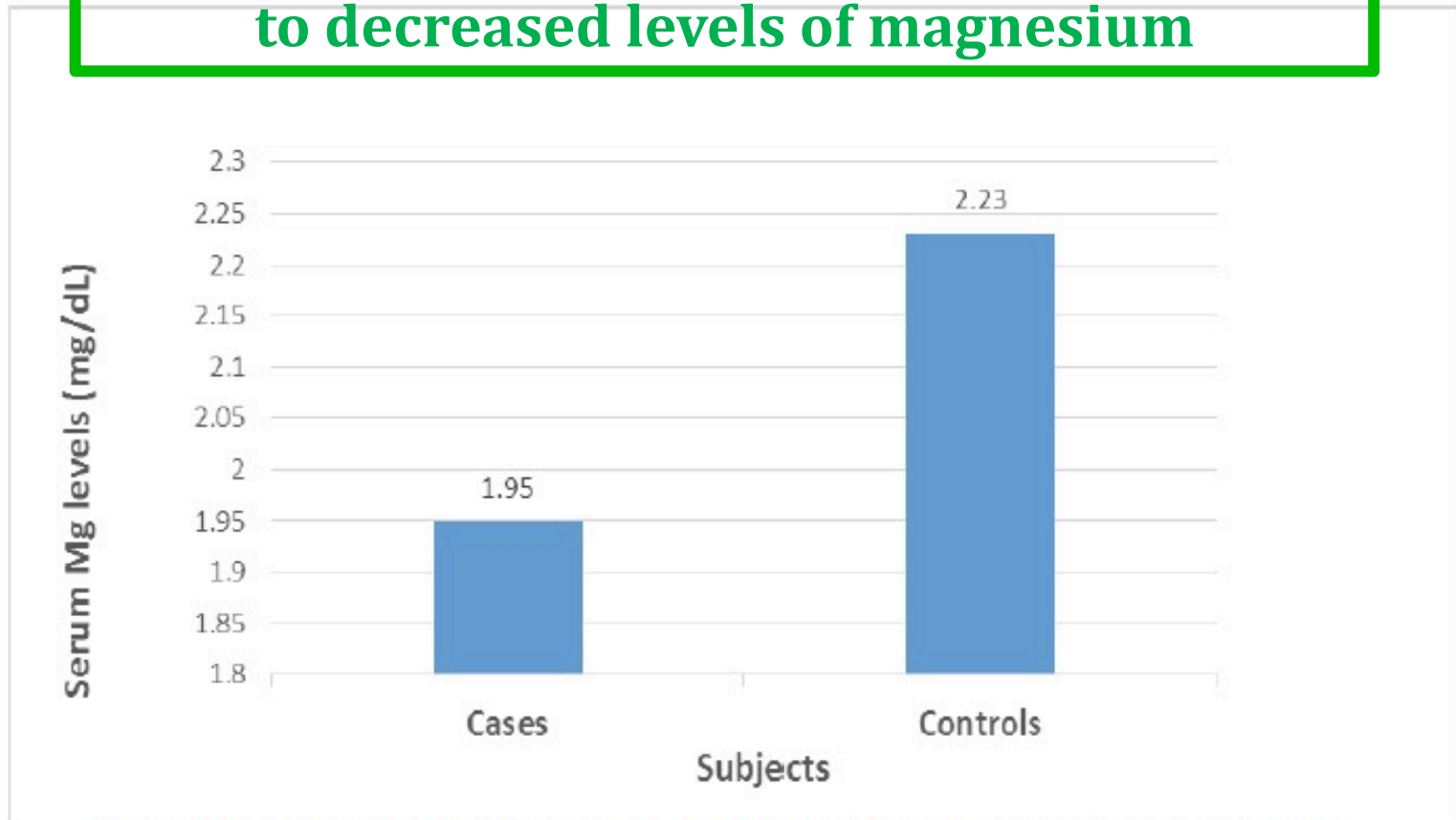


Figure 1: Serum magnesium levels in the Case and Control groups.

Anxiety & depression

The role of magnesium

Association between magnesium intake and depression and anxiety in community-dwelling adults: the Hordaland Health Study

Felice N. Jacka, Simon Overland, Robert Stewart, Grethe S. Tell, Ingvar Bjelland, Arnstein Mykletun

To our knowledge this study is the first to test the hypothesis that magnesium intake is related to anxiety and depression in an epidemiological setting.

While being cognisant of the limitations of the study, the present findings of an inverse association between magnesium intake and depression may have both public health and treatment implications. In primary

Jacka, F. N., Overland, S., Stewart, R., Tell, G. S., Bjelland, I., & Mykletun, A. (2009). Association between magnesium intake and depression and anxiety in community-dwelling adults: the Hordaland Health Study. Australian & New Zealand Journal of Psychiatry, 43(1), 45-52.

FACTS

Menopause Bone health

Premenopausal BMEV

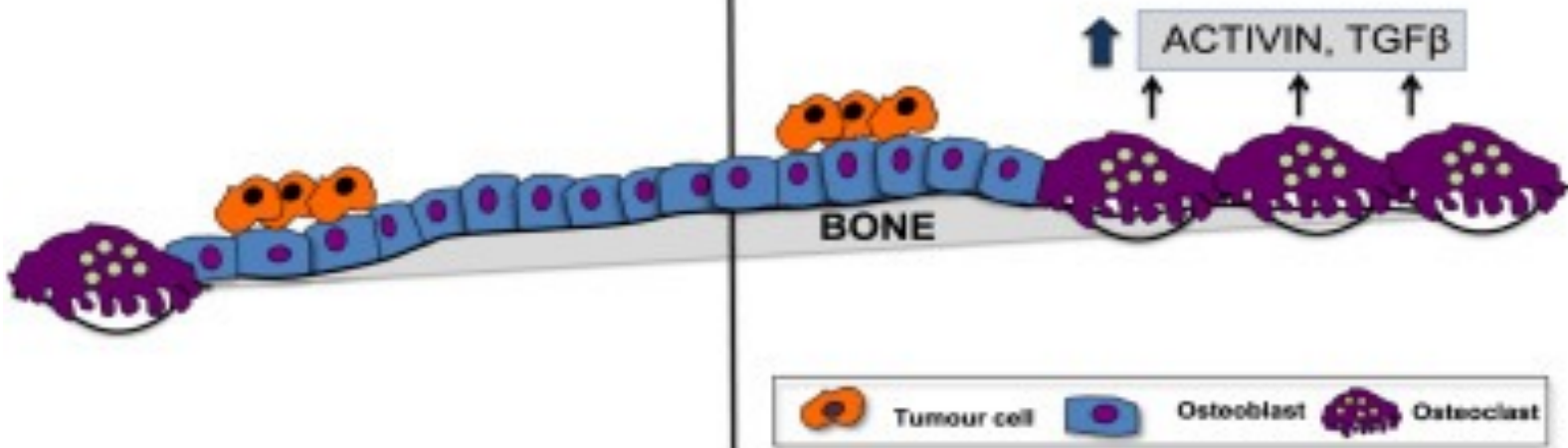


Normal bone turnover
Normal bone density

Postmenopausal BMEV



Increased bone turnover
Decreased bone density



Homocysteine & bone health

The NEW ENGLAND JOURNAL of MEDICINE

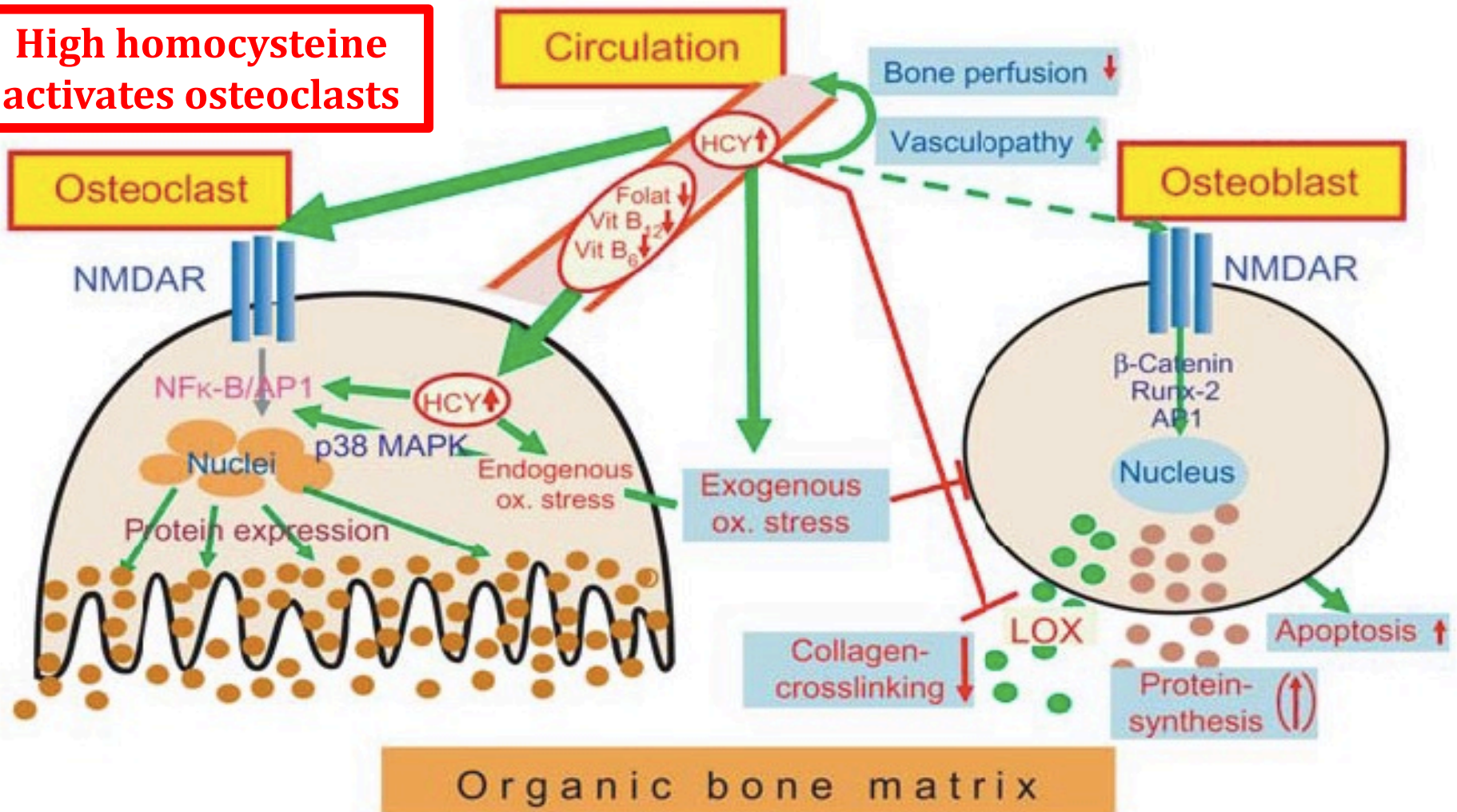
ORIGINAL ARTICLE

Homocysteine as a Predictive Factor for Hip Fracture in Older Persons

Robert R. McLean, M.P.H., Paul F. Jacques, D.Sc., Jacob Selhub, Ph.D.,
Katherine L. Tucker, Ph.D., Elizabeth J. Samelson, Ph.D., Kerry E. Broe, M.P.H.,
Marian T. Hannan, D.Sc., L. Adrienne Cupples, Ph.D., and Douglas P. Kiel, M.D.

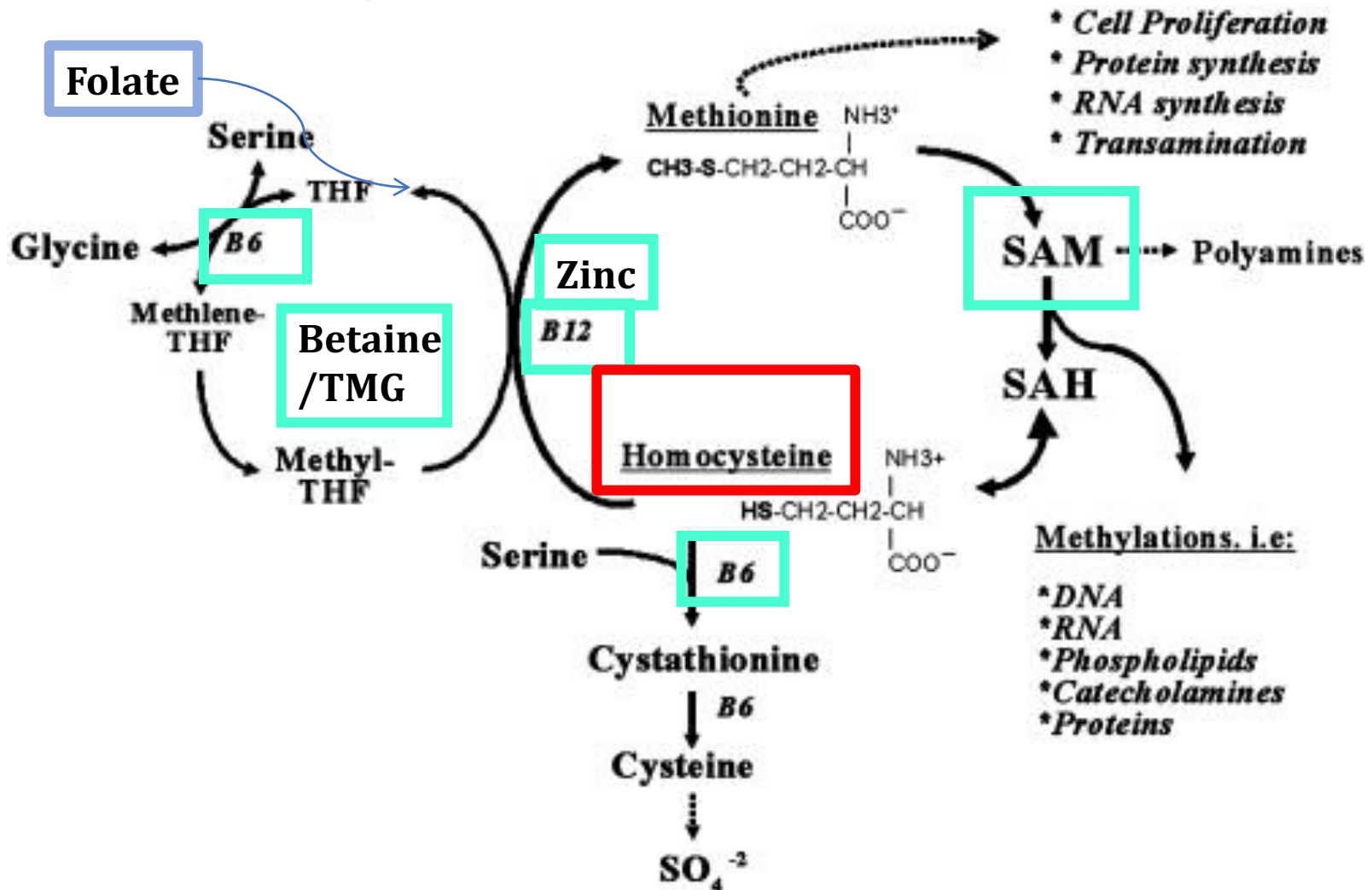
Homocysteine & bone health

High homocysteine activates osteoclasts



Herrmann, M., Peter Schmidt, J., Umanskaya, N., Wagner, A., Taban-Shomal, O., Widmann, T., ... & Herrmann, W. (2007). The role of hyperhomocysteinemia as well as folate, vitamin B6 and B12 deficiencies in osteoporosis—a systematic review. *Clinical Chemical Laboratory Medicine*, 45(12), 1621-1632.

Homocysteine & bone health



Solution(s)



Homocysteine

Vitamin D, folate, B vitamins

Clin Chem Lab Med 2007;45(12):1621–1632 © 2007 by Walter de Gruyter • Berlin • New York. DOI 10.1515/CCLM.2007.362

Review

The role of hyperhomocysteinemia as well as folate, vitamin B₆ and B₁₂ deficiencies in osteoporosis – a systematic review

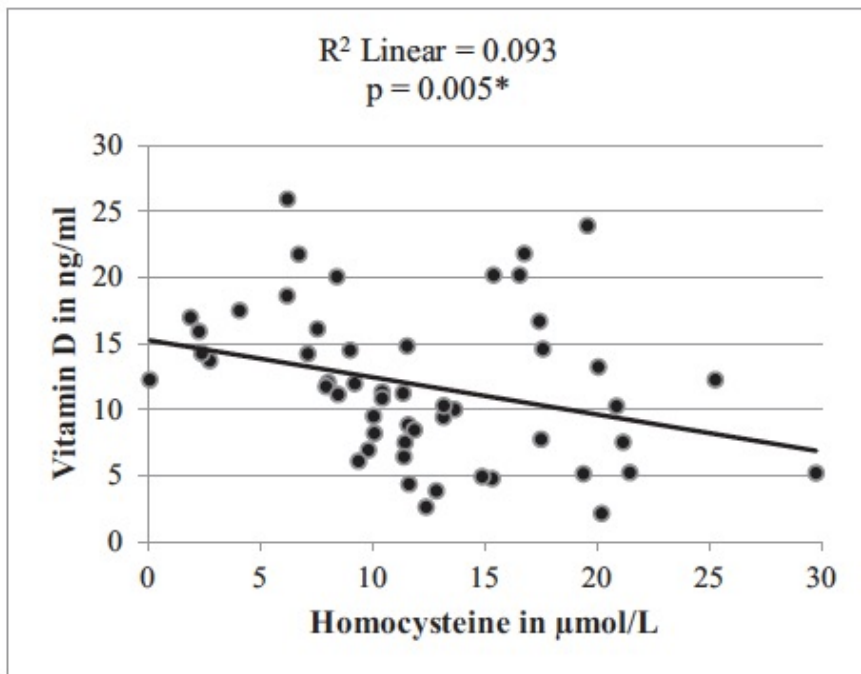
Interplay of vitamin D, vitamin B₁₂, homocysteine and bone mineral density in postmenopausal females

Sundus Tariq^{a,b}, Saba Tariq^{c,d}, and Khalid Parvez Lone^e

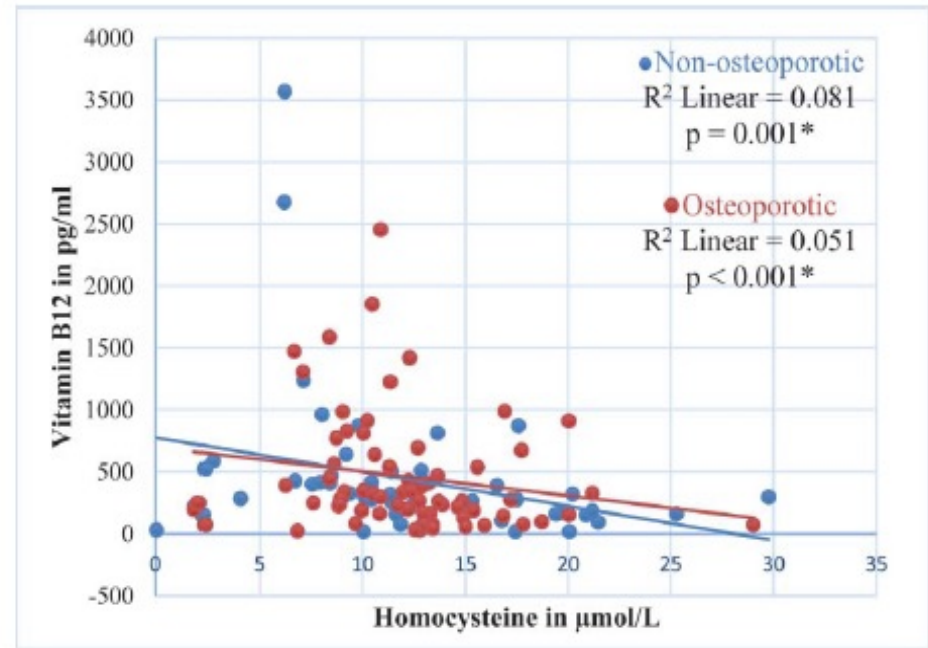
- Tariq, S., Tariq, S., & Lone, K. P. (2018). *Interplay of vitamin D, vitamin B12, homocysteine and bone mineral density in postmenopausal females. Health care for women international, 39(12), 1340-1349.*
- Herrmann, M., Peter Schmidt, J., Umanskaya, N., Wagner, A., Taban-Shomal, O., Widmann, T., ... & Herrmann, W. (2007). *The role of hyperhomocysteinemia as well as folate, vitamin B6 and B12 deficiencies in osteoporosis—a systematic review. Clinical Chemical Laboratory Medicine, 45(12), 1621-1632.*

Homocysteine

Vitamin D, folate, B vitamins



* p -value ≤ 0.05 is considered statistically significant



Vitamin D₃ and B₁₂ levels are inversely correlated to homocysteine levels

HOMOCYSTEINE BALANCE

- POWERFUL COMBINATION
- 7 SPECIFIC NUTRIENTS:
 - *TMG (BETAINE)*
 - *INOSITOL*
 - *CHOLINE*
 - *VITAMIN B6, B12 AND FOLATE*
 - *ZINC*
- NATURAL LEMON-LIME FLAVOUR
- **DOSE:**
 - **2 SCOOPS/DAY**



Important notice...

☐ B₁₂

Proton pump inhibitors (PPI), aspirin, antibiotics and metformin cause vitamin B₁₂ deficit



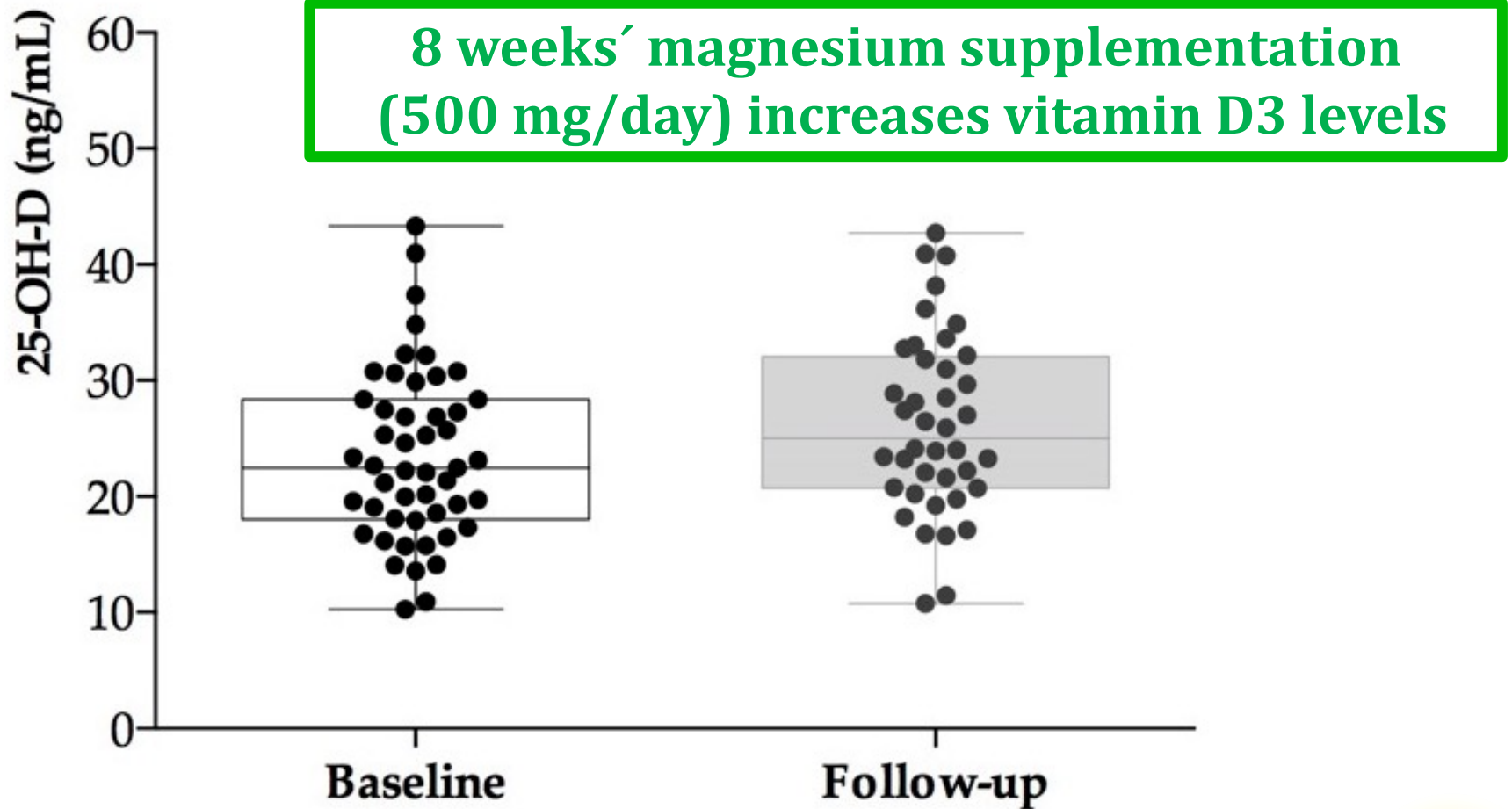
☐ Folate

☐ Metotrexate and alcoholism cause folate deficit

**SUPPLEMENTATION
IS NEEDED**

Osteoporosis

Vitamin D3 & magnesium



Vázquez-Lorente, H., Herrera-Quintana, L., Molina-López, J., Gamarra-Morales, Y., López-González, B., Miralles-Adell, C., & Planells, E. (2020). Response of vitamin D after magnesium intervention in a postmenopausal population from the province of Granada, Spain. *Nutrients*, 12(8), 2283.

Osteoporosis

Vitamin D₃ and magnesium

Vitamin D insufficiency and the blunted PTH response in established osteoporosis: the role of magnesium deficiency

O. Sahota · M. K. Munday · P. San ·
I. M. Godber · D. J. Hosking

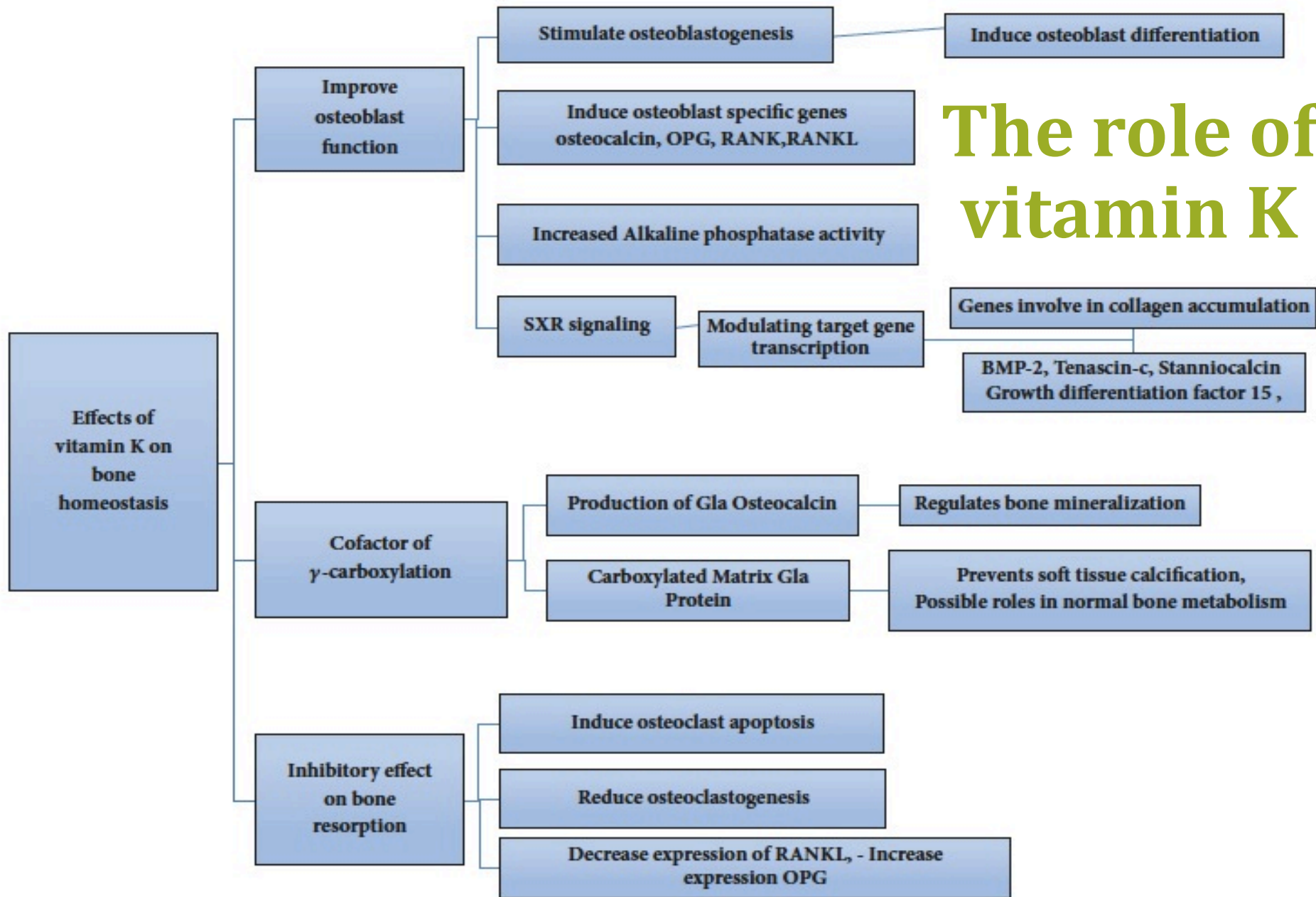
Conclusions: This study confirms that in patients with established osteoporosis, there is also a distinct group with a low vitamin D and a blunted PTH level and that Mg deficiency (as measured by the Mg loading test) is an important contributing factor.

DOSE:

VITAMIN D₃: 4.000-10.000 UI

MAGNESIUM: 300-1000 mg

The role of vitamin K



VITAMIN K2+D3

- POWERFUL
COMBINATION
- WITH MCT
- 1 000 IU VITAMIN D3
PER 4 DROPS
- **DOSE:**
 - 4-16 DROPS/DAY



Anxiety & depression

The role of magnesium

Association between magnesium intake and depression and anxiety in community-dwelling adults: the Hordaland Health Study

Felice N. Jacka, Simon Overland, Robert Stewart, Grethe S. Tell, Ingvar Bjelland, Arnstein Mykletun

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Jacka, F. N., Overland, S., Stewart, R., Tell, G. S., Bjelland, I., & Mykletun, A. (2009). Association between magnesium intake and depression and anxiety in community-dwelling adults: the Hordaland Health Study. Australian & New Zealand Journal of Psychiatry, 43(1), 45-52.

MAGNESIUM BISGLYCINATE PLUS

- 150 mg ELEMENTAL Mg PER CAPSULE
- MOST BIOAVAILABLE FORM OF MAGNESIUM
- FORMULA INCLUDES L-TAURINE FOR INCREASED ABSORPTION
- **DOSE:**
 - 2-6 CAPS/DAY

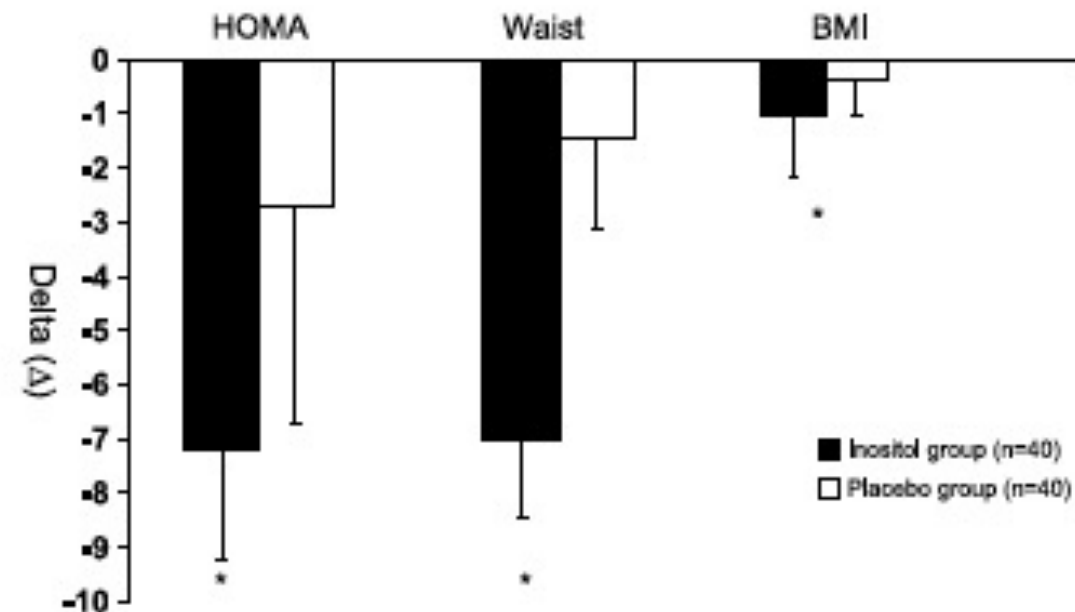


Myo-inositol

BRIEF REPORT **Metabolic syndrome**

Effects of myo-inositol supplementation in postmenopausal women with metabolic syndrome: a perspective, randomized, placebo-controlled study

Domenico Giordano, MD,¹ Francesco Corrado, MD,² Angelo Santamaria, MD,² Simona Quattrone, MD,¹ Basilio Pintaudi, MD,³ Antonino Di Benedetto, MD,³ and Rosario D'Anna, MD²



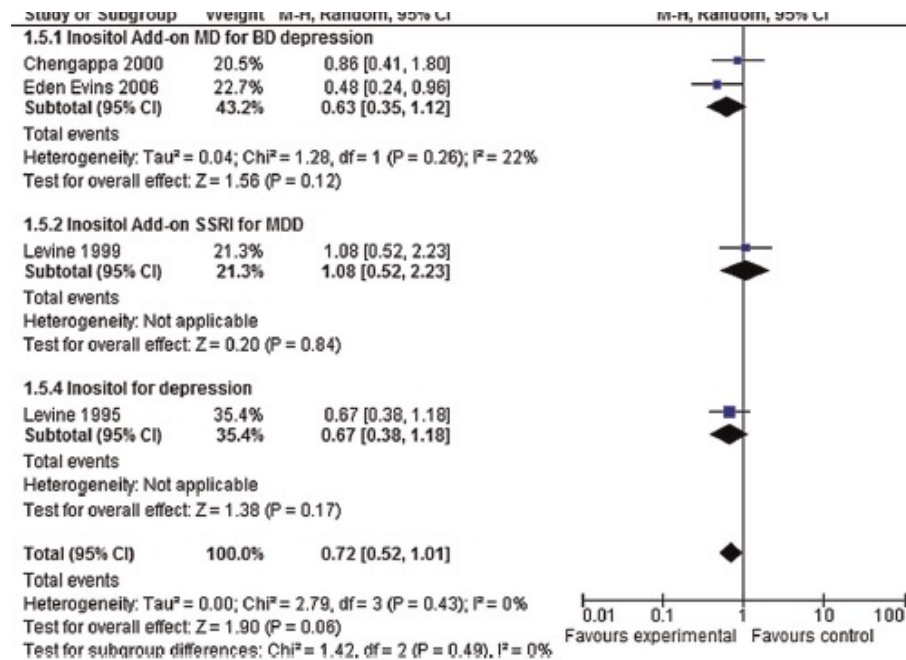
- Improves**
- ✓ HOMA-IR
 - ✓ Waist circumference
 - ✓ BMI
 - ✓ Systolic and diastolic BP
 - ✓ Triglycerides
 - ✓ HDL-C

Myo-inositol Anxiety & depression

A meta-analysis of inositol for depression and anxiety disorders

Tomohiko Mukai[†], Taro Kishi^{*†}, Yuki Matsuda and Nakao Iwata

Department of Psychiatry, School of Medicine, Fujita Health University, Toyoake, Aichi, Japan



Mukai, T., Kishi, T., Matsuda, Y., & Iwata, N. (2014). A meta-analysis of inositol for depression and anxiety disorders. *Human Psychopharmacology: Clinical and Experimental*, 29(1), 55-63.

INOSITOL

- MYO-INOSITOL
- FROM NON-GMO RICE
- 100% NATURAL

- DOSE:
 - 1/2- 2 SCOOPS/DAY

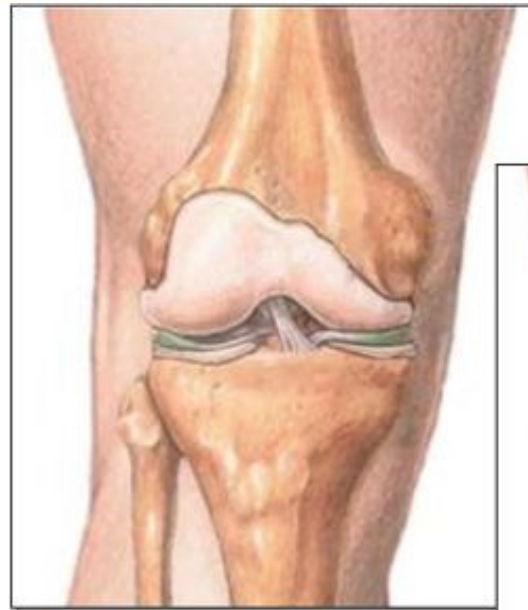


Cimicifuga racemosa

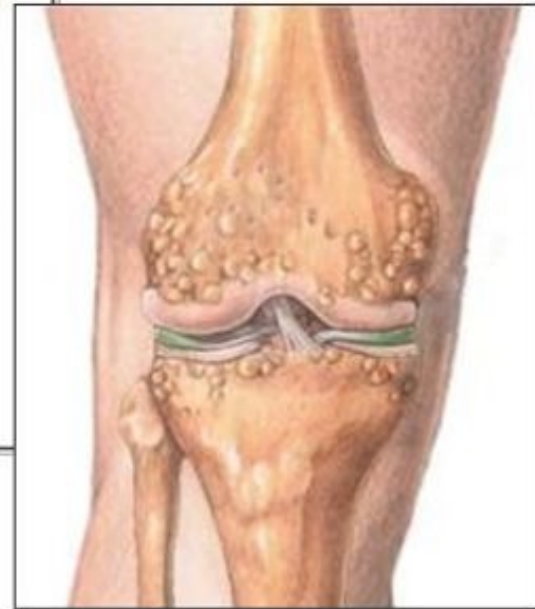
Menopause and bone/joint health

Cimicifuga racemosa and its triterpene-saponins prevent the Metabolic Syndrome and deterioration of cartilage in the knee joint of ovariectomized rats by similar mechanisms

Dana Seidlova-Wuttke^a, Nicole Eder^a, Vera Stahnke^a, Markus Kammann^c, Günter Stecher^c, Jutta Haunschild^c, Johannes T. Wessels^b, Wolfgang Wuttke^{a,*}



Healthy Knee



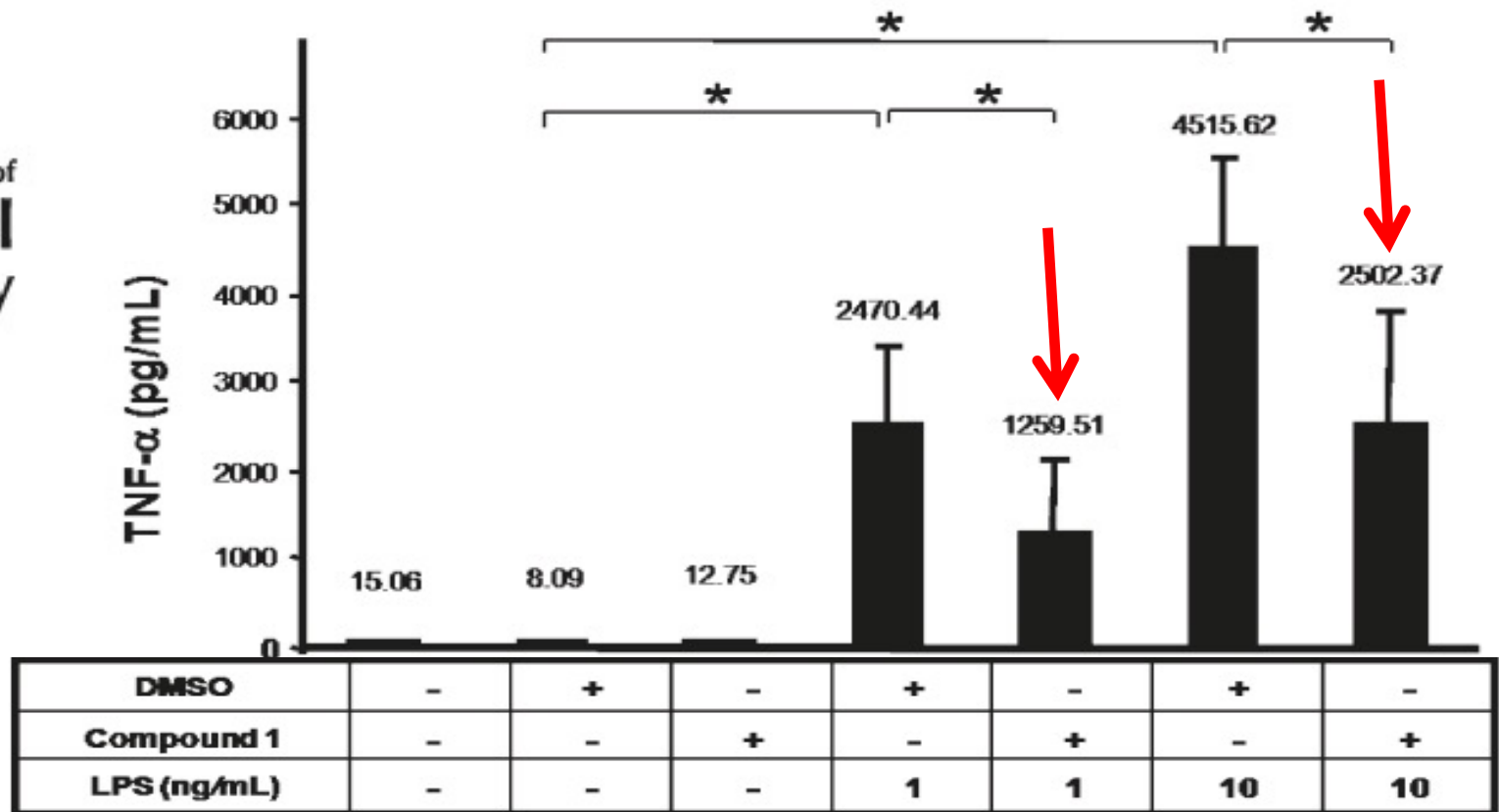
Arthritic Knee

Cimicifuga racemosa

Menopause and bone/joint health

Identification of the Bioactive Constituent and Its Mechanisms of Action in Mediating the Anti-Inflammatory Effects of Black Cohosh and Related *Cimicifuga* species on Human Primary Blood Macrophages

Journal of
**Medicinal
Chemistry**
Article



Yang, C. L., Chik, S. C., Li, J. C., Cheung, B. K., & Lau, A. S. (2009). Identification of the bioactive constituent and its mechanisms of action in mediating the anti-inflammatory effects of black cohosh and related *Cimicifuga* species on human primary blood macrophages. *Journal of medicinal chemistry*, 52(21), 6707-6715.

Cimicifuga racemosa

Hot flushes

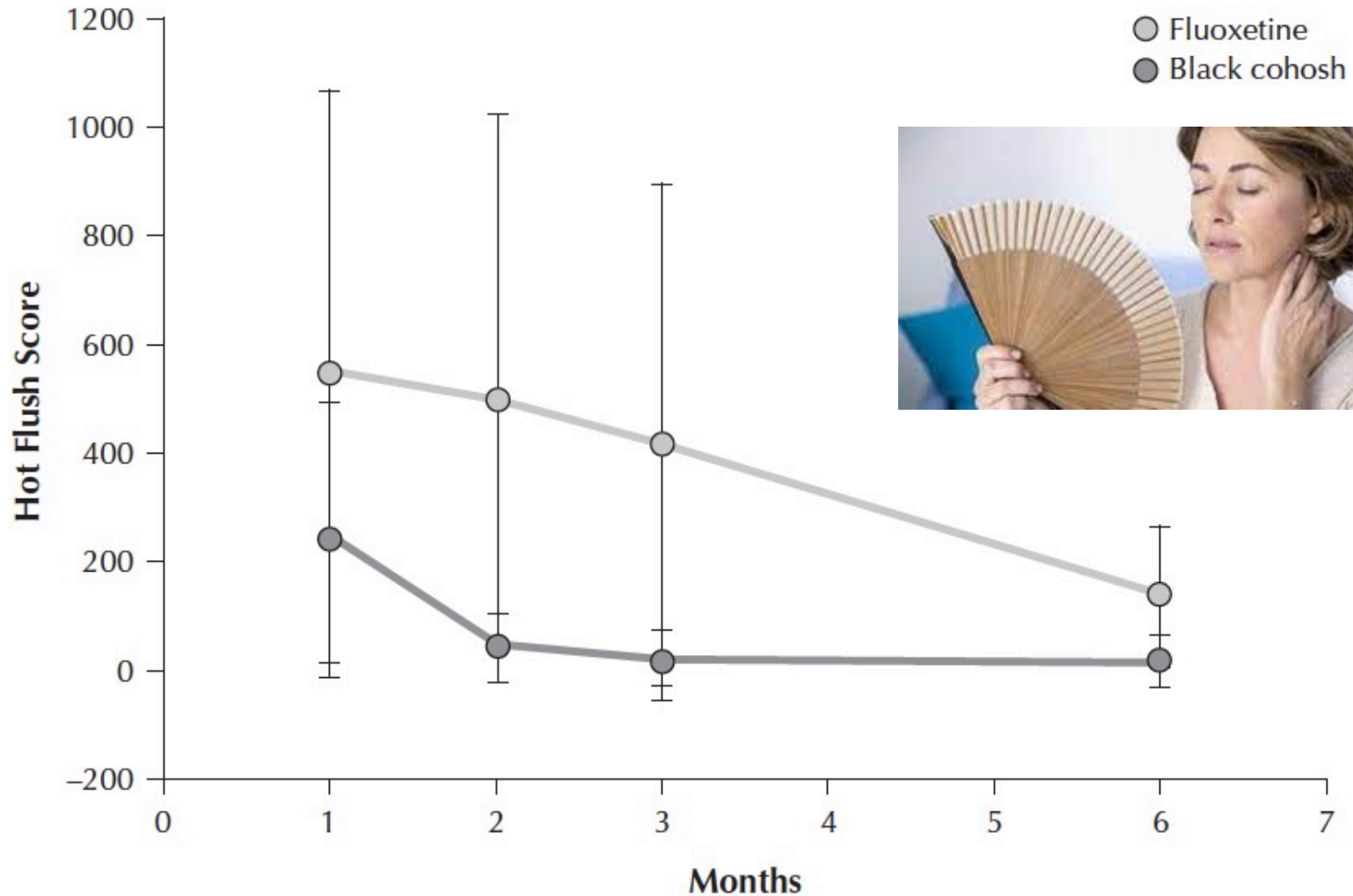
Variable	Black Cohosh (n=40)				Fluoxetine (n=40)			
	1 mo	2 mo	3 mo	6 mo	1 mo	2 mo	3 mo	6 mo
Hot flush score	239.3±256.2	39.8±67.2	19.2±51.9	18.7±49.0	542.6±528.7	499.6±527.1	417.8±481.0	134.8±133.7
Night sweat score	211.2±265.9	43.4±91.5	30.5±86.9	21.3±60.1	267.8±182.1	228.2±150.5	197.7±152.8	106.3±150.3



Oktem, M., Eroglu, D., Karahan, H. B., Taskintuna, N., Kuscu, E., & Zeyneloglu, H. B. (2007). Black cohosh and fluoxetine in the treatment of postmenopausal symptoms: a prospective, randomized trial. *Advances in therapy*, 24(2), 448-461.

Cimicifuga racemosa

Hot flushes



Oktem, M., Eroglu, D., Karahan, H. B., Taskintuna, N., Kuscu, E., & Zeyneloglu, H. B. (2007). Black cohosh and fluoxetine in the treatment of postmenopausal symptoms: a prospective, randomized trial. *Advances in therapy*, 24(2), 448-461.

Cimicifuga racemosa

Woman's health

- ✓ Improves hot flushes and night sweats
- ✓ Improves vaginal dryness
- ✓ Oestrogenic activity (regulatory)
- ✓ Anti-histamine properties
- ✓ Serotonergic activity
- ✓ Improves mood disturbances
- ✓ Improves osteoblast and chondroblast turnover



- *Wuttke, W., Gorkow, C., & Seidlová-Wuttke, D. (2006). Effects of black cohosh (Cimicifuga racemosa) on bone turnover, vaginal mucosa, and various blood parameters in postmenopausal women: a double-blind, placebo-controlled, and conjugated estrogens-controlled study. Menopause, 13(2), 185-196.*
- *Wuttke, W., Seidlova-Wuttke, D., & Gorkow, C. (2003). The Cimicifuga preparation BNO 1055 vs. conjugated estrogens in a double-blind placebo-controlled study: effects on menopause symptoms and bone markers. Maturitas, 44, S67-S77.*

Menopause Isoflavones

Review

Utilization of Isoflavones in Soybeans for Women with Menopausal Syndrome: An Overview

Li-Ru Chen ^{1,2} and Kuo-Hu Chen ^{3,4,*} 

Regarding the therapeutic effects on menopausal syndrome or other diseases, isoflavones have been found to alleviate vasomotor syndromes even after considering placebo effects, reduce bone loss in the spine and ameliorate hypertension and in vitro glycemic control. They may also alleviate

**Useful for hot flushes, night sweats and
bone loss in the spine**

Phytotherapy in menopause

- ✓ *Vitex agnus castus* (chasteberry)
- ✓ *Dong quai*
- ✓ *Cimicifuga racemosa*
- ✓ *Salvia officinalis*
- ✓ Dandelion
- ✓ Isoflavones from: red clover, *dioscorea*
- ✓ *Humulus lupulus* (hops)

- TAGHIZADEH, ZIBA, et al. *The study of the effect of vitex agnus-castus on the early menopausal complications in menopausal women.* 2006.
- Pitkin, J. (2012). *Alternative and complementary therapies for the menopause.* *Menopause international*, 18(1), 20-27.

MENOPEACE

- POWERFUL SYNERGY OF 6 PLANTS:
 - DONG QUAI
 - CHASTE-TREE (AGNUS CASTUS)
 - RED CLOVER
 - DANDELION
 - BLACK COHOSH
 - DIOSCOREA VILOSA
- DOSE:
 - 1-4 DAY



Women's health

Ashwagandha

Body Weight Management in Adults Under Chronic Stress Through Treatment With Ashwagandha Root Extract: A Double-Blind, Randomized, Placebo-Controlled Trial

- ✓ Improves cortisol & ACTH levels
- ✓ Improves perceived stress and cravings
- ✓ Decreases BMI & weight

Choudhary, D., Bhattacharyya, S., & Joshi, K. (2017). Body weight management in adults under chronic stress through treatment with Ashwagandha root extract: a double-blind, randomized, placebo-controlled trial. Journal of evidence-based complementary & alternative medicine, 22(1), 96-106.

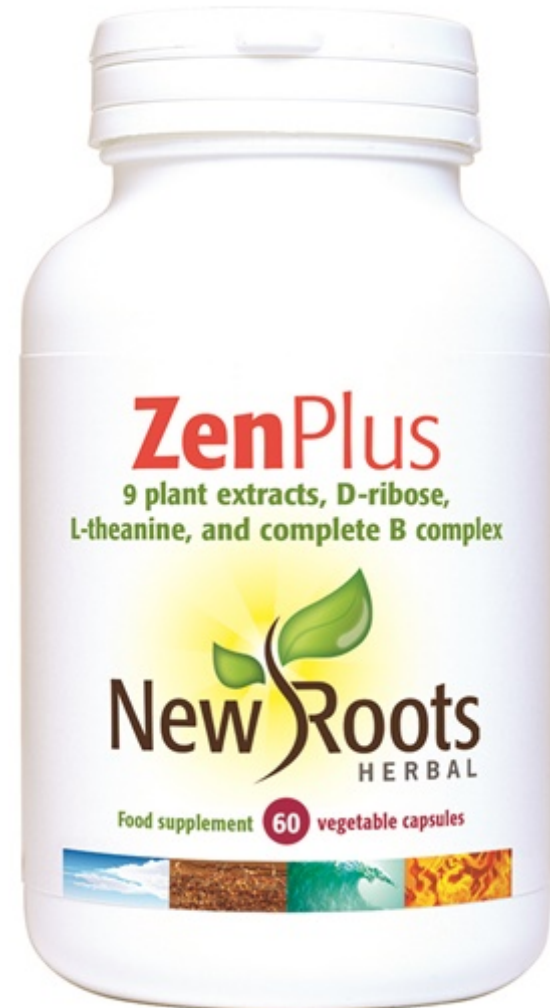
ZEN PLUS

- POWERFUL COMBINATION:

- *L-THEANINE*
- *ASHWAGHANDHA*
- *ASTRAGALUS*
- *MAGNOLIA*
- *PASSIFLORA*
- *RHODIOLA*
- *VITAMIN B COMPLEX*

- DOSE:

- **2 CAPS/DAY**





Case study

- ✓ *María, 50 years old*
- ✓ *Lack of period since 2 years*
- ✓ *Hot flushes and night sweats*
- ✓ *Sadness & higher anxiety perception*
- ✓ *Sleep disturbances (wakes up 3-4 times)*
- ✓ *Diet: low protein & fat, high carbohydrate intake*
- ✓ *Blood tests: high glucose, low estradiol, low progesterone, low vitamin D levels*

Case study

Intervention

- ✓ *Increase protein & fat in the diet*
- ✓ *Lower carbohydrate intake*
- ✓ *Educate with sleep routine*
- ✓ *Meditation and increase physical activity*
- ✓ *2 g EPA+DHA/day WITH a meal*
- ✓ *VITAMIN D₃+K₂, 16 drops/day WITH a meal*
- ✓ *MENOPEACE, 3 caps/day*
- ✓ *INOSITOL, 4 g/day (divided in 2-3 meals)*

Case study

After 2 months

- ✓ *Maintain eating habits*
- ✓ *Incorporate strength training and dancing*
- ✓ *VITAMIN D₃+K₂, 3 drops/day WITH a meal*
- ✓ *MENOPEACE, 2 caps/day*
- ✓ *Continue helping with stress management
(anxiety levels have decreased), ZEN PLUS, 2/day*
- ✓ *Add MAGNESIUM BISGLYCINATE, 3 caps/day*

What is the solution?

RELATIONSHIPS

BIOLOGY

FOOD & NUTRITION

GENETICS



ENVIRONMENT

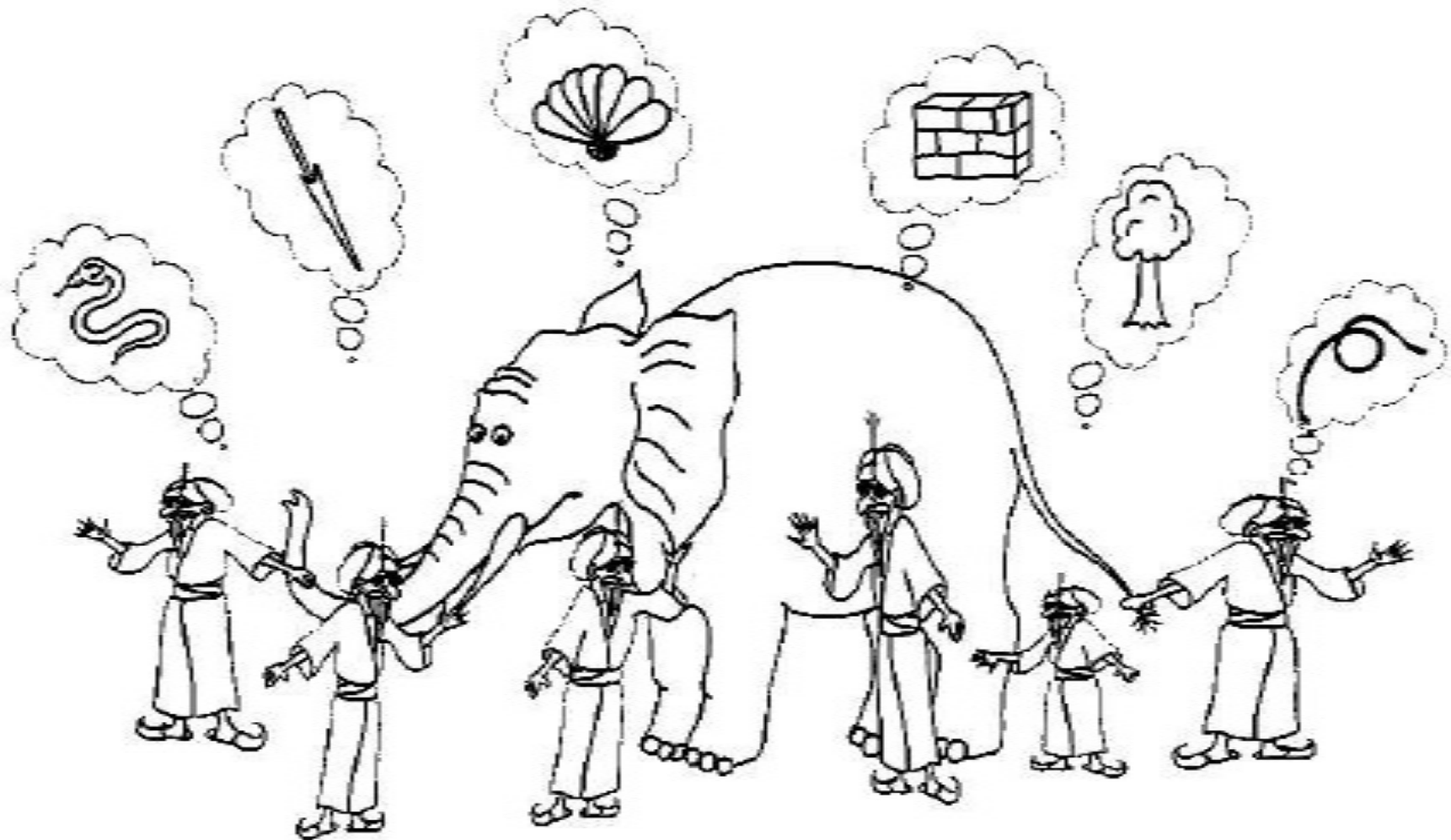
IKIGAI

MOVEMENT

EVOLUTIONARY COHERENCE!

Ruiz-Núñez, B et al.(2013). Lifestyle and nutritional imbalances associated with Western diseases: causes and consequences of chronic systemic low-grade inflammation in an evolutionary context. The Journal of nutritional biochemistry, 24(7), 1183-1201.

The Blind man and the Elephant





Dr Begoña Ruiz Núñez



Iker Martínez Pérez



Verónica Martínez Vázquez



Javier Morán Tiesta



Beatriz Amusategui Moya



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Where can I buy the products from New Roots Herbal?

- ✓ In Ireland through 'Pure & Natural' ---> www.pureandnatural.ie
- ✓ In the UK through 'Amrita Nutrition' & 'The Natural Dispensary'
www.amritanutrition.co.uk & www.naturaldispensary.co.uk



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