



2. kongres
klinične prehrane in
presnovne podpore
z mednarodno udeležbo

2nd Congress of clinical nutrition and metabolic care

with international participation

Grand Hotel Bernardin Portorož, 15th to 17th November 2013



Proteins and physical activity: specificities in different age groups

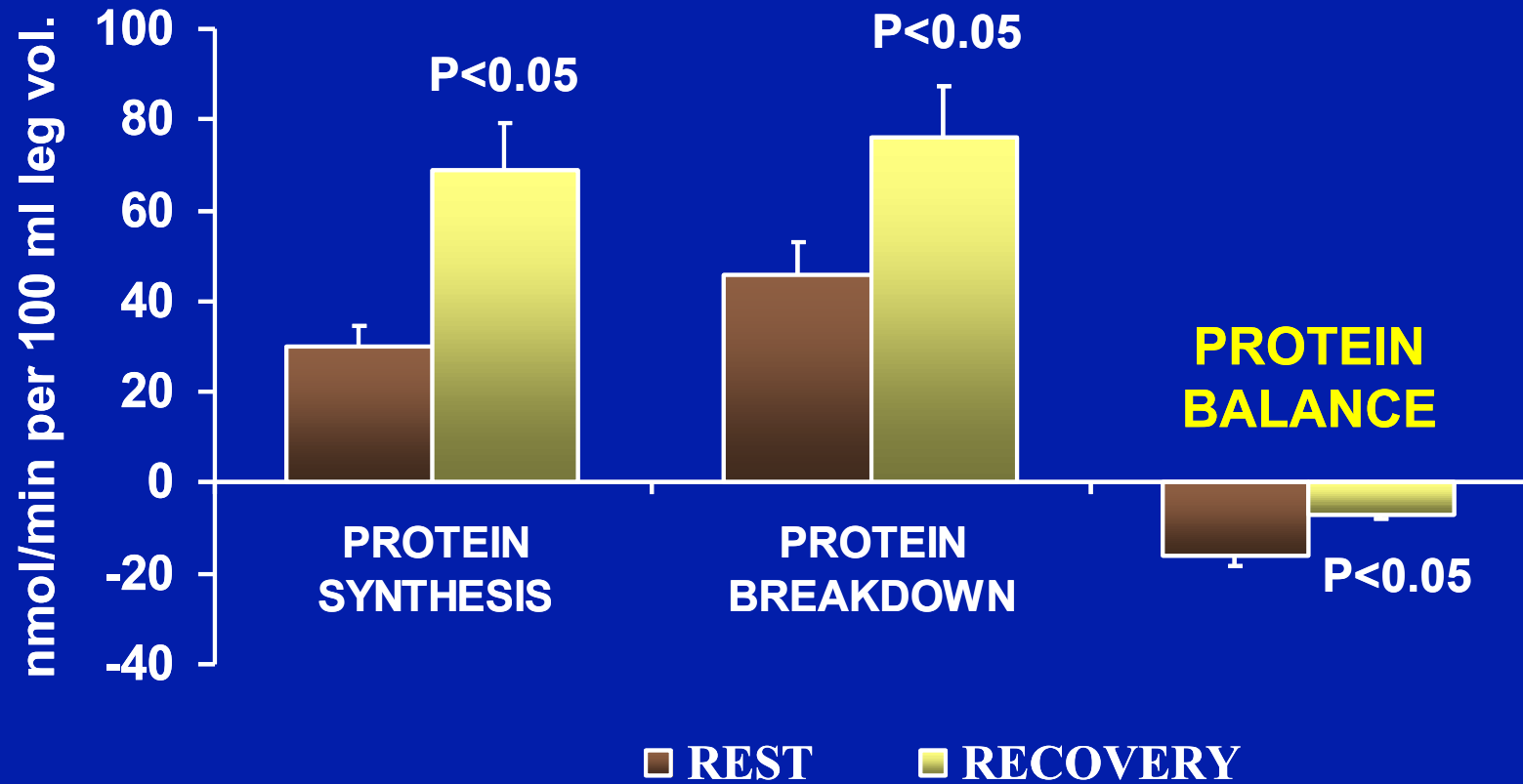
Gianni Biolo

Department of Medical Sciences

Clinica Medica AOUST

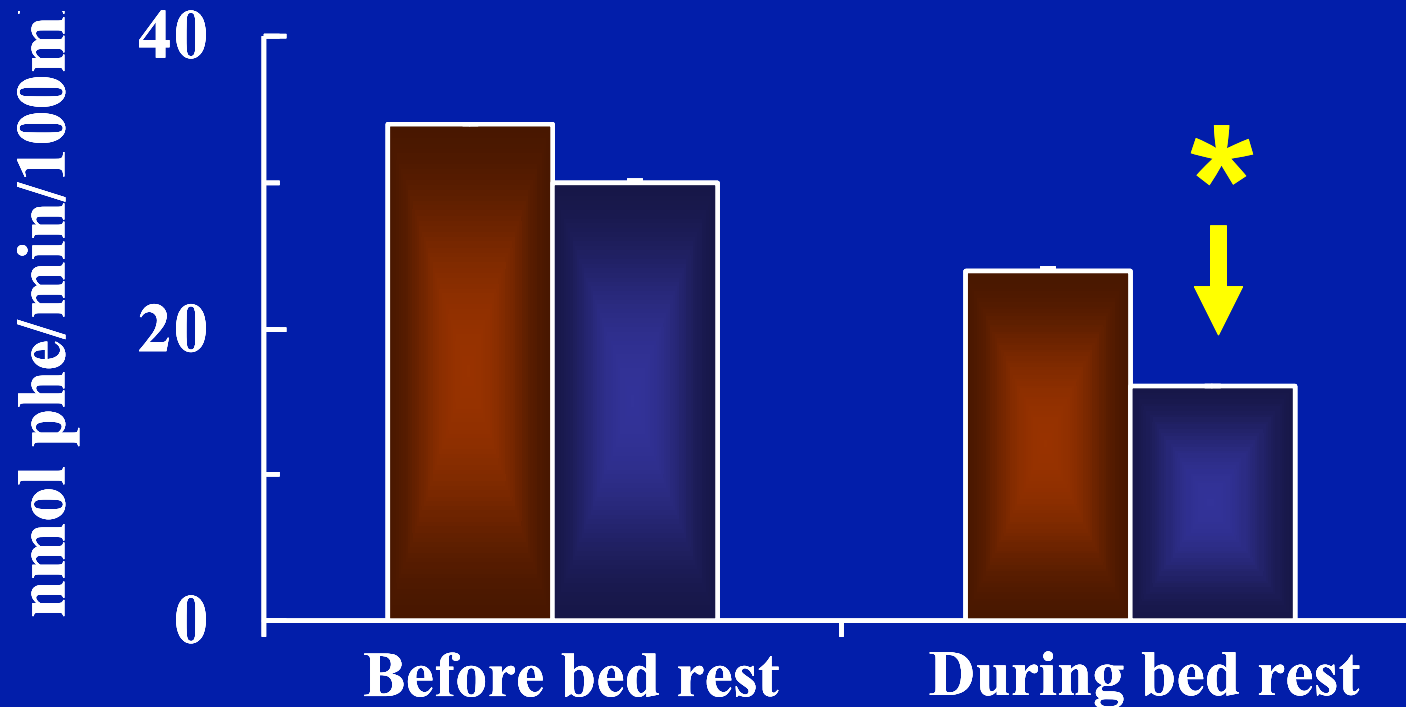
University of Trieste - Italy

Muscle protein kinetics during post-exercise recovery



Prolonged Bed Rest Decreases Skeletal Muscle and Whole Body Protein Synthesis

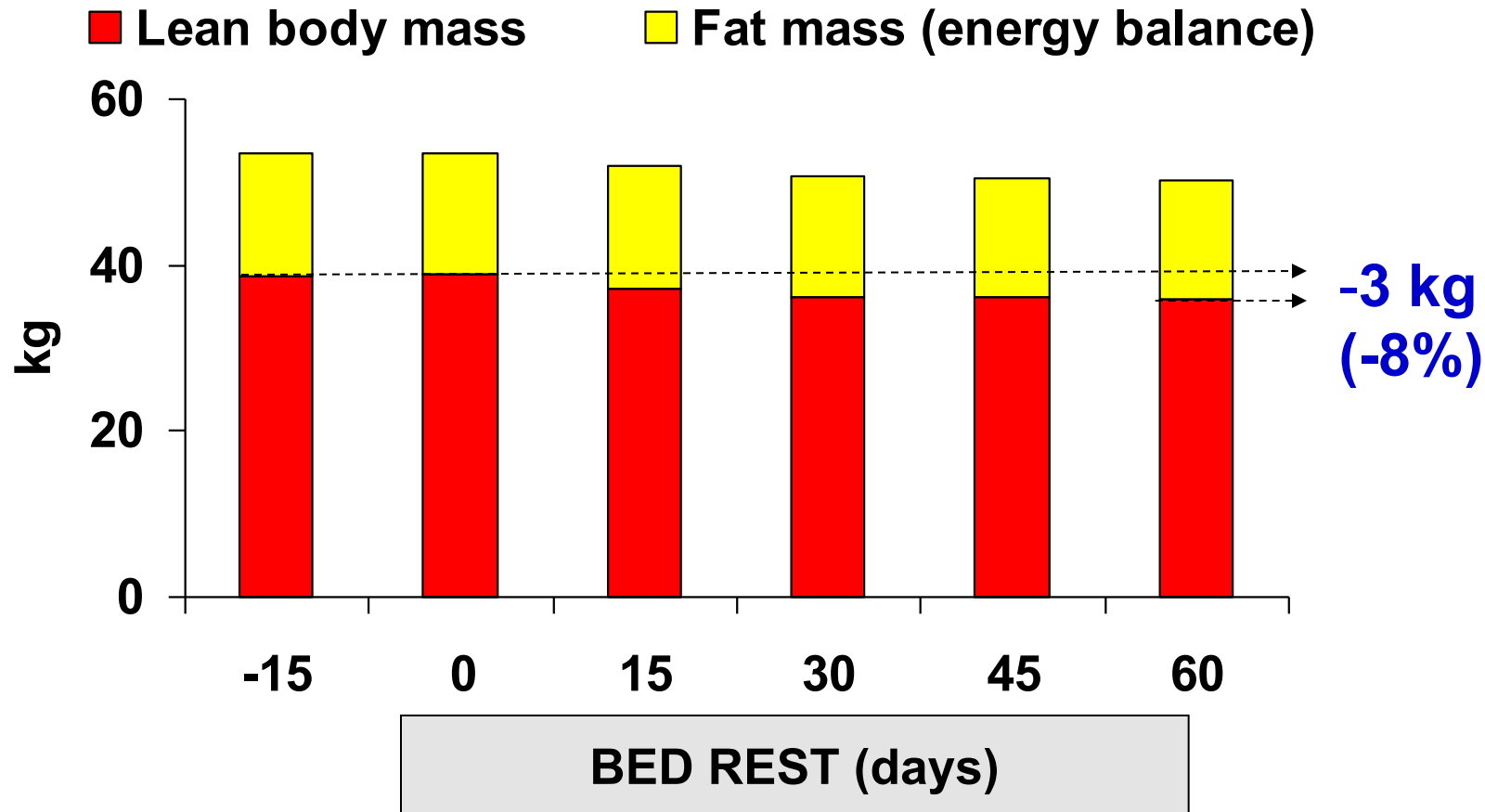
A. Ferrando AJPENDO 1996



■ Protein Degradation ■ Protein Synthesis

Dual energy X-ray absorptiometry (DXA)

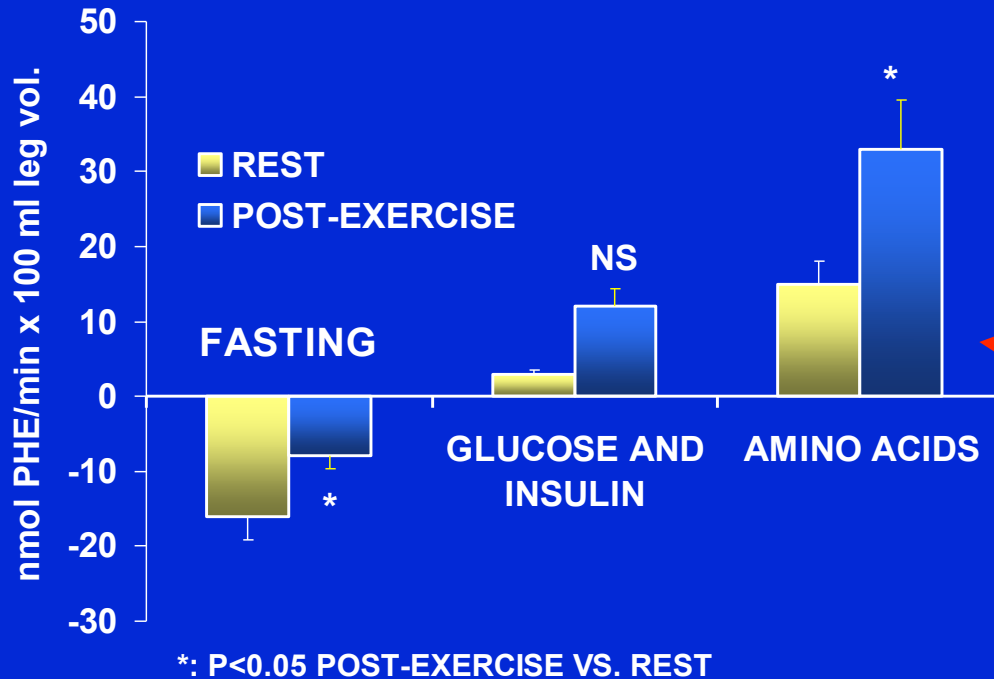
CHANGES IN BODY COMPOSITION
DURING TWO-MONTH EXPERIMENTAL BED REST IN WOMEN (n=8)



**EFFECTS OF EXERCISE AND
BED REST ON
POSTPRANDIAL ANABOLIC
EFFICIENCY**

NUTRITION AND POST-EXERCISE MUSCLE ANABOLISM

MUSCLE PROTEIN BALANCE



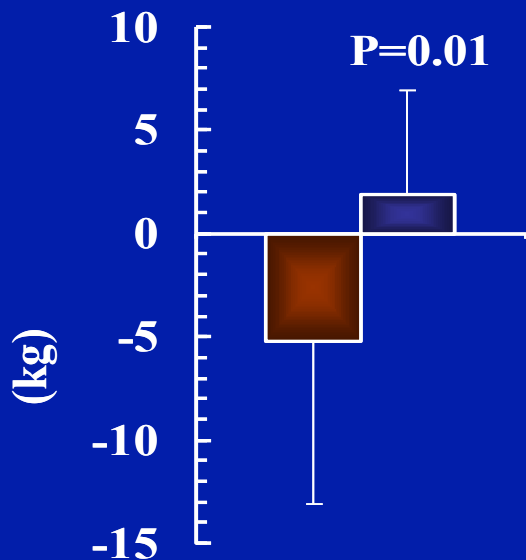
The rates of post-prandial amino acid deposition into body protein is greatly accelerated after resistance exercise.

RESISTANCE TRAINING TO COUNTERACT THE CATABOLISM OF A LOW-PROTEIN DIET IN PATIENTS WITH CHRONIC RENAL INSUFFICIENCY

Castaneda et al., Ann Intern Med 2001

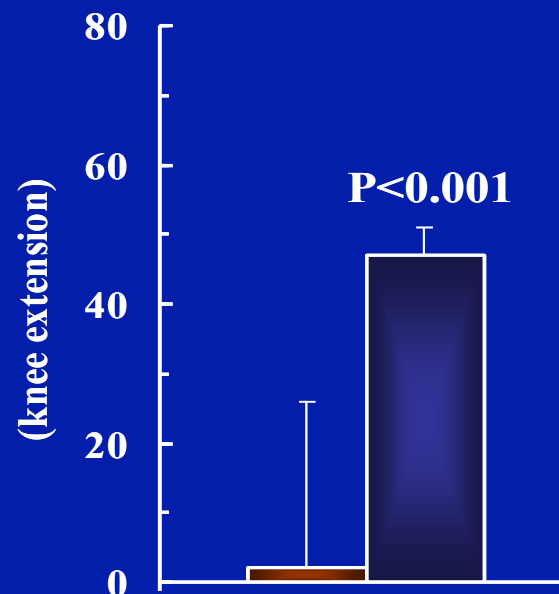
26 OLDER PATIENTS (ABOUT 65 YR) WITH MODERATE RENAL INSUFFICIENCY WHO HAD ACHIEVED STABILIZATION ON A LOW-PROTEIN DIET (0.6 G/KG/DAY) WERE RANDOMLY ASSIGNED TO RESISTANCE EXERCISE TRAINING OR NO INTERVENTION FOR 12 WEEKS.

Absolute change in total body potassium (kg)



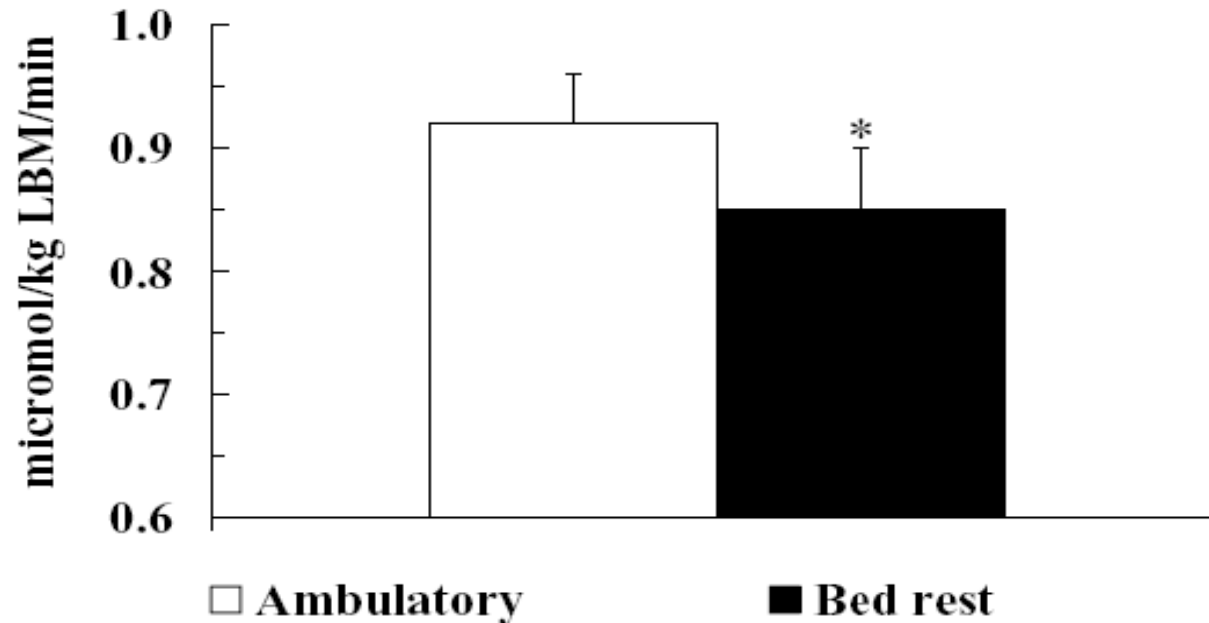
- Low-protein diet alone
- Low-protein diet plus resistance training

Percent change in lower body strength (knee extension)



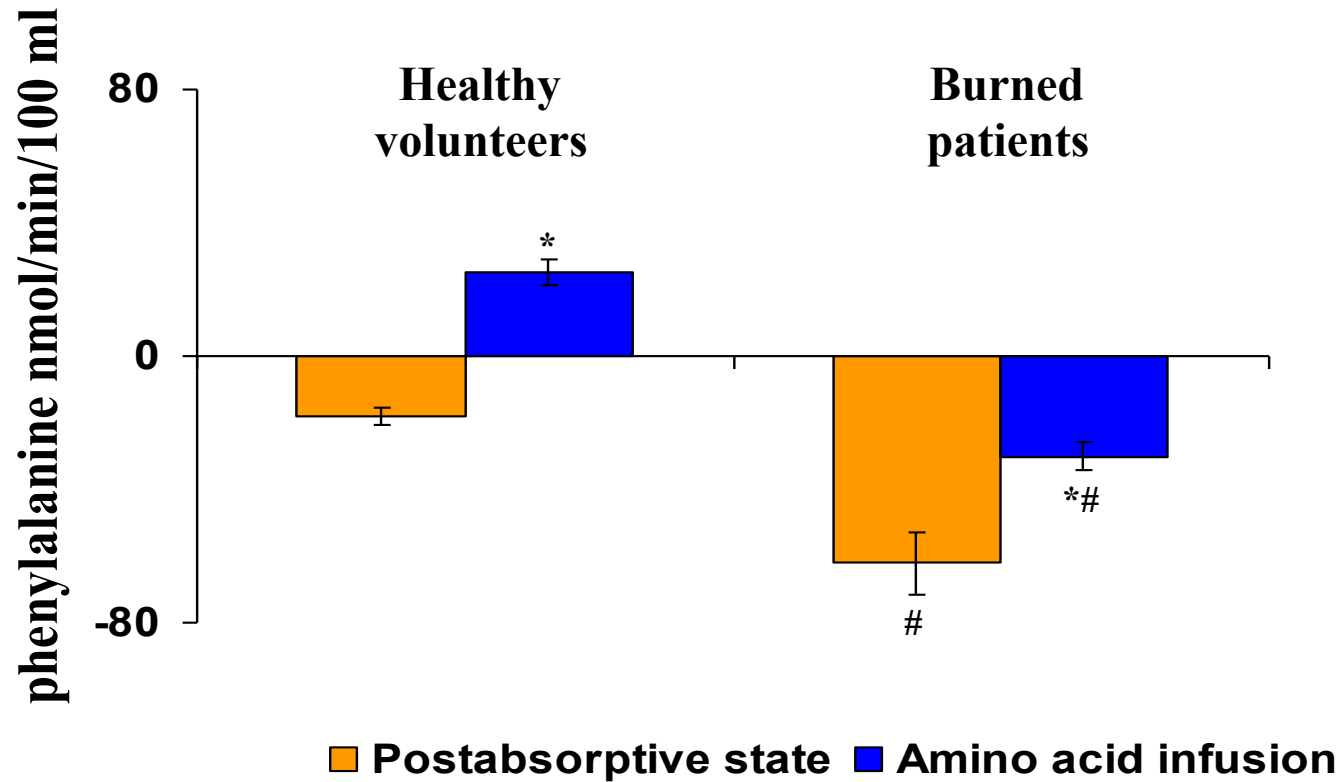
SHORT-TERM BED REST IMPAIRS AMINO ACID-INDUCED PROTEIN ANABOLISM IN HUMANS

Biolo et al., J Physiol 2004



The rates of post-prandial amino acid deposition into body protein is impaired in bed rest conditions.

EFFECTS OF AMINO ACID INFUSION ON SKELETAL MUSCLE PROTEIN BALANCE IN SEVERELY BURNED PATIENTS



*, $P < 0.05$ vs. postabsorptive state #, $P < 0.05$ vs. healthy volunteers

Aging is associated with diminished accretion of muscle proteins after the ingestion of a small bolus of essential amino acids¹⁻³

Christos S Katsanos, Hisamine Kobayashi, Melinda Sheffield-Moore, Asle Aarland, and Robert R Wolfe

Am J Clin Nutr 2005;82:1065-73.

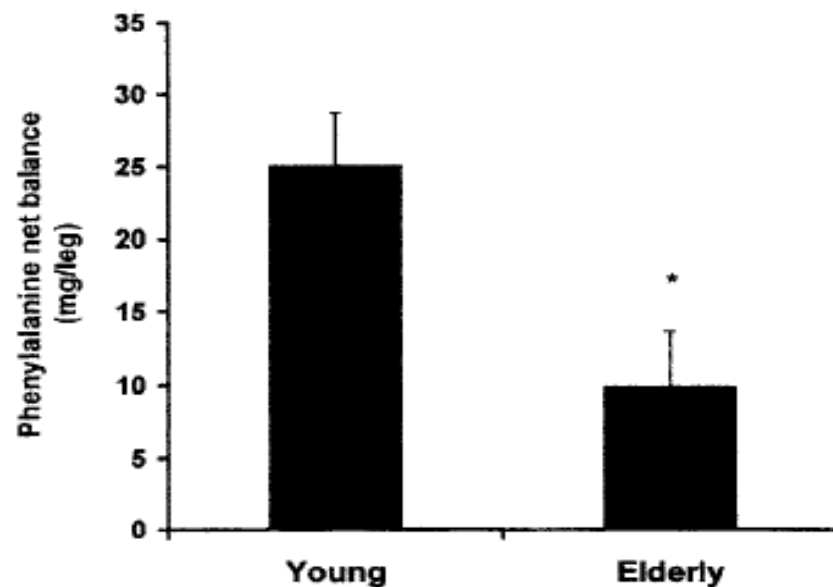


FIGURE 5. Mean (\pm SEM) leg phenylalanine net balance 3.5 h after the ingestion of essential amino acids calculated by measuring the area under the phenylalanine net balance response curve (in the calculations, basal net balance was taken as zero) in the elderly ($n = 11$) and the young ($n = 8$). Data were analyzed with a *t* test. *Significantly different from the young, $P = 0.010$.

Protein-containing nutrient supplementation following strength training enhances the effect on muscle mass, strength, and bone formation in postmenopausal women

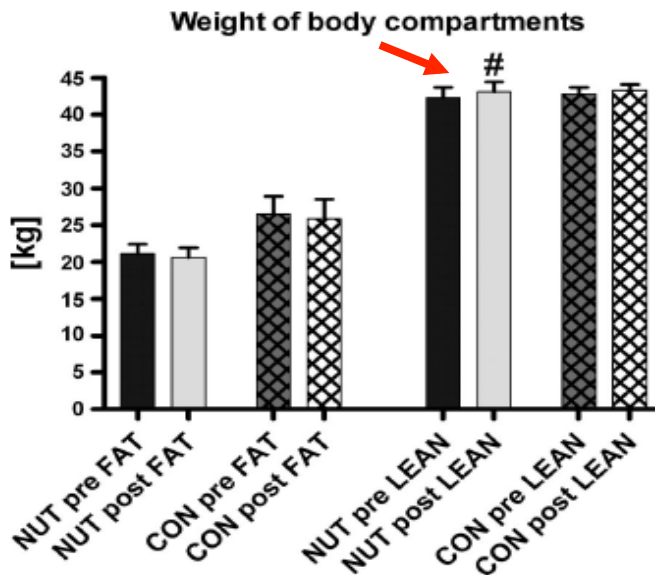
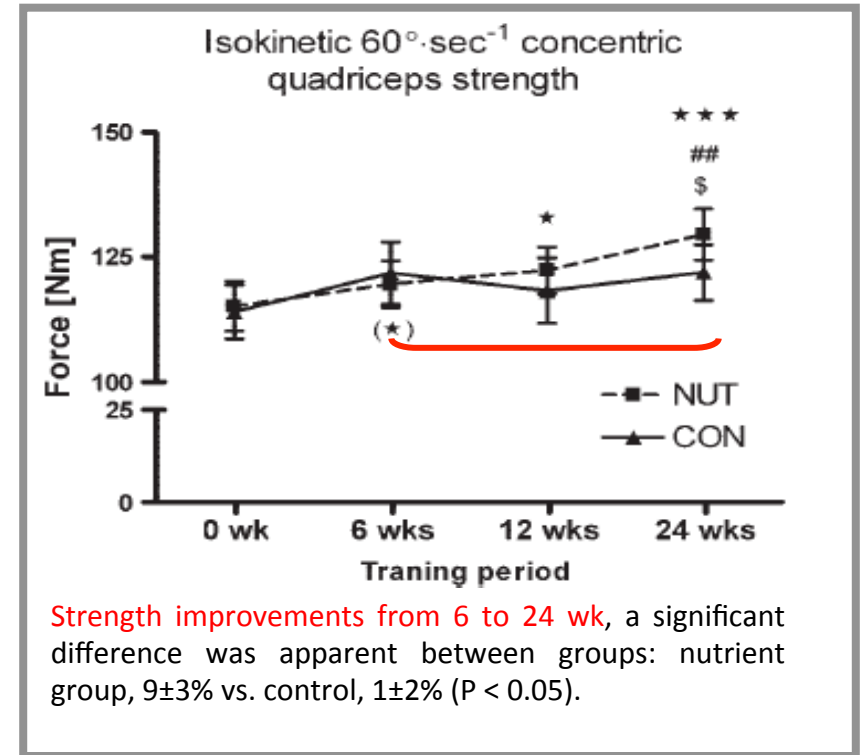
J Appl Physiol 105: 274–281, 2008.

Table 4. Bone mineral density

	Control Group	Nutrient Group
Total BMD, g/mm ³		
0 wk	1.117±0.022	1.113±0.027
24 wk	1.122±0.023	1.116±0.027
Femoral neck BMD, g/mm ³		
0 wk	0.943±0.028	0.953±0.051
24 wk	0.930±0.024	0.978±0.043
L2–L4 BMD, g/mm ³		
0 wk	1.043±0.032	1.084±0.053
24 wk	1.068±0.038*	1.108±0.049*

Values are means ± SE of bone mineral density (BMD) at whole body, femoral neck, and lumbar spine (L2–L4). *P < 0.05 compared with 0 wk.

Adjusting for covariates (age at inclusion, BMI at inclusion, and BMD of the femoral neck at inclusion) a significant (P < 0.05) difference was seen in the response to training between the two groups.



Dietary omega-3 fatty acid supplementation increases the rate of muscle protein synthesis in older adults: a randomized controlled trial¹⁻³

Gordon I Smith, Philip Atherton, Dominic N Reeds, B Selma Mohammed, Debbie Rankin, Michael J Rennie, and Bettina Mittendorfer

Am J Clin Nutr 2011;93:402-12.

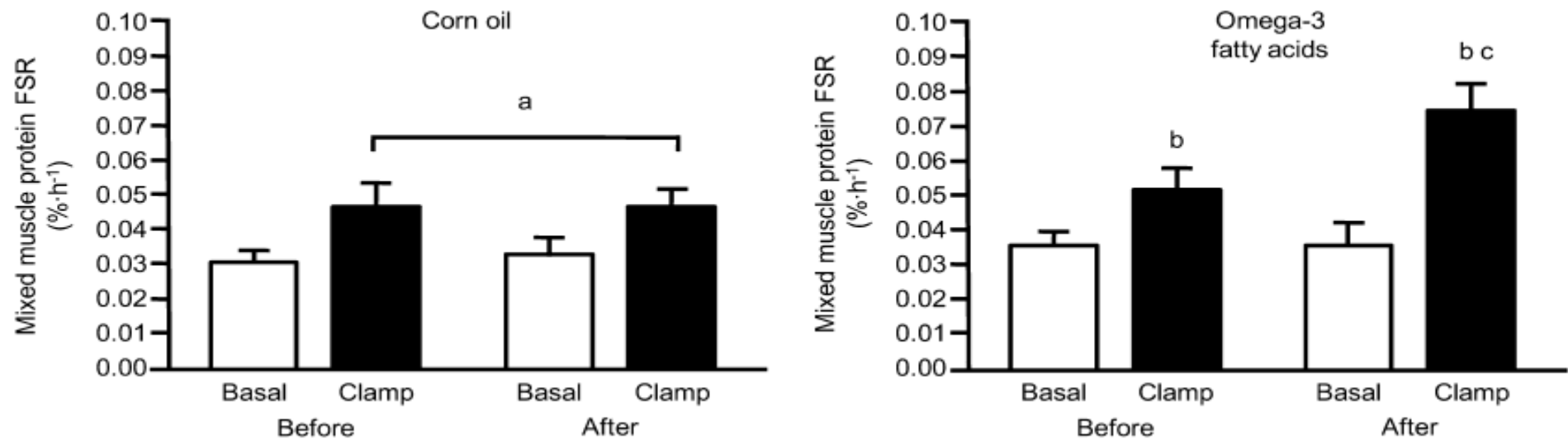


FIGURE 1. Mean (\pm SEM) mixed skeletal muscle protein fractional synthesis rate (FSR), calculated by using the average plasma free phenylalanine enrichment as the precursor pool enrichment, during basal, postabsorptive conditions and during the hyperaminoacidemic-hyperinsulinemic clamp before and after 8 wk of supplementation with either corn oil ($n = 7$) or omega-3 fatty acids ($n = 8$). There was no difference in the muscle protein FSR between the omega-3 fatty acid and corn oil groups before the intervention [ANOVA showed a significant effect of clamp ($P < 0.001$), no significant effect of group ($P = 0.47$), and no interaction ($P = 0.60$)]. ^aIn the corn oil group, ANOVA showed a significant main effect of clamp ($P < 0.01$). In the omega-3 fatty acid group, ANOVA showed a significant effect of clamp ($P < 0.01$) and an interaction ($P < 0.001$), which was followed by Tukey's post hoc analysis. ^bSignificantly different from the corresponding basal value, $P < 0.01$. ^cSignificantly different from the corresponding value before omega-3 fatty acid supplementation, $P < 0.01$. Furthermore, the before-after intervention change in the anabolic response (increase in the muscle protein FSR from basal values) was significantly greater in the omega-3 fatty acid group than in the corn oil group ($P = 0.01$, Student's t test for independent samples).

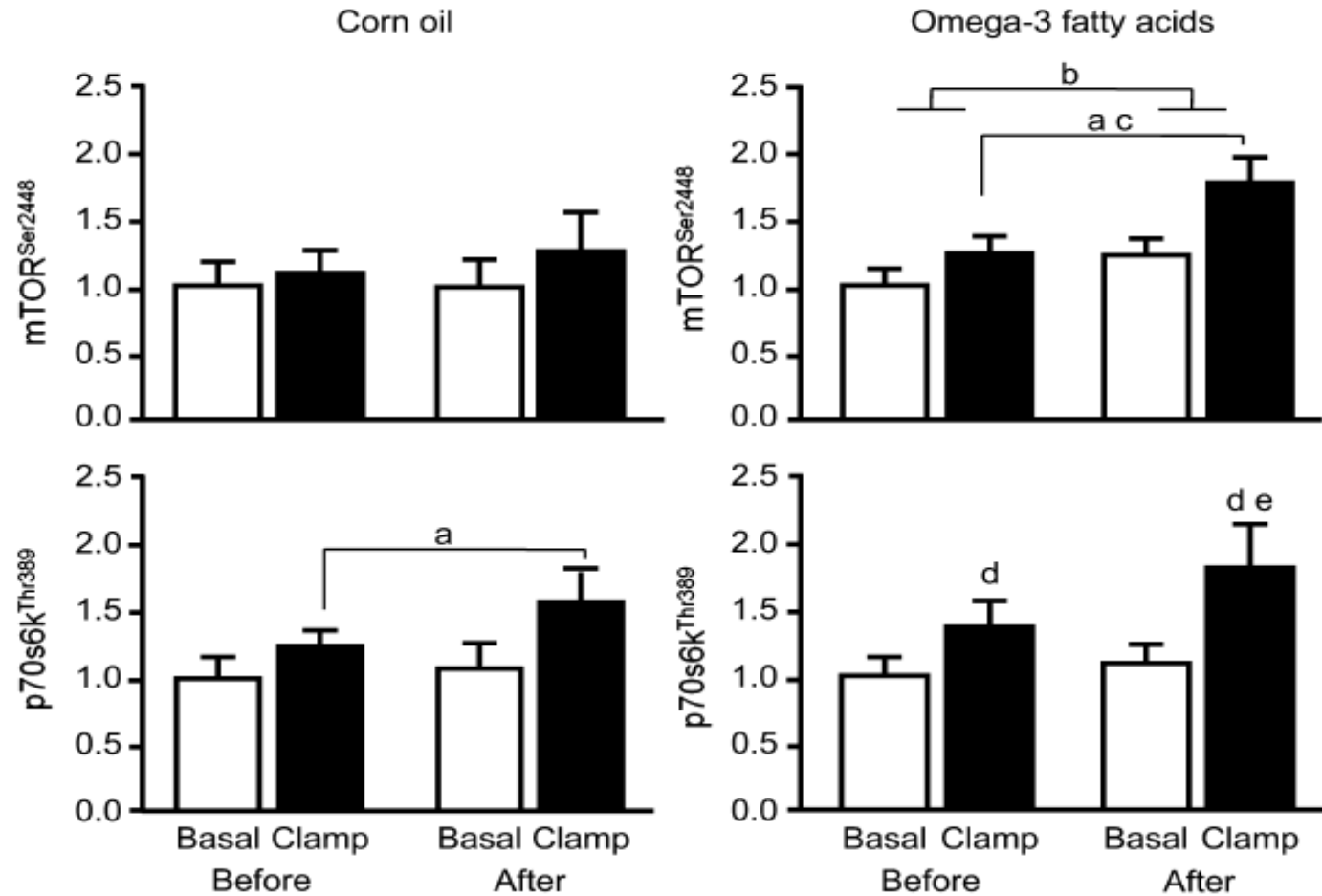
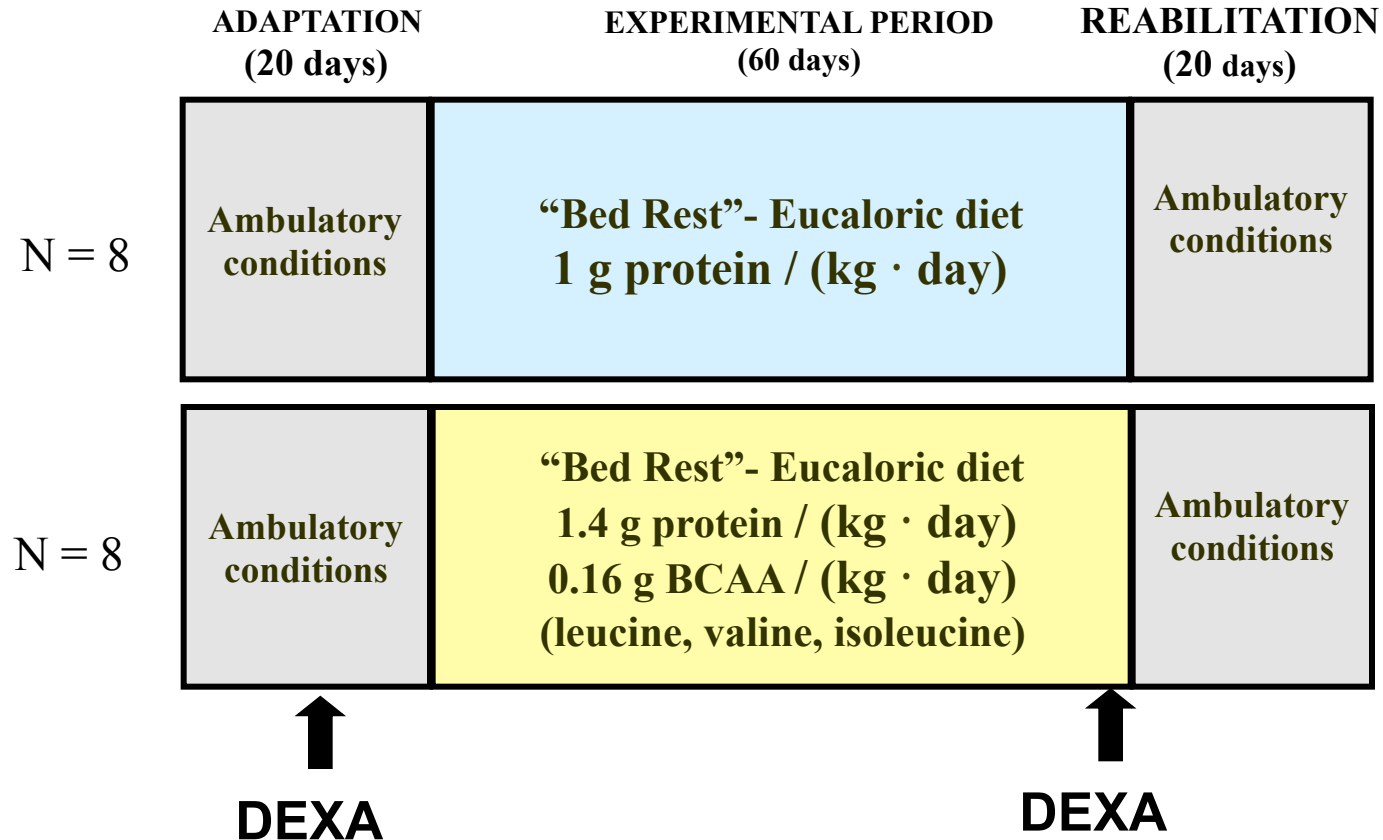
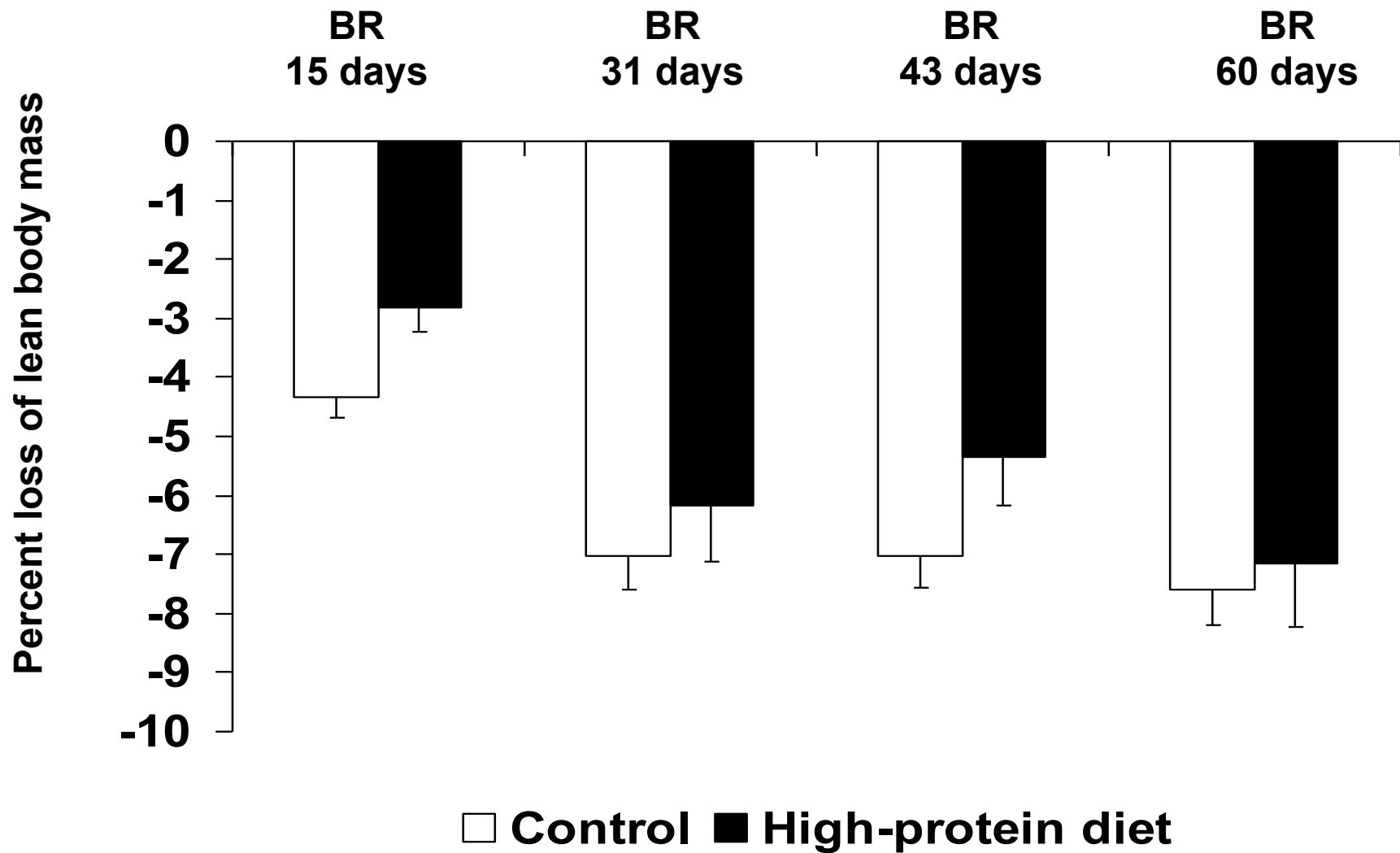


FIGURE 2. Mean (\pm SEM) concentrations (arbitrary units) of mTOR^{Ser2448} and p70s6k^{Thr389} during basal, postabsorptive conditions and during the hyperaminoacidemic-hyperinsulinemic clamp before and after 8 wk of supplementation with either corn oil ($n = 7$) or omega-3 fatty acids ($n = 8$). ^aANOVA showed a significant main effect of clamp ($P < 0.01$). ^bANOVA showed a significant main effect of time ($P < 0.05$). ^cThere was a trend for a greater clamp-induced increase in mTOR^{Ser2448} after omega-3 fatty acid supplementation than before supplementation (interaction: $P = 0.08$). ^{d,e}ANOVA showed a significant interaction ($P < 0.05$), which was followed by Tukey's post hoc analysis. ^dSignificantly different from corresponding basal value, $P < 0.05$. ^eSignificantly different from corresponding value before omega-3 fatty acid supplementation, $P < 0.05$. Furthermore, the before-after intervention changes in the insulin/amino acid-mediated increase in p70s6k and mTOR phosphorylation above basal values were greater in the omega-3 fatty acid group than in the corn oil group ($P < 0.05$ and $P = 0.07$, respectively; Mann-Whitney U test).

Long-term Bed-rest

*WISE 2005 (Women International Space Simulation for Exploration) ESA/CNES/NASA/CSA
Toulouse France*





Bed rest effect: $p = 0.01$; bed rest \times diet interaction: $p = 0.01$ (repeated measures ANCOVA)

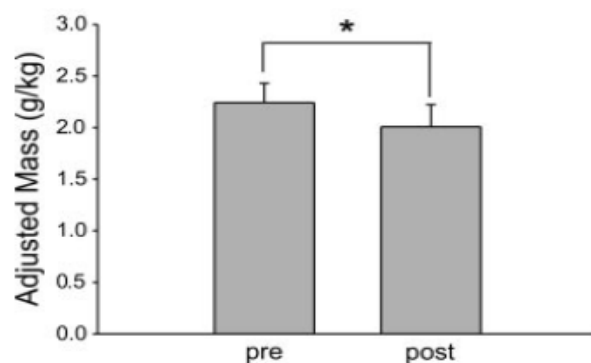
Cardiac atrophy in women following bed rest

Todd A. Dorfman,^{1,2} Benjamin D. Levine,^{1,2*} Tommy Tillery,² Ronald M. Peshock,² Jeff L. Hastings,^{1,2} Suzanne M. Schneider,² Brandon R. Macias,² Gianni Biolo,⁴ and Alan R. Hargens^{2*}

¹Institute for Exercise and Environmental Medicine, Presbyterian Hospital of Dallas, and ²Division of Cardiology, Department of Internal Medicine, University of Texas Southwestern Medical Center, Dallas, Texas; ³Division of Physical Performance and Development, University of New Mexico, Albuquerque, New Mexico; ⁴Department of Clinical, Technological and Morphological Sciences, and Division of Internal Medicine, University of Trieste, Trieste, Italy; and ⁵Department of Orthopaedic Surgery, University of California, San Diego, California

CONTROL

Adjusted LV Mass - Control



HIGH-PROTEIN

Adjusted LV Mass - Nutrition

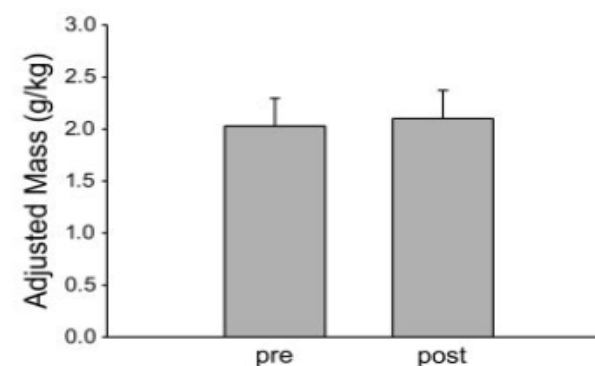
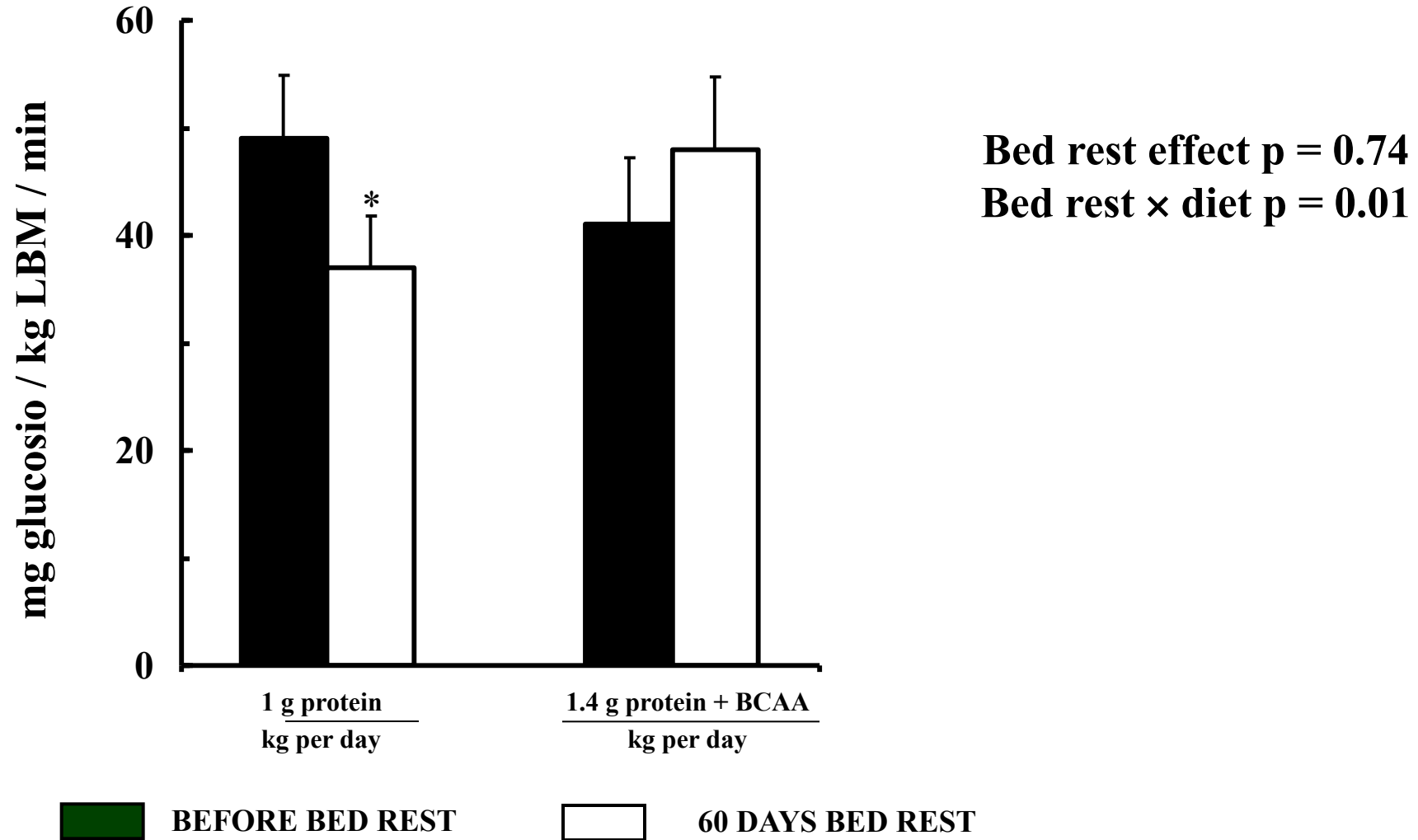


Fig. 2. A: adjusted left ventricular (LV) mass at baseline (pre) and after sedentary prolonged bed rest (post). B: adjusted LV mass at baseline and following protein supplementation during bed rest. * $P < 0.05$.



Rate of glucose disappearance during euglycemic hyperinsulinemic clamp

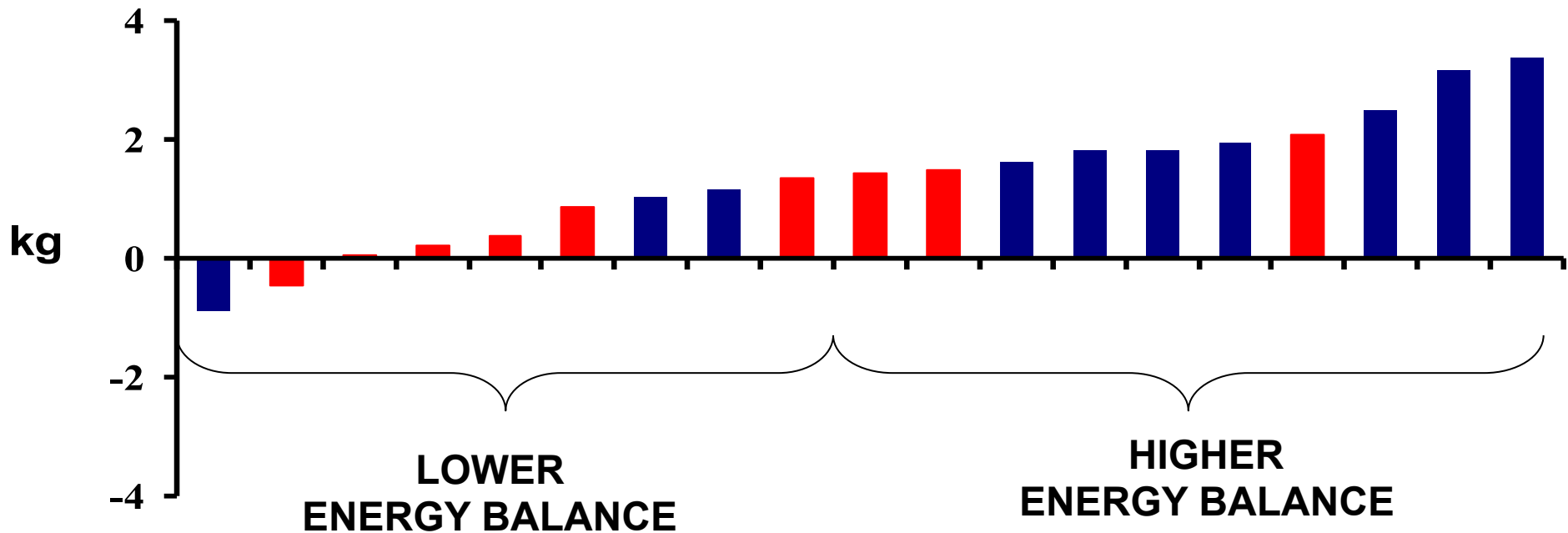


**PROTEIN KINETICS IN
RELATION TO ENERGY
AVAILABILITY**

VALDOLTRA Bed Rest Studies 2006 & 2007

INDIVIDUAL CHANGES IN FAT MASS

-  Study A (spontaneous adaptation to decreased energy requirements)
-  Study B (activity-matched eucaloric diet)



Positive energy balance is associated with accelerated muscle atrophy and increased erythrocyte glutathione turnover during 5 wk of bed rest¹⁻³

Gianni Biolo, Francesco Agostini, Bostjan Simunic, Mariella Sturma, Lucio Torelli, Jean Charles Preiser, Ginette Deby-Dupont, Paolo Magni, Felice Strollo, Pietro di Prampero, Gianfranco Guarneri, Igor B Mekjavic, Rado Pišot, and Marco V Narici

Am J Clin Nutr 2008;88:950–8.



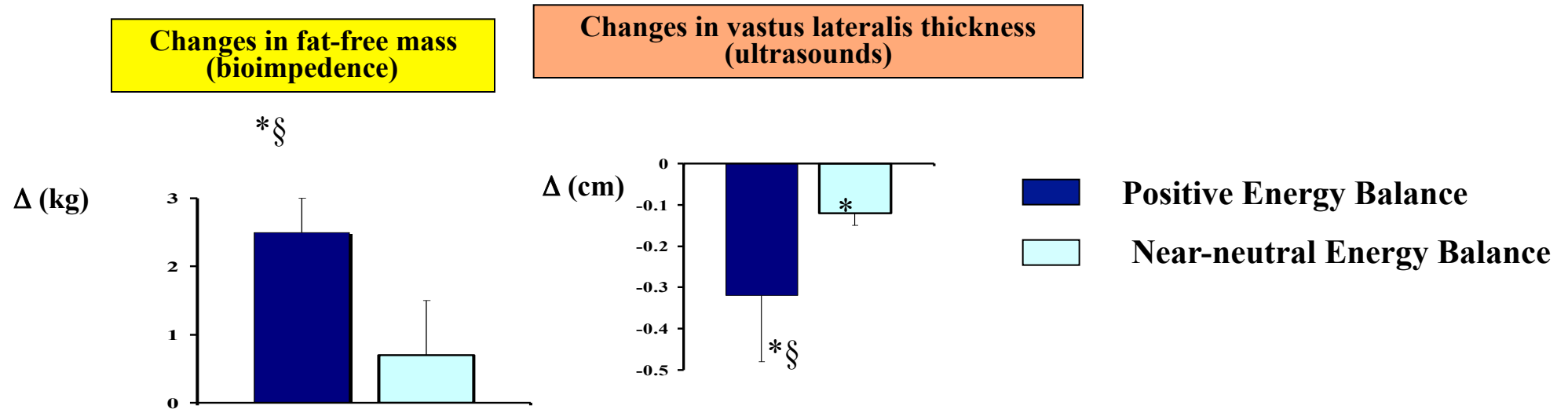
*, $p < 0.05$ significant different from zero;

§, $p < 0.05$ versus lower energy balance

Positive energy balance is associated with accelerated muscle atrophy and increased erythrocyte glutathione turnover during 5 wk of bed rest¹⁻³

Gianni Biolo, Francesco Agostini, Bostjan Simunic, Mariella Sturma, Lucio Torelli, Jean Charles Preiser, Ginette Deby-Dupont, Paolo Magni, Felice Strollo, Pietro di Prampero, Gianfranco Guarneri, Igor B Mekjavic, Rado Pišot, and Marco V Narici

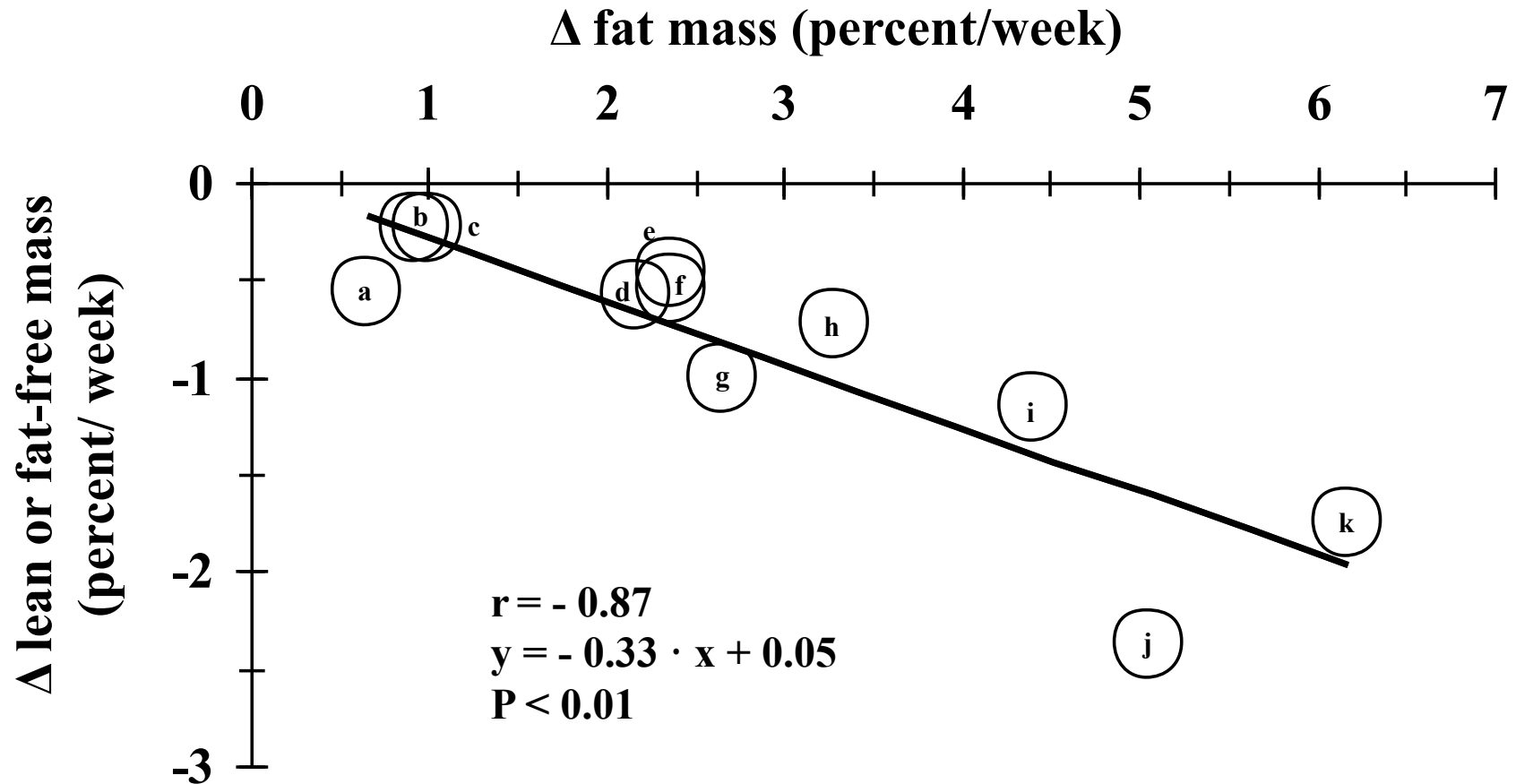
Am J Clin Nutr 2008;88:950–8.



*, $p < 0.05$ significant different from zero;
§, $p < 0.05$ versus lower energy balance

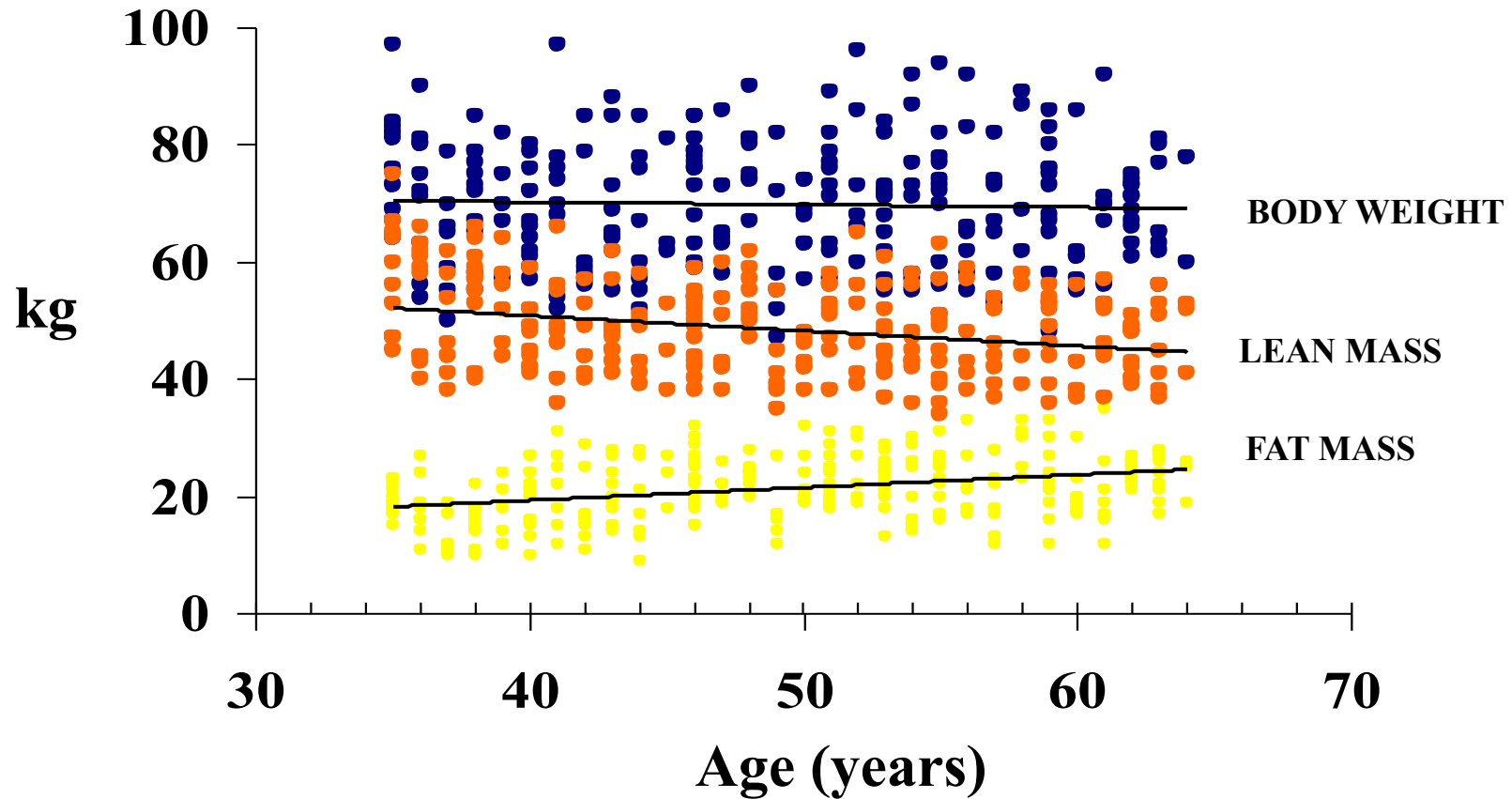
RELATIONSHIP BETWEEN CHANGES IN FAT AND LEAN MASS IN BED REST STUDIES (1-17 weeks) AT POSITIVE ENERGY BALANCE

(a) Lovejoy et al., Am J Physiol 1999; (b) Shackelford et al., J Appl Physiol 2004 ; (c) Scheld et al., Clin Chem 2001; (d) NNEB; (e) Krebs et al., Aviat Space Environ Med 1990 ; (f) Gretebeck et al., J Appl Physiol 1995 ; (g) Stein et al., Am J Physiol 1999; (h) Ferrando et al., Am J Physiol 1996; (i) PEB ; (j) Barbe et al., J Appl Physiol 1999 ; (k) Blanc et al., Am J Physiol Regul Integr Comp Physiol 2000.



CROSS-SECTIONAL STUDY
252 healthy subjects with normal body mass index, 35 to 65 years

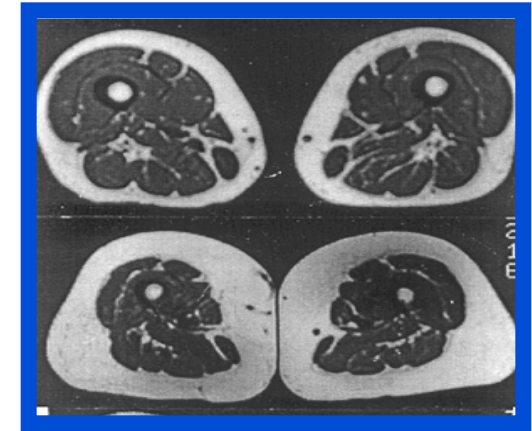
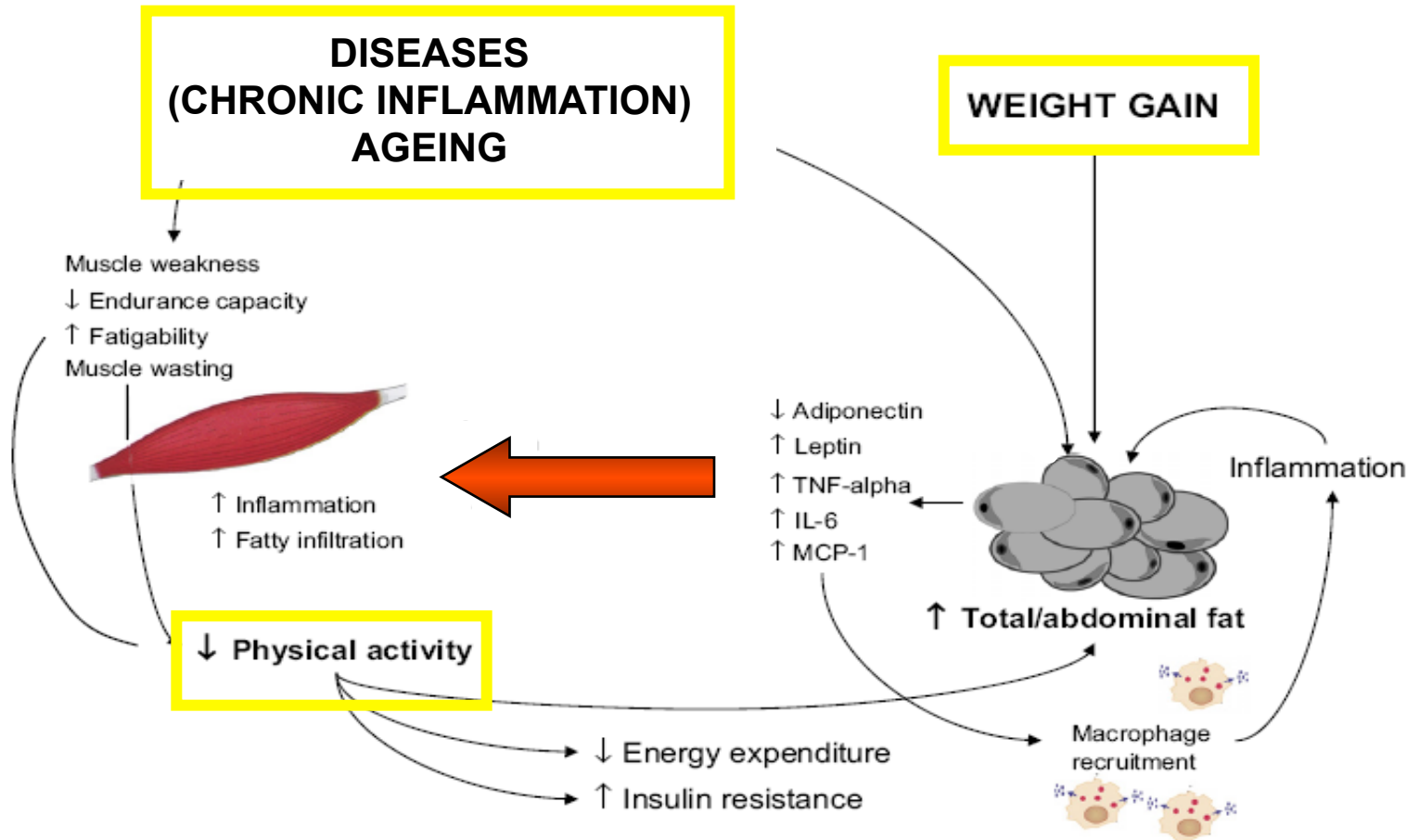
BODY WEIGHT AND COMPOSITION



Inter-relationships between adipose tissue and muscle

A mechanism leading to sarcopenic obesity

(Ageing, Critical illness, Chronic inflammatory diseases, Cancer)

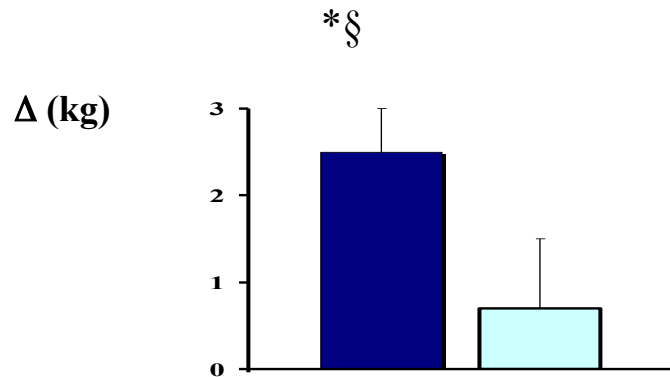


Positive energy balance is associated with accelerated muscle atrophy and increased erythrocyte glutathione turnover during 5 wk of bed rest¹⁻³

Gianni Biolo, Francesco Agostini, Bostjan Simunic, Mariella Sturma, Lucio Torelli, Jean Charles Preiser, Ginette Deby-Dupont, Paolo Magni, Felice Strollo, Pietro di Prampero, Gianfranco Guarneri, Igor B Mekjavic, Rado Pišot, and Marco V Narici

Am J Clin Nutr 2008;88:950–8.

Changes in fat-free mass (bioimpedence)

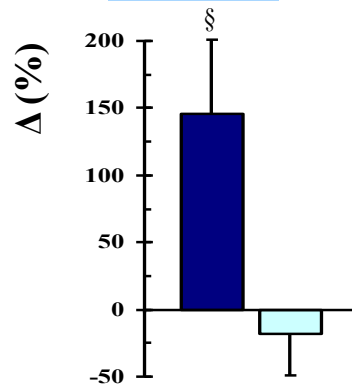


Changes in vastus lateralis thickness (ultrasounds)

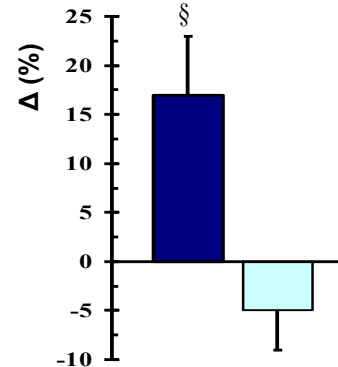


Positive Energy Balance
Near-neutral Energy Balance

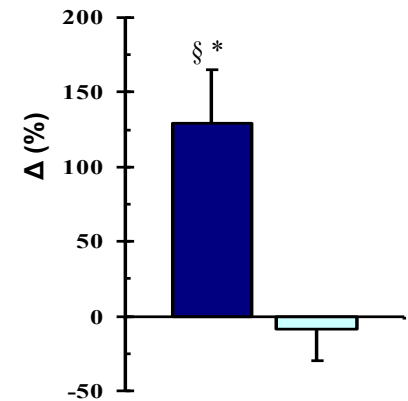
PLASMA CRP



PLASMA MYELOPEROXIDASE



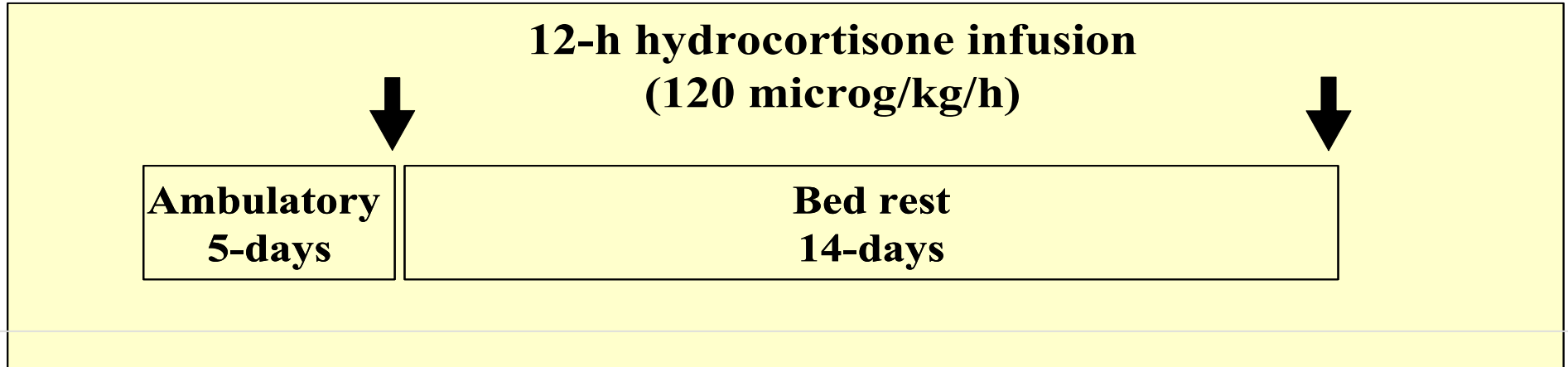
ERYTHROCYTE GLUTATHIONE SYNTHESIS RATE



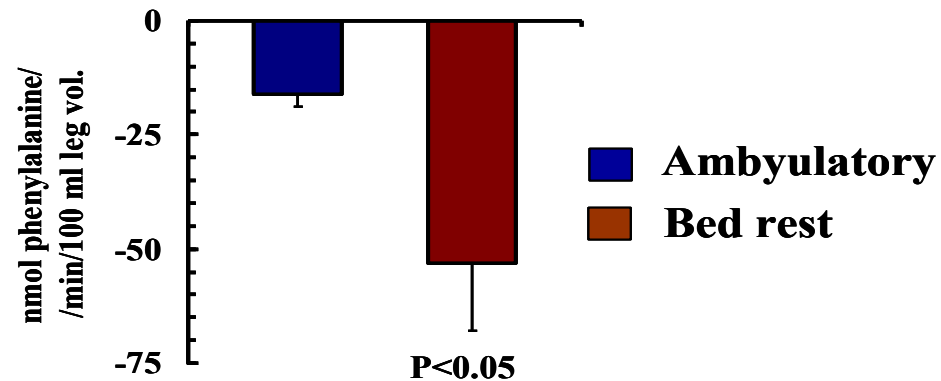
*****, $p < 0.05$ significant different from zero;
§, $p < 0.05$ versus lower energy balance

Inactivity Amplifies the Catabolic Response of Skeletal Muscle to Cortisol

Ferrando et al., J Clin Endocrinol & Metab, 1999



MUSCLE PROTEIN BALANCE

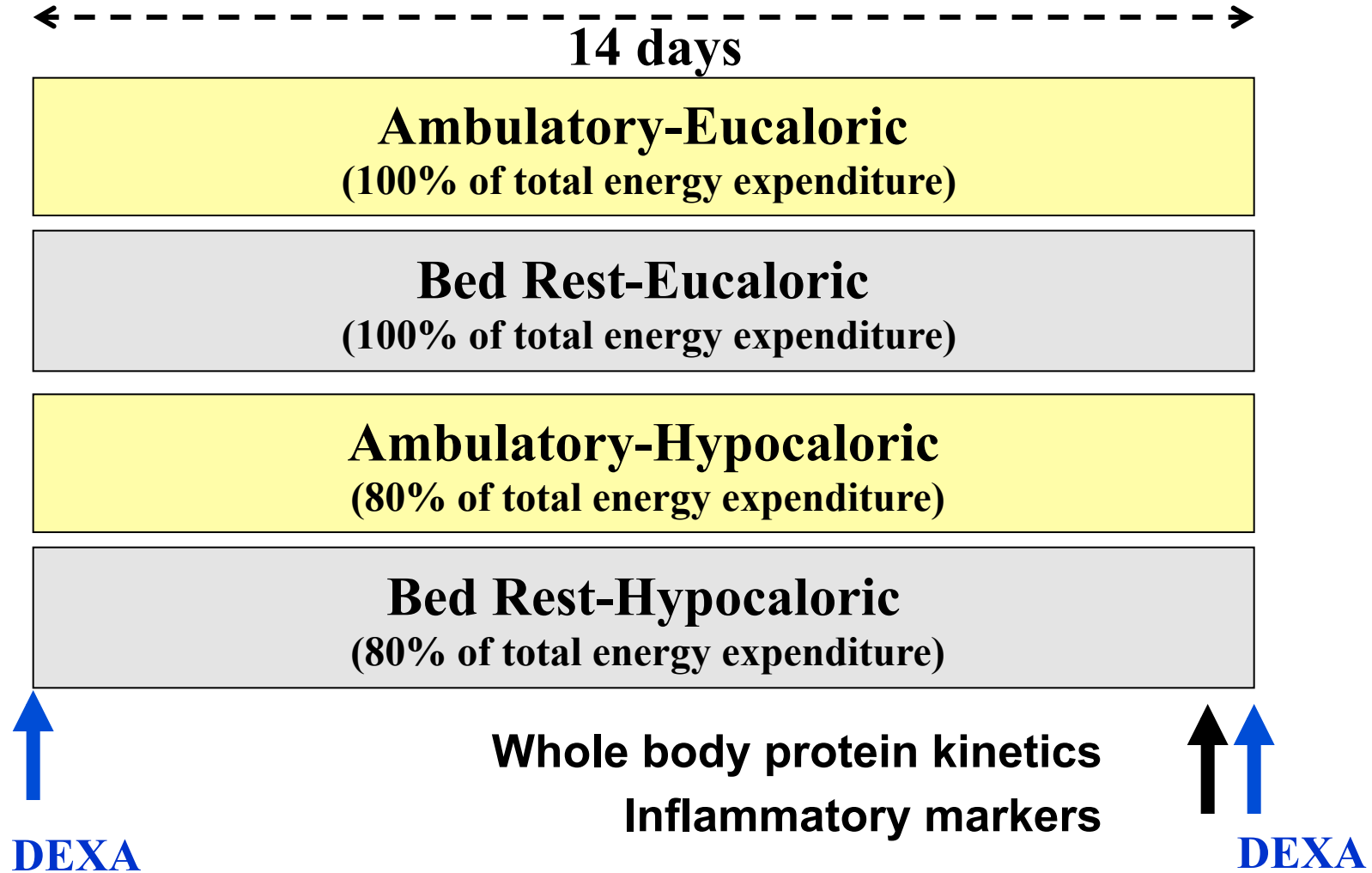


EXPERIMENTAL PROTOCOL

STBR study DLR – Cologne – Germany

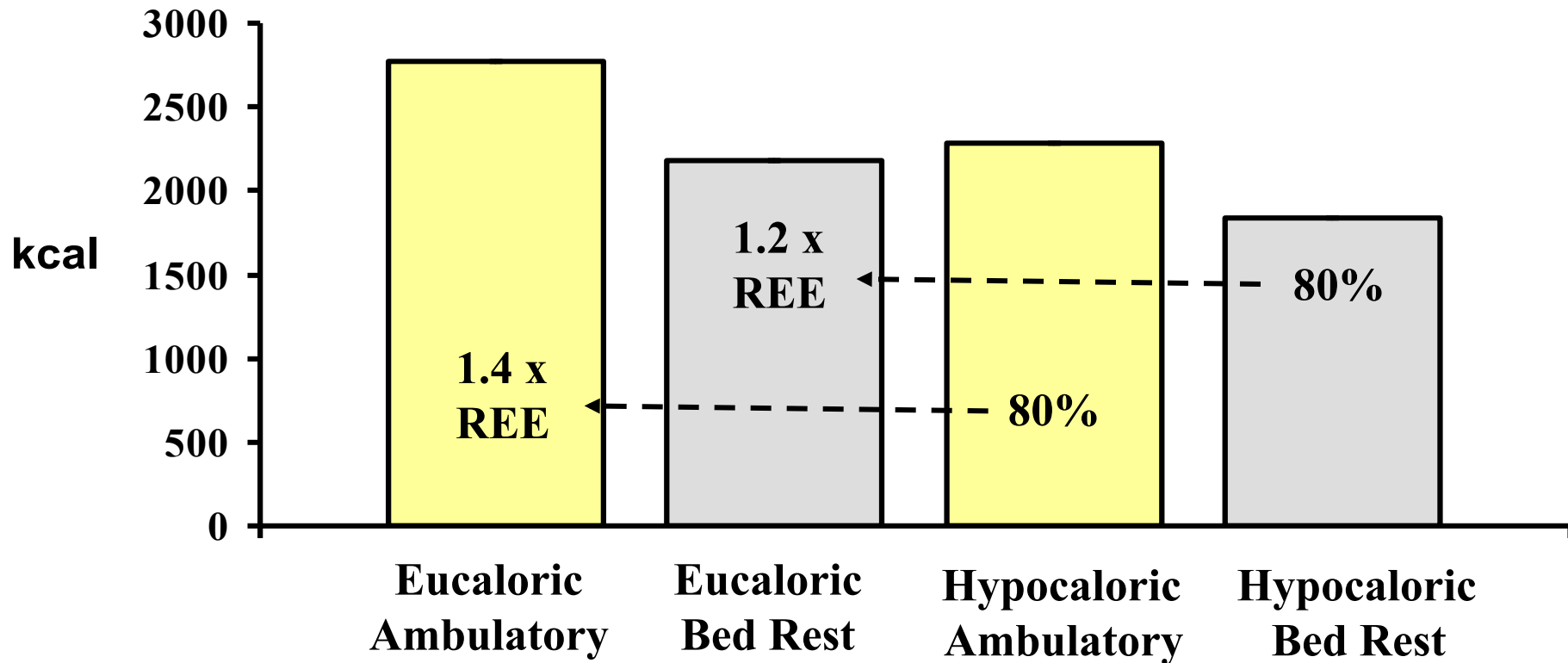
9 NORMAL MALE VOLUNTEERS

Randomized Cross-Over

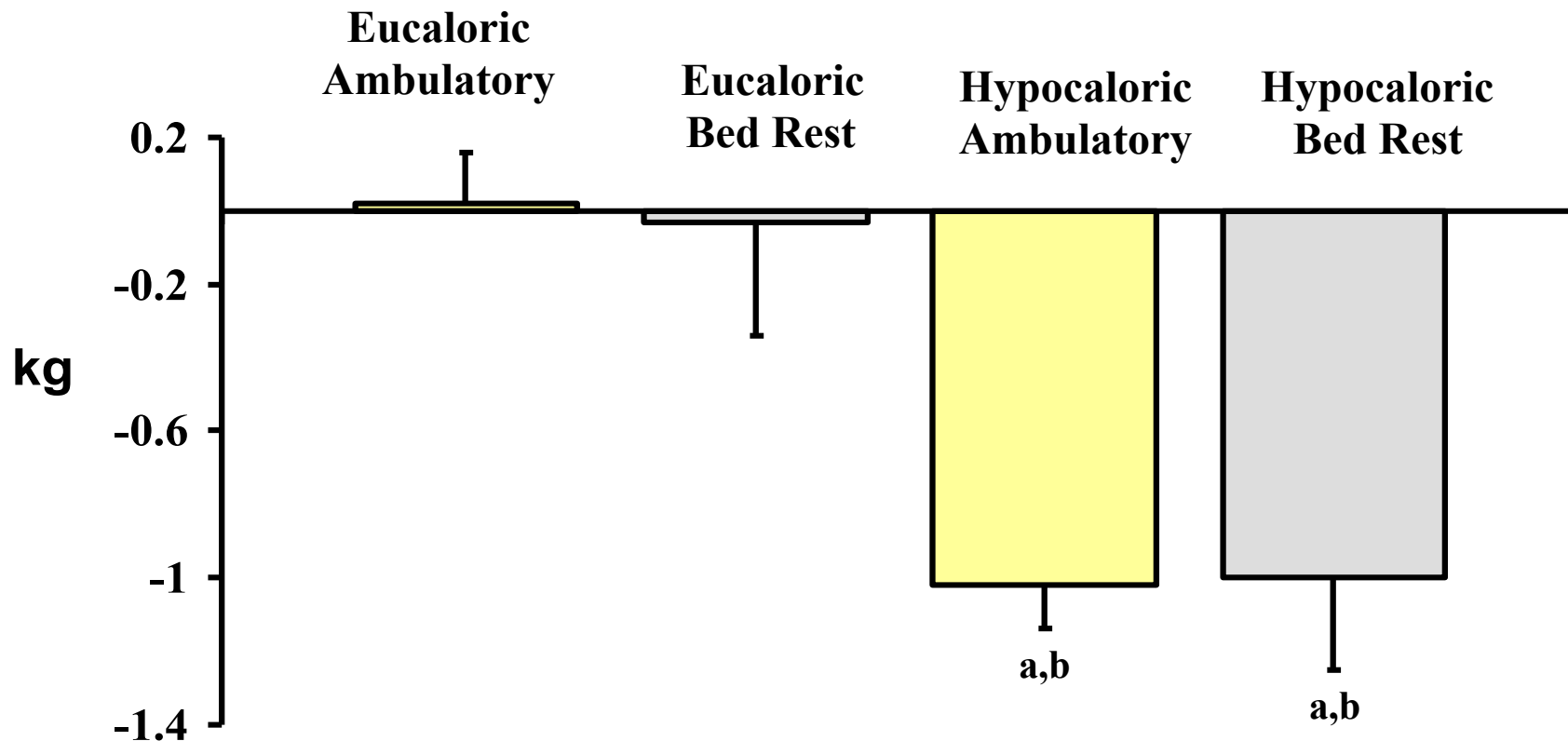


J Physiol 2005; Am J Clin Nutr 2007; J Clin Endocrinol Metab 2008

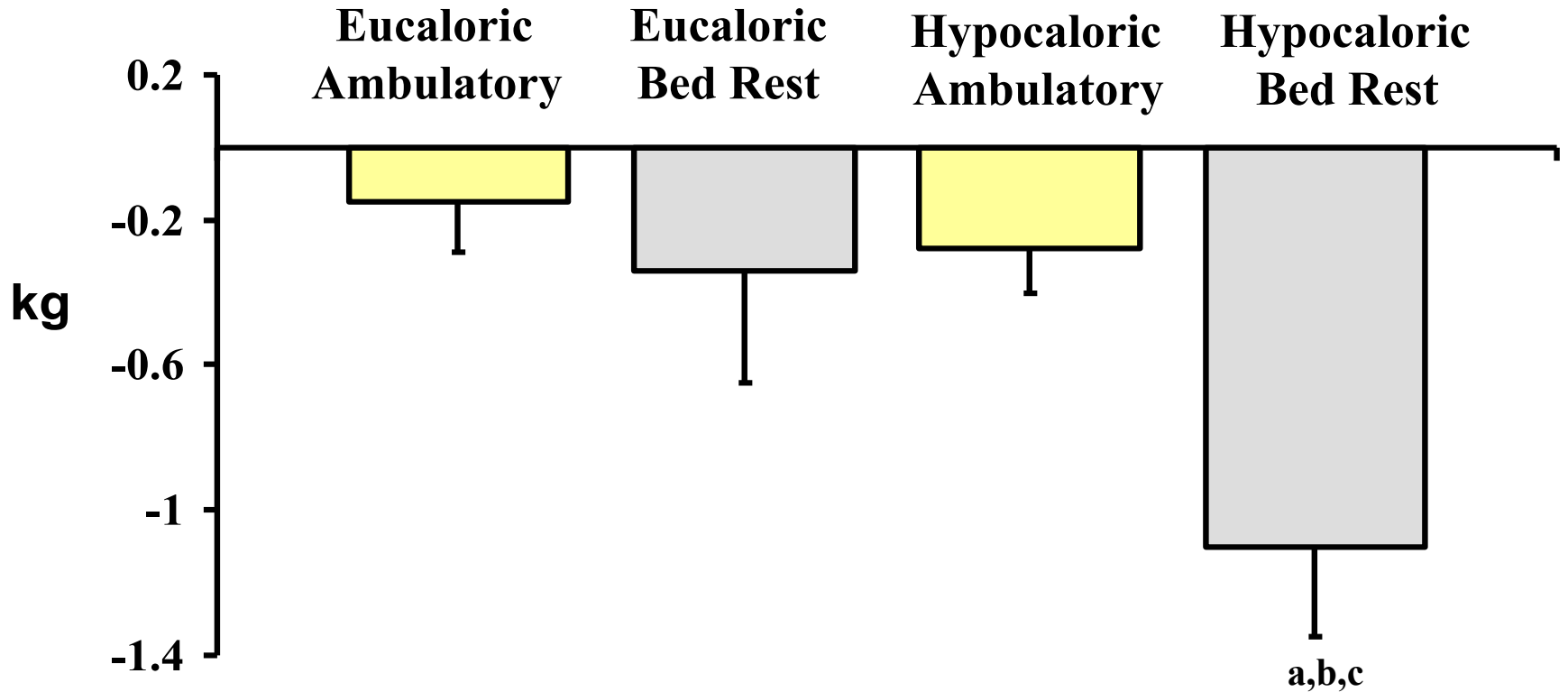
ENERGY INTAKE



CHANGES IN FAT MASS (DXA) DURING THE 14-DAY EXPERIMENTAL PERIODS (ENERGY BALANCE)



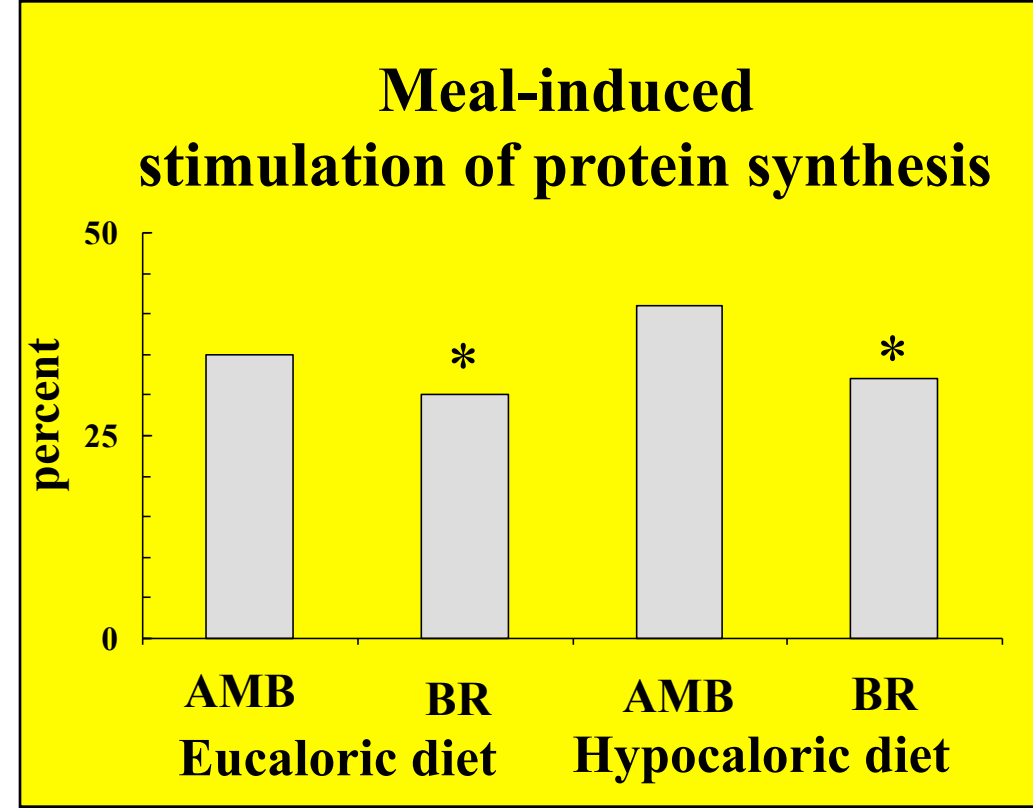
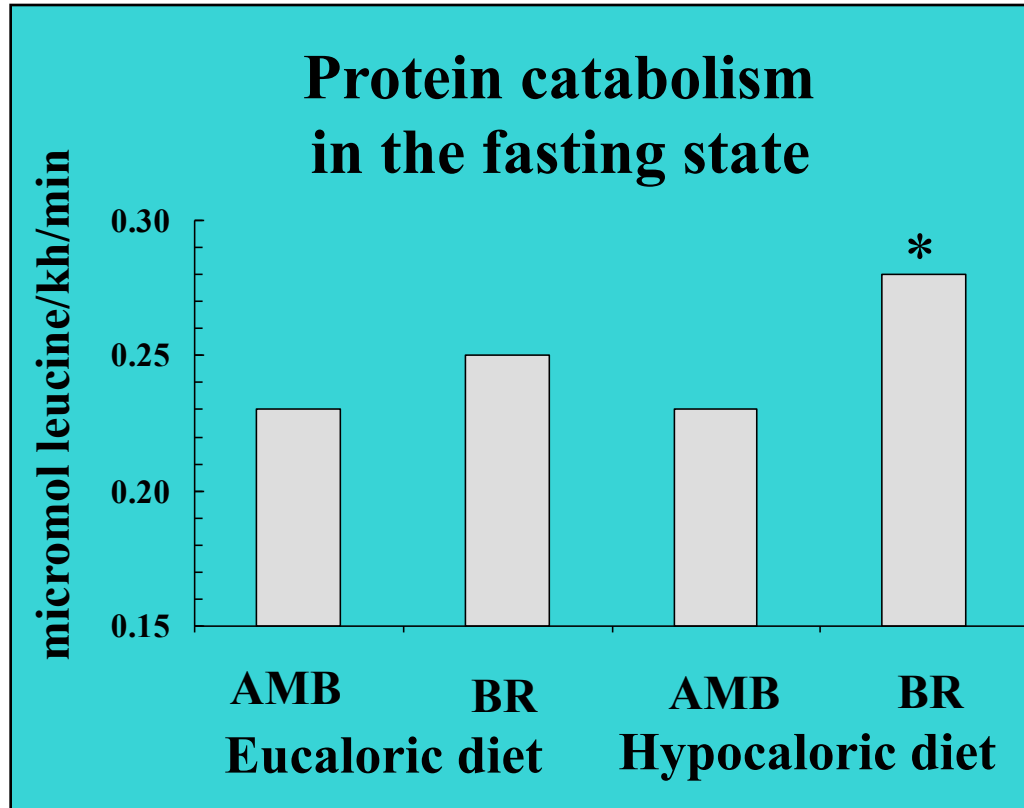
CHANGES IN LEAN MASS (DXA) DURING THE 14-DAY EXPERIMENTAL PERIODS

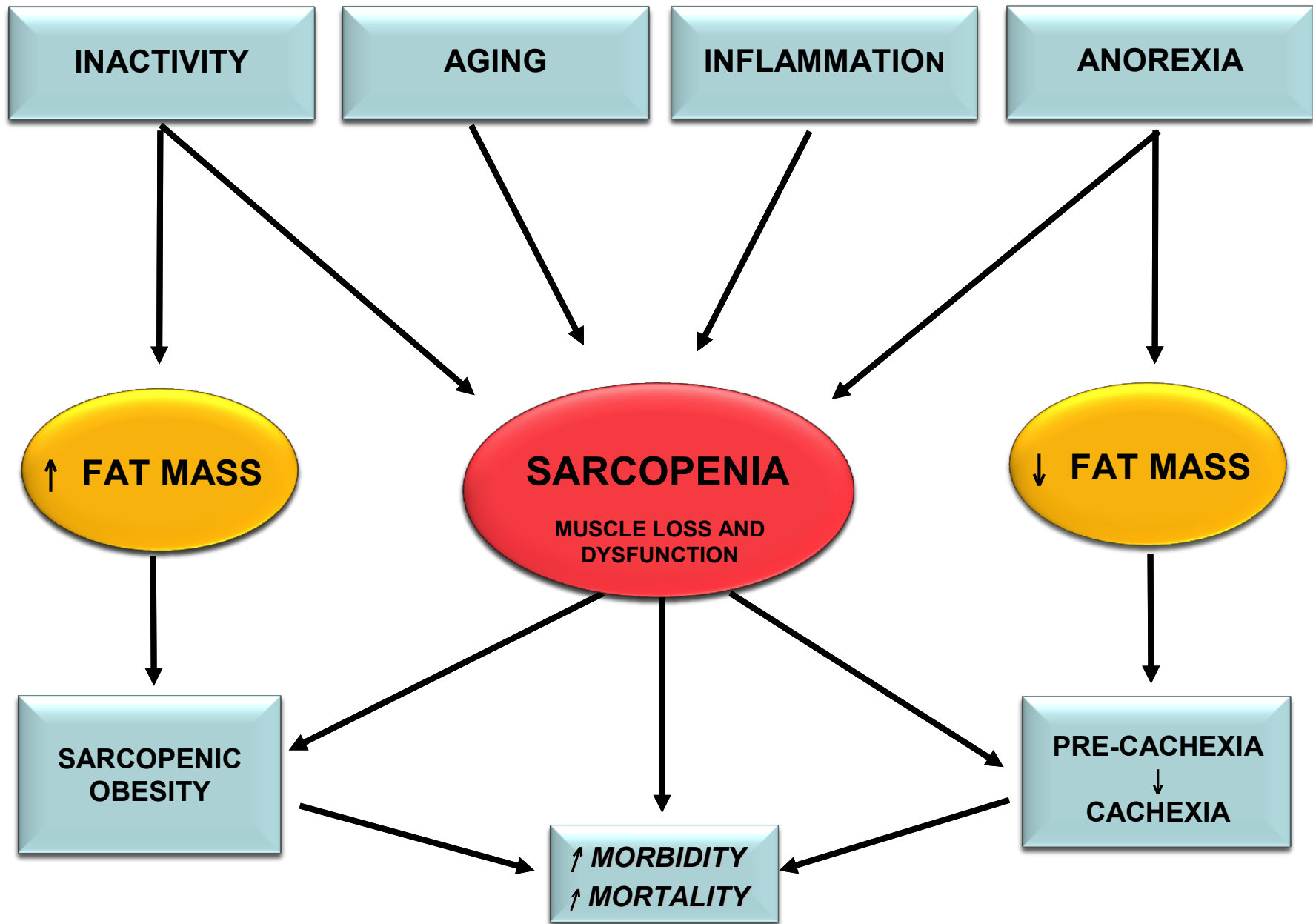


Calorie restriction accelerates the catabolism of lean body mass during 2 wk of bed rest¹⁻³

Gianni Biolo, Beniamino Ciocchi, Manuela Stulle, Alessandra Bosutti, Rocco Barazzoni, Michela Zanetti, Raffaella Antonione, Marion Lebenstedt, Petra Platen, Martina Heer, and Gianfranco Guarneri

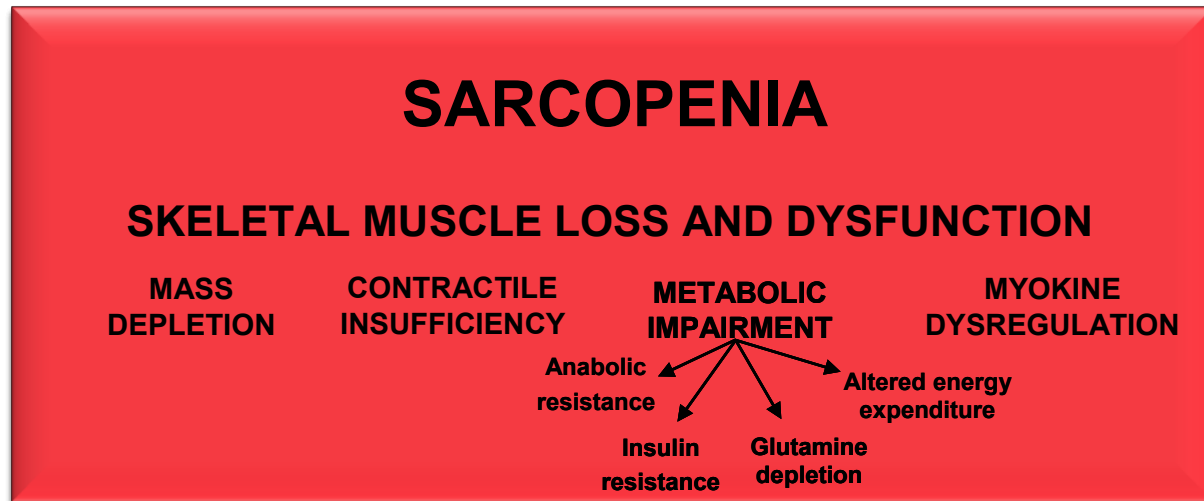
Am J Clin Nutr 2007;86:366-72.





CHRONIC DISEASES

AGING



↑ MORBIDITY

Dynapenia, fatigue, disability and falls, impaired ventilation, osteoporosis, bone fractures, dyslipidemia, metabolic syndrome, type 2 diabetes, increased cardiovascular risk, impaired immunity, infections, etc.



↑ MORTALITY

SUMMARY

- 1. Inactivity impairs amino acid-induced protein anabolism.**
- 2. Physical activity enhances anabolic utilization of amino acids.**
- 3. High protein-BCAA intake decreases inactivity-mediated loss of lean body and myocardial mass as well as prevents inactivity-mediated insulin resistance.**
- 4. Overfeeding and underfeeding accelerate inactivity-mediated muscle atrophy.**