

## Anabole resistentie bij veroudering

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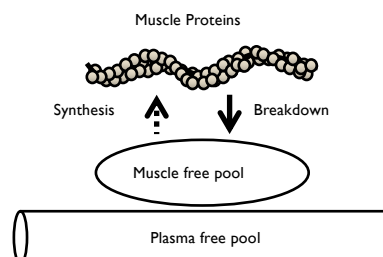
## Universiteit van Nederland



## University of the Netherlands



## Muscle protein turnover



Burd et al., Exerc Sport Sci Rev 2013

### Fractional muscle protein synthesis

1-2 % per day

(0.04 – 0.14 %·h<sup>-1</sup>)



### Muscle reconditioning



Lance Armstrong



Jay Cutler

### Muscle deconditioning



- immobilisation
- sarcopenia
- cancer cachexia
- COPD
- type 2 diabetes
- cardiovascular disease

### Muscle mass maintenance

### Main anabolic stimuli

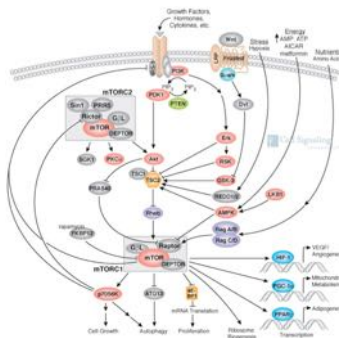
### Nutrition is an anabolic stimulus



### Amino acids



### Amino acids stimulate protein synthesis



### Post-prandial muscle protein synthesis

- protein digestion
- amino acid absorption
- plasma amino acid availability
- hormonal response
- postprandial perfusion
- muscle protein signaling proteins
- myofibrillar protein synthesis



### Research methods



### Post-prandial muscle protein synthesis

- protein digestion
- amino acid absorption
- plasma amino acid availability
- hormonal response
- postprandial perfusion
- muscle protein signaling proteins
- myofibrillar protein synthesis



### Stable isotope tracers



### Intrinsically labeled protein



van Loon et al., *J Dairy Sci*, 2009; Penning et al., *J Dairy Sci*, 2010; Burd et al., *PLOS One*, 2013

### Post-prandial protein synthesis



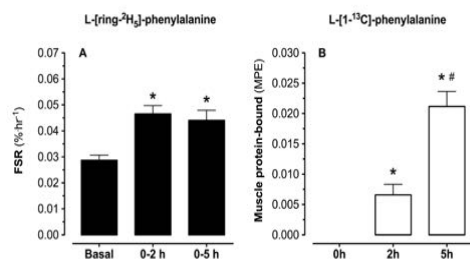
Groen et al., *PLOS one*, 2015

### Post-prandial protein synthesis



Groen et al., *PLOS one*, 2015

### Post-prandial protein synthesis



Groen et al., *PLOS one*, 2015

'You are what you just ate'

### Post-prandial protein synthesis



- source of protein



- amount of protein



- macronutrients



- timing



- food preparation

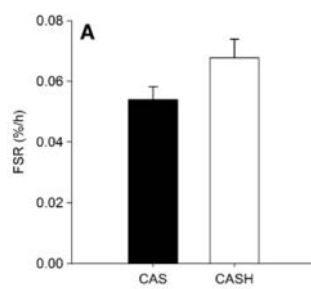


- body position

### Source of protein

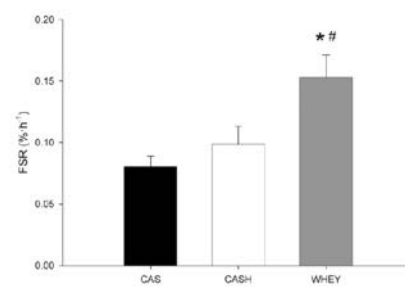


### Intact protein versus protein hydrolysate



Koopman et al., Am J Clin Nutr, 2009

### Whey versus casein

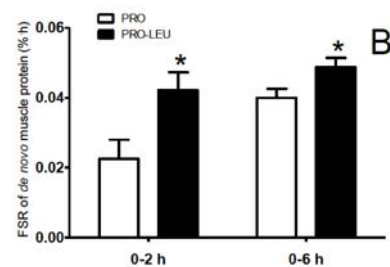


Pennington et al., Am J Clin Nutr, 2011

### Whey versus casein



### Leucine co-ingestion



Wall et al., Clin Nutrition, 2013

## Milk versus Beef



Burd et al., *Am J Clin Nutr*, 2015

## Plant-derived proteins: game changers?



## Plant based proteins

### The Skeletal Muscle Anabolic Response to Plant- versus Animal-Based Protein Consumption<sup>1</sup>

Stephan van Vliet,<sup>2,3</sup> Nicholas A Burd,<sup>2,3</sup> and Luc JC van Loon<sup>2\*</sup>

<sup>2</sup>Department of Kinesiology and Community Health, University of Illinois at Urbana-Champaign, Urbana, IL and <sup>3</sup>Department of Human Movement Sciences, Faculty of Health, Medicine, and Life Sciences, School for Nutrition and Translational Research in Metabolism (NUTRIM), Maastricht University, Maastricht, Netherlands

van Vliet et al., *J Nutr*, 2015

## Plant based proteins



van Vliet et al., *J Nutr*, 2015

## Muscle gain on plant based protein consumption



## Muscle gain on plant based protein consumption



Bill Pearl - 1971



## Soy



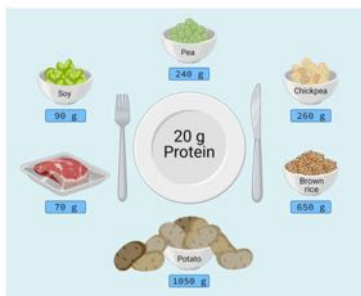
Tang et al., J Appl Physiol, 2009  
Yang et al., Nutr Metab, 2012

## Wheat



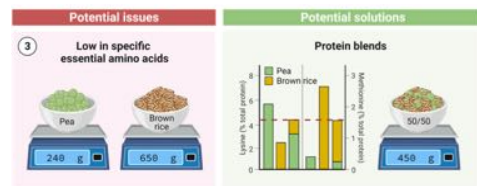
Gorissen et al., J Nutr, 2016

## Plant-based whole-foods



Pinckaers et al., Sports Med, 2021

## Anabolic properties of plant-based proteins

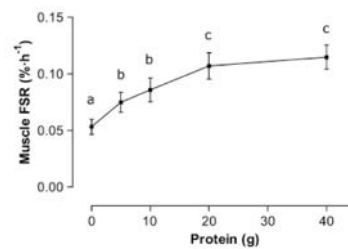


Pinckaers et al., Sports Med, 2021

## Amount of dietary protein



## Amount of dietary protein

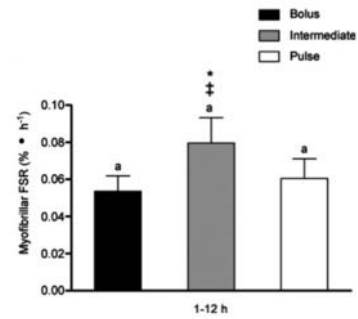


Moore et al., J Physiol, 2009

### Timing of protein ingestion

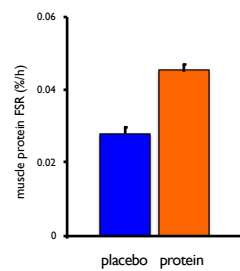


### Protein intake distribution



Areza et al., J Physiol, 2013.

### Muscle protein synthesis during sleep

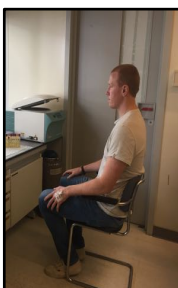


Groen et al., Am J Physiol, 2012.

### Food preparation



### Body position

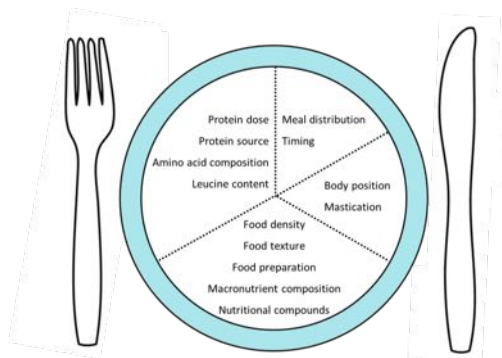


### Whole foods and food processing





## Post-prandial protein synthesis



Kouw et al., PhD thesis, 2018.

## Muscle contraction is an anabolic stimulus



## Muscle contraction

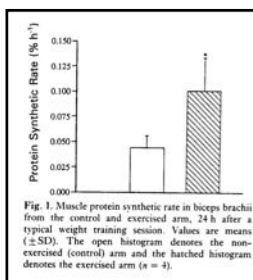
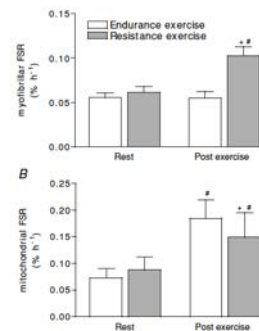


Fig. 1. Muscle protein synthetic rate in biceps brachii from the control and exercised arm, 24 h after a typical weight training session. Values are means ( $\pm$ SD). The open histogram denotes the non-exercised (control) arm and the hatched histogram denotes the exercised arm ( $n = 4$ ).

Chesley et al., J Appl Physiol, 1992.

## Post-exercise muscle protein synthesis



Wilkinson et al., J Physiol, 2008.

## Muscle reconditioning



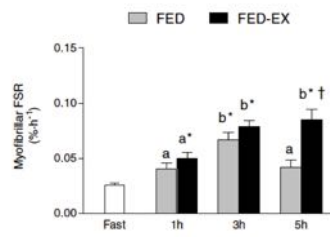
Lance Armstrong



Jay Cutler

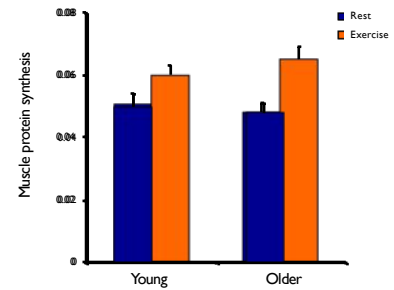
## Synergy between exercise and food intake

### Exercise and nutrition



Moore et al., J Physiol, 2009

### Post-prandial muscle protein synthesis



Pennington et al., Am J Clin Nutr, 2010

Physical activity prior to food intake

'You are more of what you just ate'

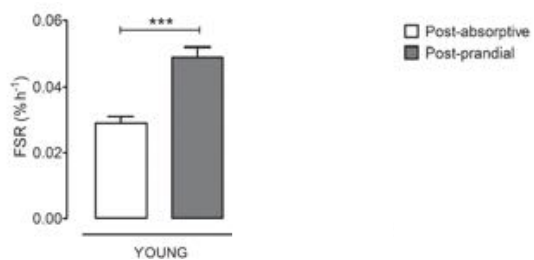
### Muscle deconditioning



- sarcopenia
- cancer cachexia
- COPD
- type 2 diabetes
- cardiovascular disease

What causes muscle loss with aging

### Anabolic resistance



Wall et al., PLOS one, 2015

### Concept of anabolic resistance

Burd et al., J Appl Physiol, 2012

#### Physiological factors



Protein digestion



Amino acid absorption



Postprandial hormonal response



Microvascular perfusion



Amino acid uptake in muscle



Intramuscular signalling



Muscle protein synthesis

Anabolic  
resistance of muscle  
protein synthesis

Lower level of physical activity



Changes in food intake

Bedrest



Hospitalization

Insulin resistance



Obesity



Aging

Inflammation

Health status

Medication

#### Clinical factors

### Decline in physical activity

Burd et al., J Appl Physiol, 2012

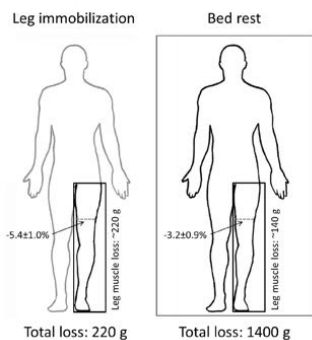
### Muscle disuse



Wall et al., Acta Physiol Scand, 2013

### Muscle deconditioning





Dirks et al. J Appl Physiol 2017

Marlou L. Dirks,<sup>1</sup> Benjamin T. Wall,<sup>1</sup> Bas van de Valk,<sup>1</sup> Tanya M. Holloway,<sup>2</sup>  
Graham P. Holloway,<sup>2</sup> Adrian Chabowski,<sup>3</sup> Gijs H. Goossens,<sup>1</sup> and Luc J.C. van Loon<sup>1</sup>

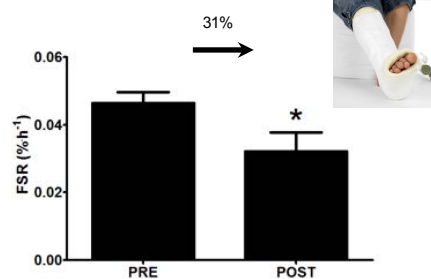
### One Week of Bed Rest Leads to Substantial Muscle Atrophy and Induces Whole-Body Insulin Resistance in the Absence of Skeletal Muscle Lipid Accumulation

Diabetes 2016;85:2862–2875 | DOI: 10.2337/db15-1861



Dirks et al. Diabetes, 2016

### Anabolic resistance to protein ingestion



Wall et al. J Clin Endocrinol Metab, 2013

Decline in physical activity

'You are less of what you just ate'

## Clinical relevance

## Exercise training in the elderly

Muscle mass and strength

Endurance capacity

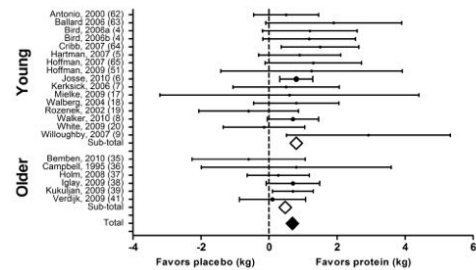
Functional capacity



Franssen et al., 1988, 1990, 2003; Fitts et al., 1999, 1994; Chouinard et al., 1991; Lexell et al., 1992; Allen et al., 1994; Vasselj et al., 2002; Baskaran et al., 2002; Bouter et al., 2002; Toot et al., 2002; Bui et al., 2002; Steiner et al., 2002, 2004; Young et al., 2009

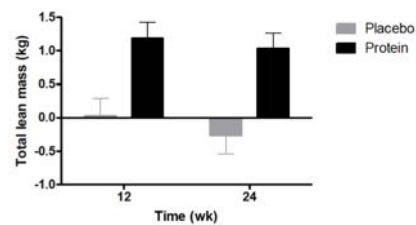


## Exercise training and protein supplementation



Cermak et al., Am J Clin Nutr, 2012.

## Protein supplementation in frail elderly



Tieland et al., JAMDA, 2012-8



Dirks et al. Diabetes, 2016

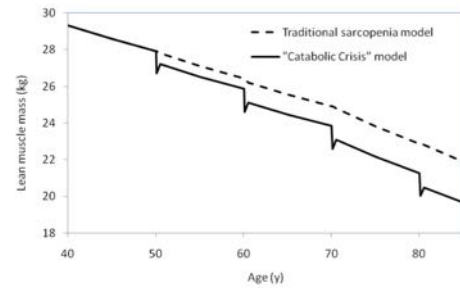
### Hospital admission



Most older patients are typically hospitalized for 5-7 days

Fisher et al., Arch Intern Med, 2010

### Catabolic crisis model



English et al., Curr Opin Clin Nutr Metab Care, 2010

### Hospital admission

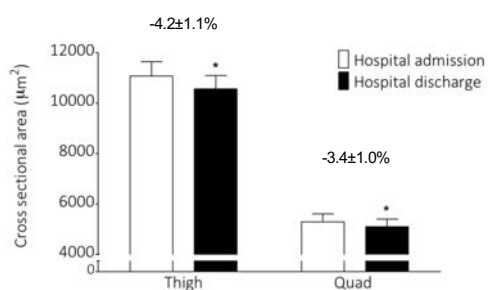


Short periods of bedrest following disease or injury contribute substantially to the loss of muscle mass with aging

Wall et al., Aging Res. Rev., 2013



### Muscle loss during hospitalisation



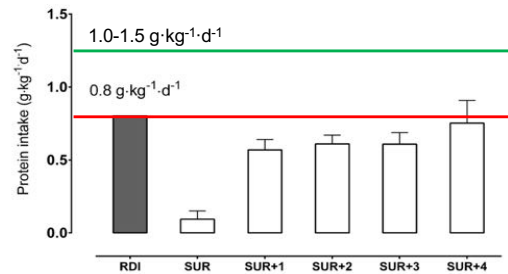
Kouw et al., JAMDA, 2018

Attenuate muscle mass and/or strength loss





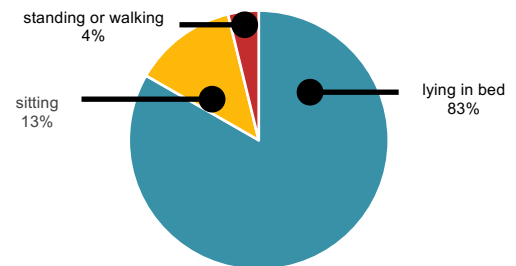
### Protein intake during hospitalisation



Weijzen et al., J Nutr Health Aging, 2019



### Physical activity during hospitalisation



Brown et al., J Am Ger Soc, 2009



### Conclusions

Protein ingestion and muscle contraction stimulate muscle tissue protein synthesis

Physical (in)activity (de)sensitizes skeletal muscle tissue to the anabolic properties of dietary protein ingestion

Protein is required to support muscle conditioning in both health and disease

## Attenuate muscle loss during disuse

Remain physically active as much as possible

Apply exercise mimetics when appropriate

Consume a more protein dense diet

Commit to an active rehabilitation program

## Collaborators and sponsors

