

Apports hyperprotéiques chez le sujet âgé: bénéfiques et risques

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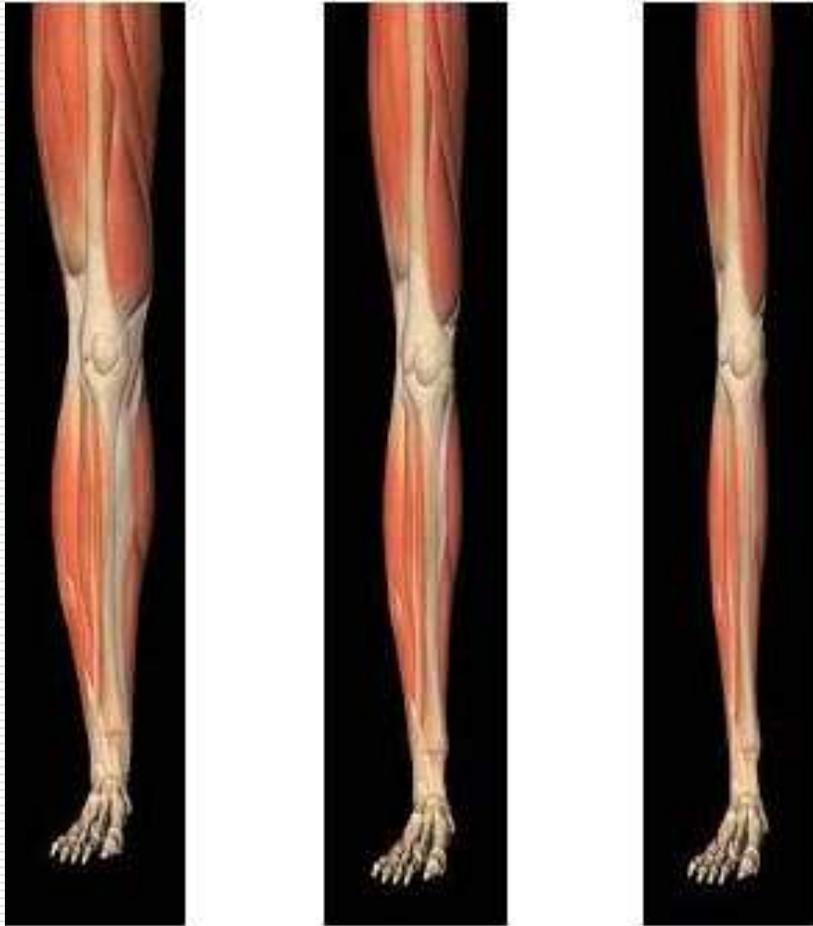


Centre universitaire de santé McGill
McGill University Health Centre

Divulgation

- Membre du Comité aviseur Pfizer
 - Reçu des honoraires de Pfizer pour des présentations
 - Membre du Comité d'experts sur l'implémentation du Plan Alzheimer Québec du MSSS
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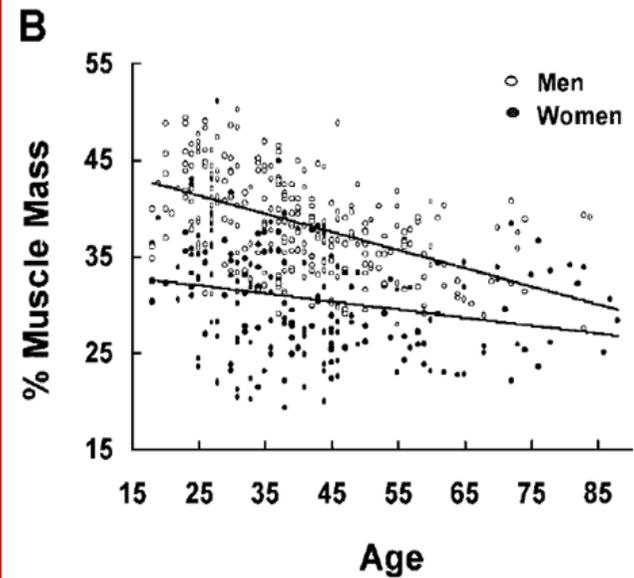
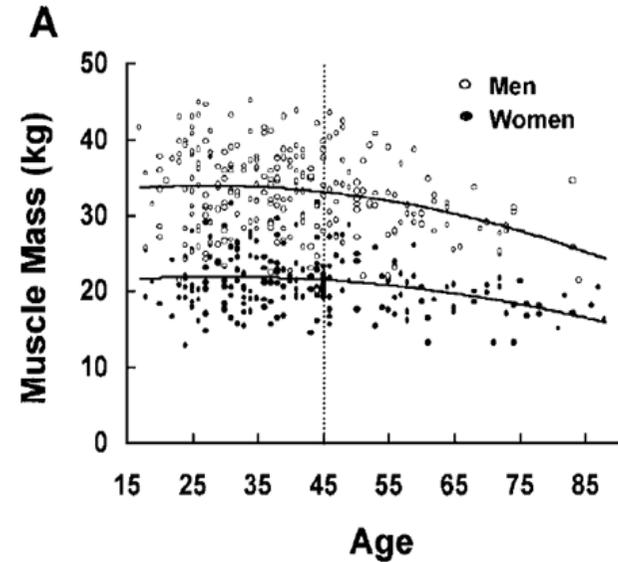
Diminution de la masse musculaire avec l'âge



20 y.o

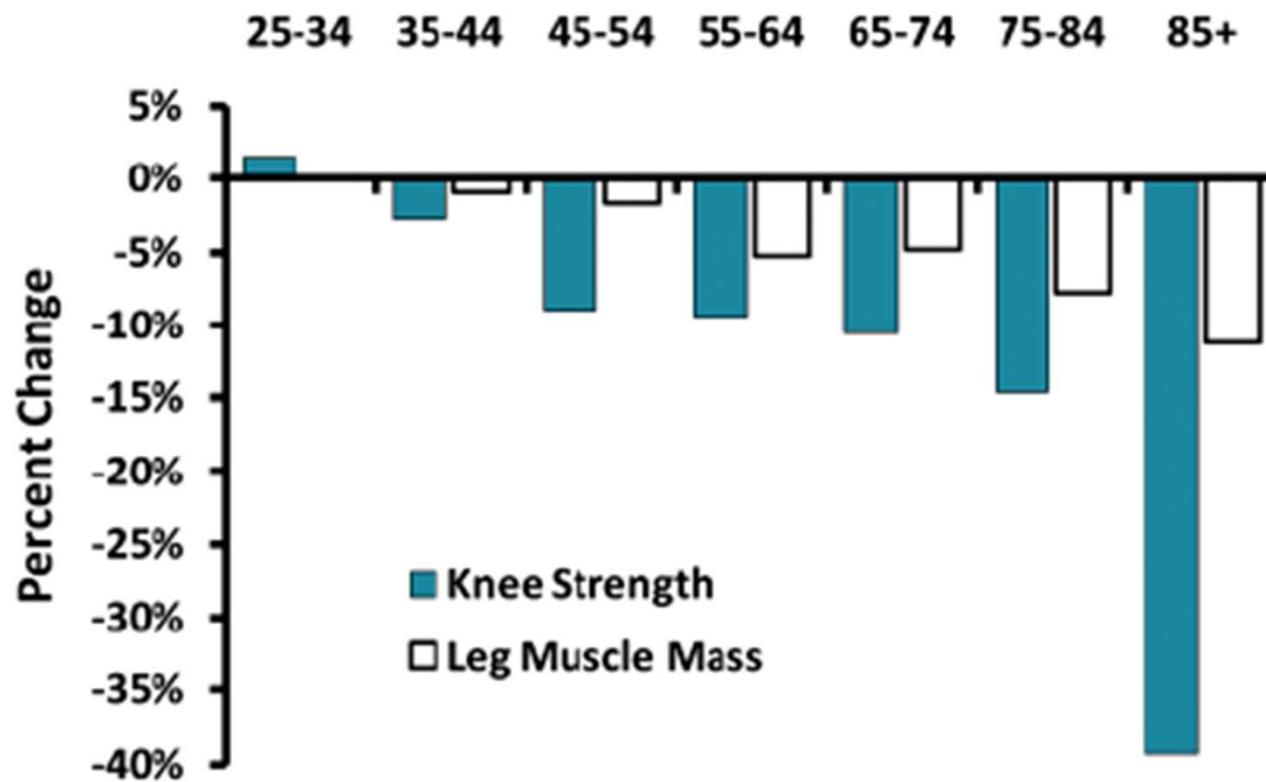
70 y.o

85 y.o

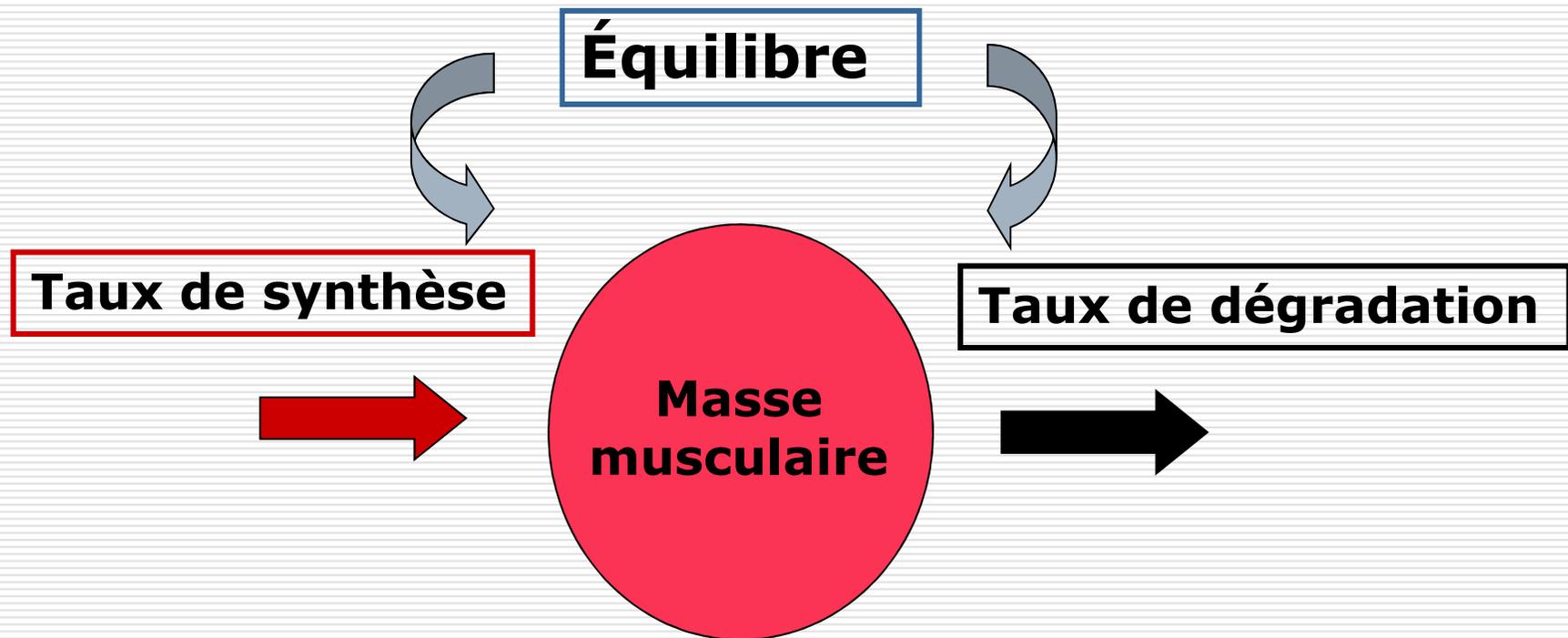


Janssen et al *J Appl Physiol* 88, 2000

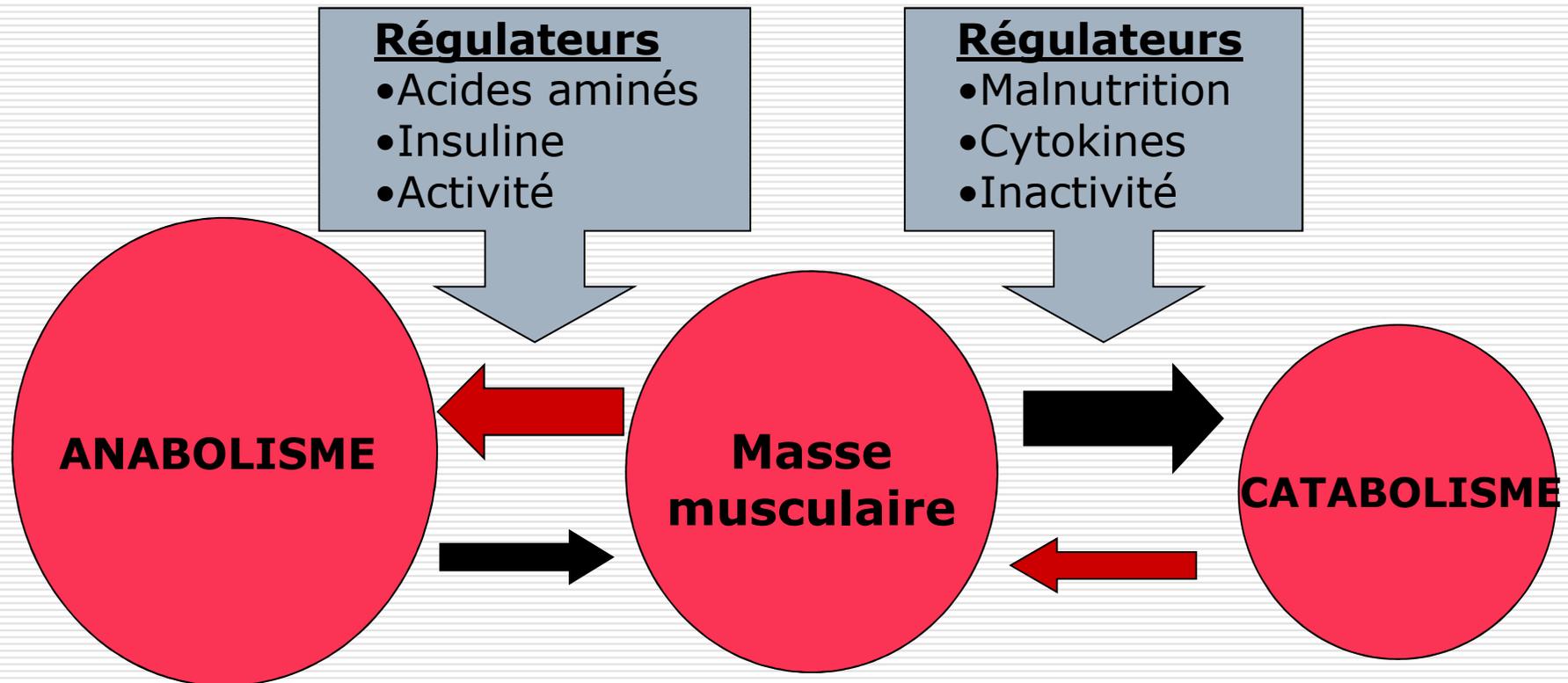
Perte musculaire versus perte de la force



Maintien de la masse musculaire



Contrôle de la masse musculaire



Taux de renouvellement protéique corporel

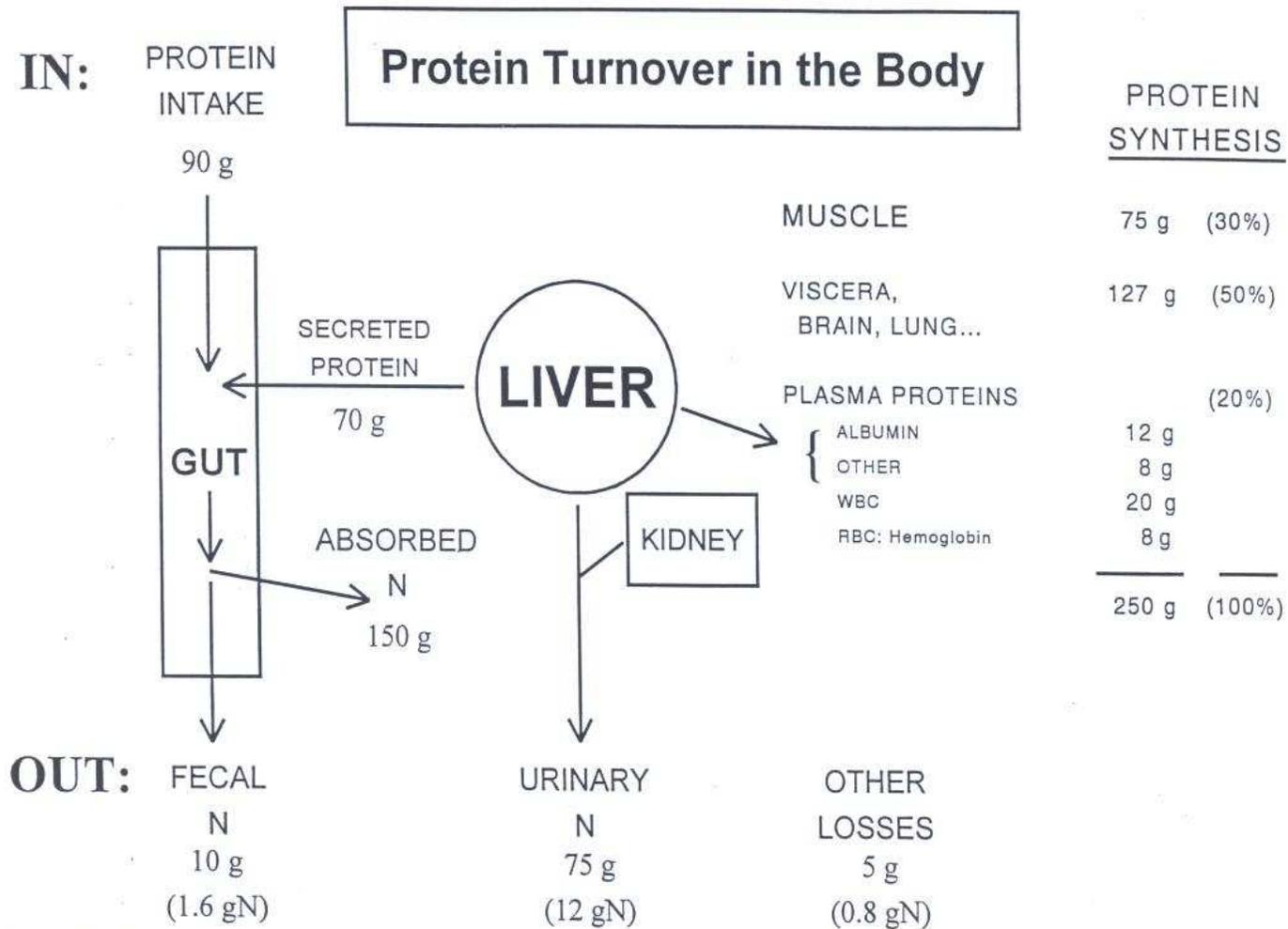
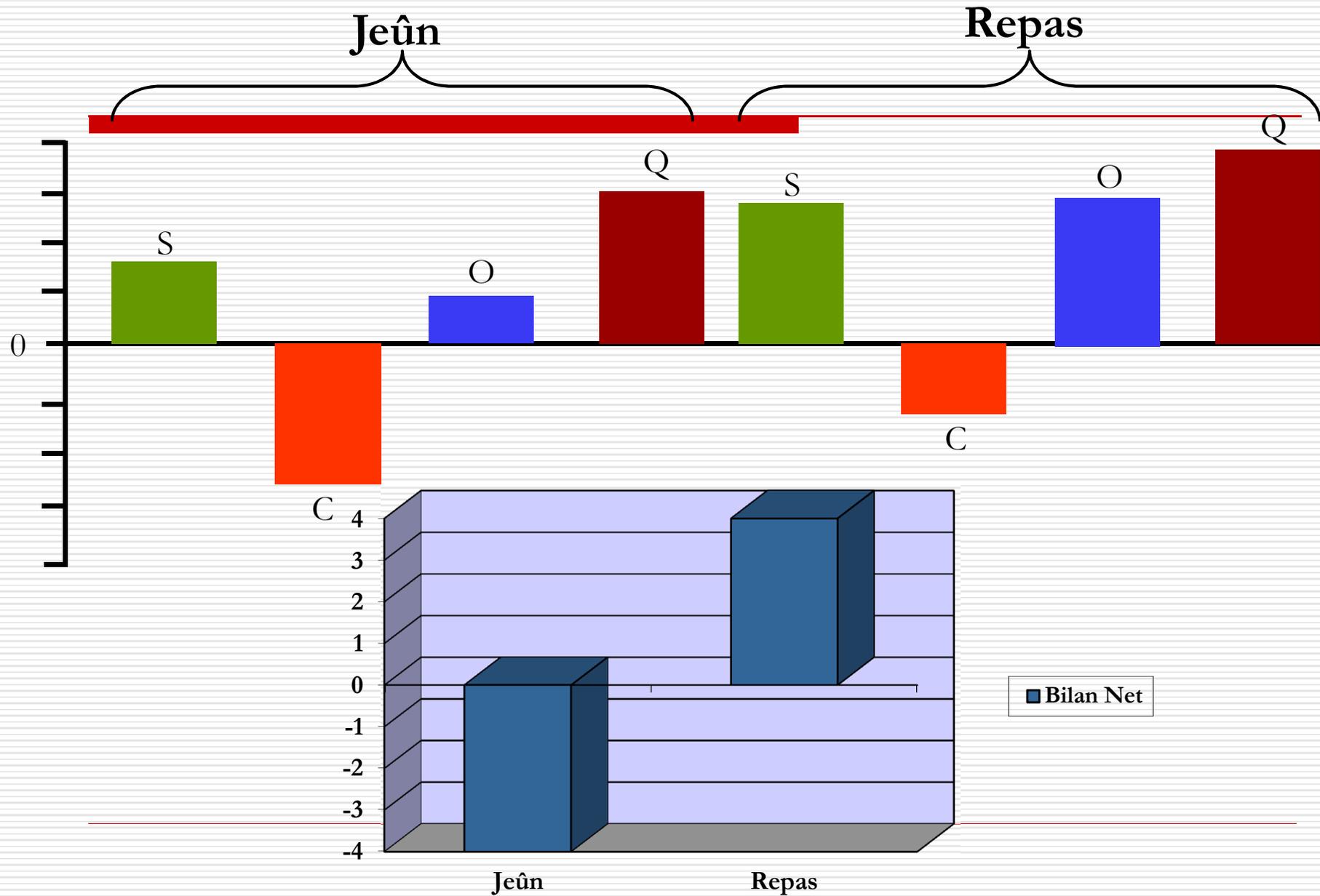


Figure 2.6. Relative rates of protein turnover and intake in a healthy 70-kg human. Under normal circumstances, dietary intake (IN = 90 g) matches N losses (OUT = 90 g). Protein breakdown then matches synthesis. Protein intake is only 90/(90 + 250) ≈ 25% of total turnover of N in the body per day. (Redrawn from Hellerstein MK, Munro HN. Interaction of liver and muscle in the regulation of metabolism in response to nutritional and other factors. In: Arias IM, Jakoby WB, Popper H, et al., eds. The liver: biology and pathobiology. 2nd ed. New York: Raven Press, 1988;965-83.)

Le cycle journalier des protéines corporelles



Besoins en protéines des adultes

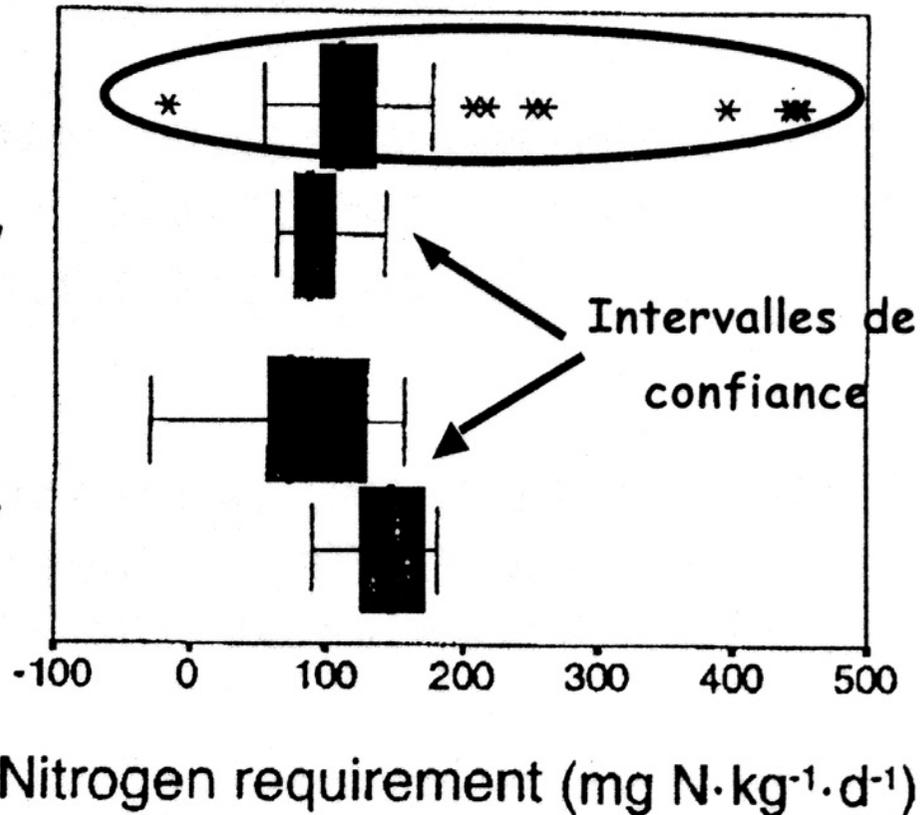
BILANS AZOTÉS: INFLUENCE DE L'ÂGE

Men < 40 y
(n = 174)

Women < 40 y
(n = 47)

Men > 67 y
(n = 7)

Women > 67 y
(n = 7)



Rand WM et al 2003

PAS DE RECOMMANDATIONS SPÉCIFIQUES
POUR LES PERSONNES ÂGÉES

= 0,8 g/kg/j

Détermination des besoins protéiques par la méthode de l'IAAO

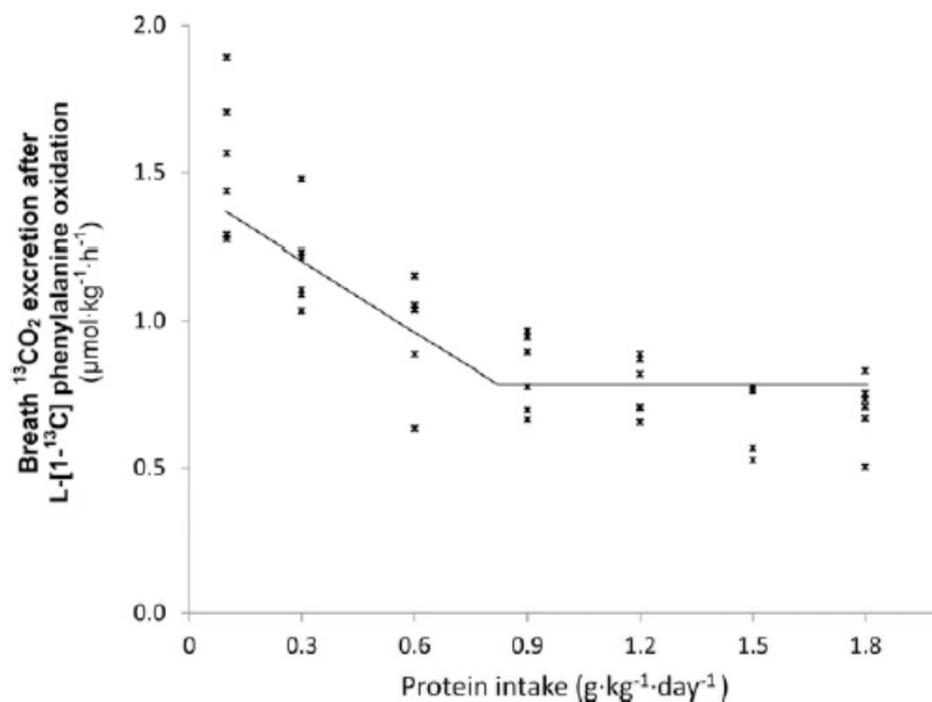


FIGURE 1. The relation between protein intake and breath ¹³CO₂ production in 6 octogenarian women. The data points represent the ¹³CO₂ excretion of individual subjects at each protein intake. The breakpoint represents the estimated mean protein requirement. The breakpoint was determined by using a nonlinear mixed model. The estimate for the mean protein requirement is 0.85 g · kg⁻¹ · d⁻¹, and the corresponding adequate protein allowance is 1.15 g · kg⁻¹ · d⁻¹.

Enquête néerlandaise

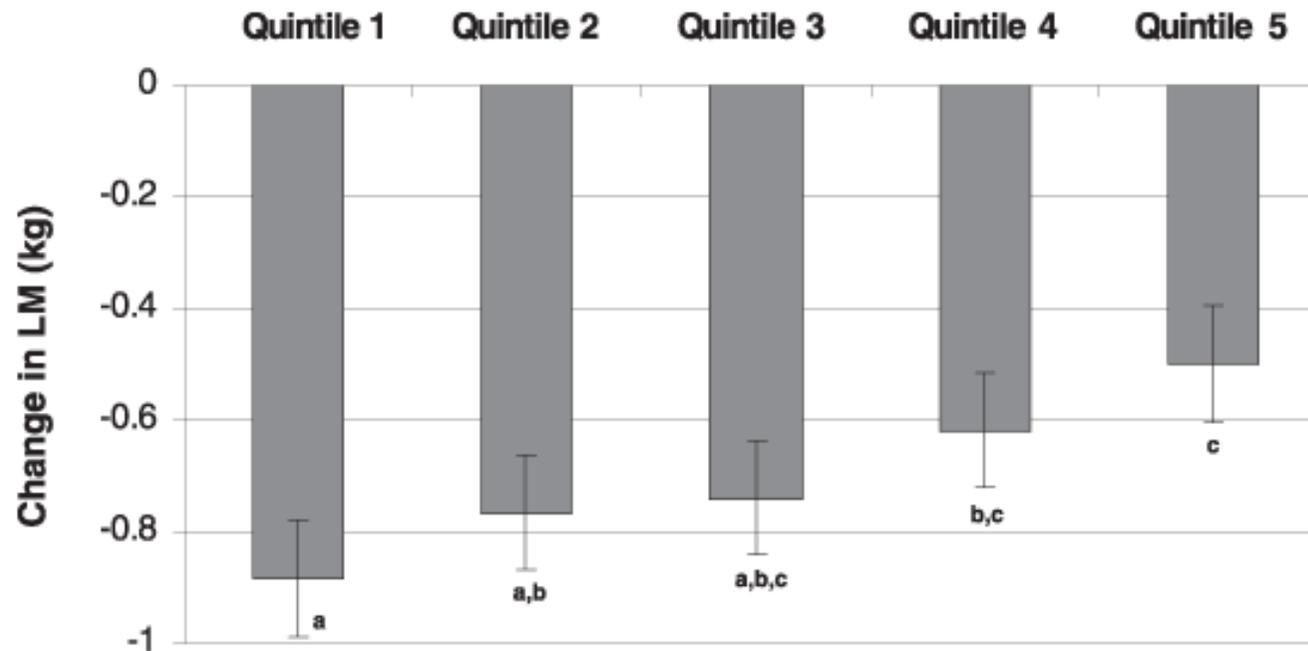
Table 2 Energy and protein intake in community-dwelling, frail, and institutionalized elderly people

	Community-dwelling		Frail FRAIL (n = 194)	Institutional	
	DNFCS (n = 400) 65–74 years	DNFCS (n = 307) 75–97 years		INST-1 (n = 60)	INST-2 (n = 216)
<i>Energy intake (MJ/day)</i>					
Men	9.4 ± 2.4 ^a	9.2 ± 2.5 ^a	8.7 ± 2.0 ^a	8.2 ± 1.6 ^a	5.8 ± 1.5 ^b
Women	7.5 ± 1.9 ^a	7.5 ± 1.8 ^{ab}	7.0 ± 1.5 ^{bc}	6.2 ± 1.5 ^{cd}	5.9 ± 1.6 ^d
<i>Protein intake (g/day)</i>					
Men	85.9 ± 23.9 ^a	81.9 ± 25.2 ^{ab}	75.4 ± 21.3 ^b	66.9 ± 18.8 ^{bc}	56.3 ± 17.1 ^c
Women	72.9 ± 18.2 ^a	71.6 ± 18.8 ^a	62.4 ± 14.9 ^b	54.0 ± 12.9 ^c	55.5 ± 15.4 ^c
<i>Protein intake (g/kg-bw/day)</i>					
Men	1.11 ± 0.31 ^a	1.07 ± 0.35 ^{ab}	1.04 ± 0.29 ^{ab}	0.86 ± 0.22 ^{bc}	0.78 ± 0.28 ^c
Women	1.03 ± 0.35 ^a	1.05 ± 0.32 ^a	1.00 ± 0.27 ^a	0.85 ± 0.20 ^b	0.81 ± 0.29 ^b
<i>Protein intake (en%)</i>					
Men	15.8 ± 3.5 ^{ab}	15.3 ± 3.2 ^{ab}	14.8 ± 2.9 ^a	13.9 ± 2.8 ^a	16.4 ± 2.6 ^b
Women	16.9 ± 3.9 ^a	16.5 ± 3.5 ^{ab}	15.3 ± 2.7 ^c	15.0 ± 3.0 ^{bc}	16.3 ± 2.5 ^{ac}

10%

35%

Données de l'étude de cohorte Health ABC



Median total protein intake as a percentage of total energy intake ($\text{g.kg}^{-1}.\text{d}^{-1}$) by quintile (from quintile 1 to quintile 5) was 11.2% ($0.7 \text{ g.kg}^{-1}.\text{d}^{-1}$), 12.7% ($0.7 \text{ g.kg}^{-1}.\text{d}^{-1}$), 14.1% ($0.8 \text{ g.kg}^{-1}.\text{d}^{-1}$), 15.8% ($0.9 \text{ g.kg}^{-1}.\text{d}^{-1}$), and 18.2% ($1.1 \text{ g.kg}^{-1}.\text{d}^{-1}$)

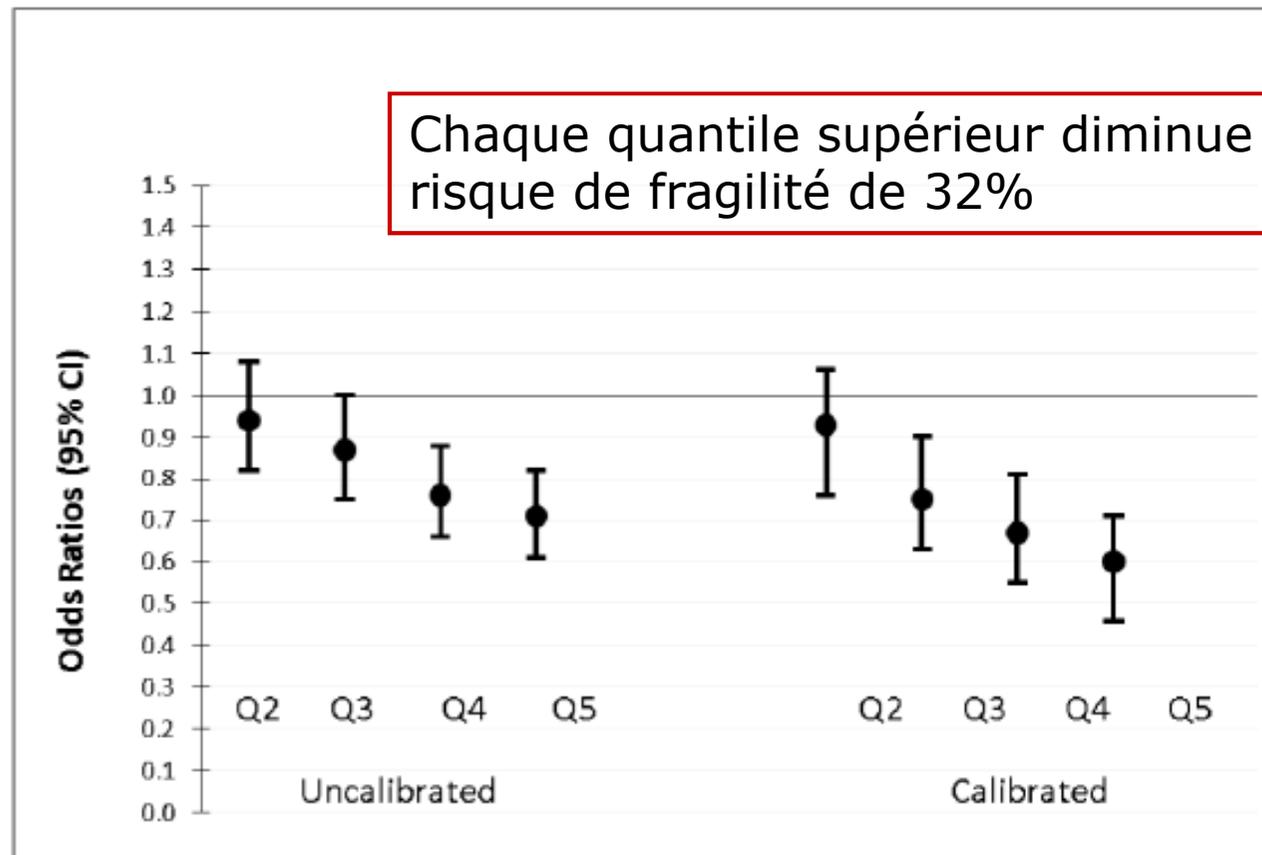
Apports protéiques et force

	Handgrip strength (kg/BW)			P
	Non-dynapenic n=20	Type I dynapenic n=23	Type II dynapenic n=30	
Lean body mass (%)	25.4±2.9	25.0±3.4	23.9±2.2	NS
Protein intake (g/d)	86.7±23.0	83.8±18.1	77.6±19.7	NS
Protein intake (%)	17.4±4.1 †	16.4±2.6	15.2±2.8	0.05
Protein intake (g/d/BW)	1.44±0.46 †	1.30±0.36 *	1.05±0.24	0.001

Devis: Cas-témoins

♀ postmenopausées

Apport protéiques et fragilité (WHS)



Études d'expérimentales





From: **Effect of 10 Days of Bed Rest on Skeletal Muscle in Healthy Older Adults**

JAMA. 2007;297(16):1769-1774. doi:10.1001/jama.297.16.1772-b

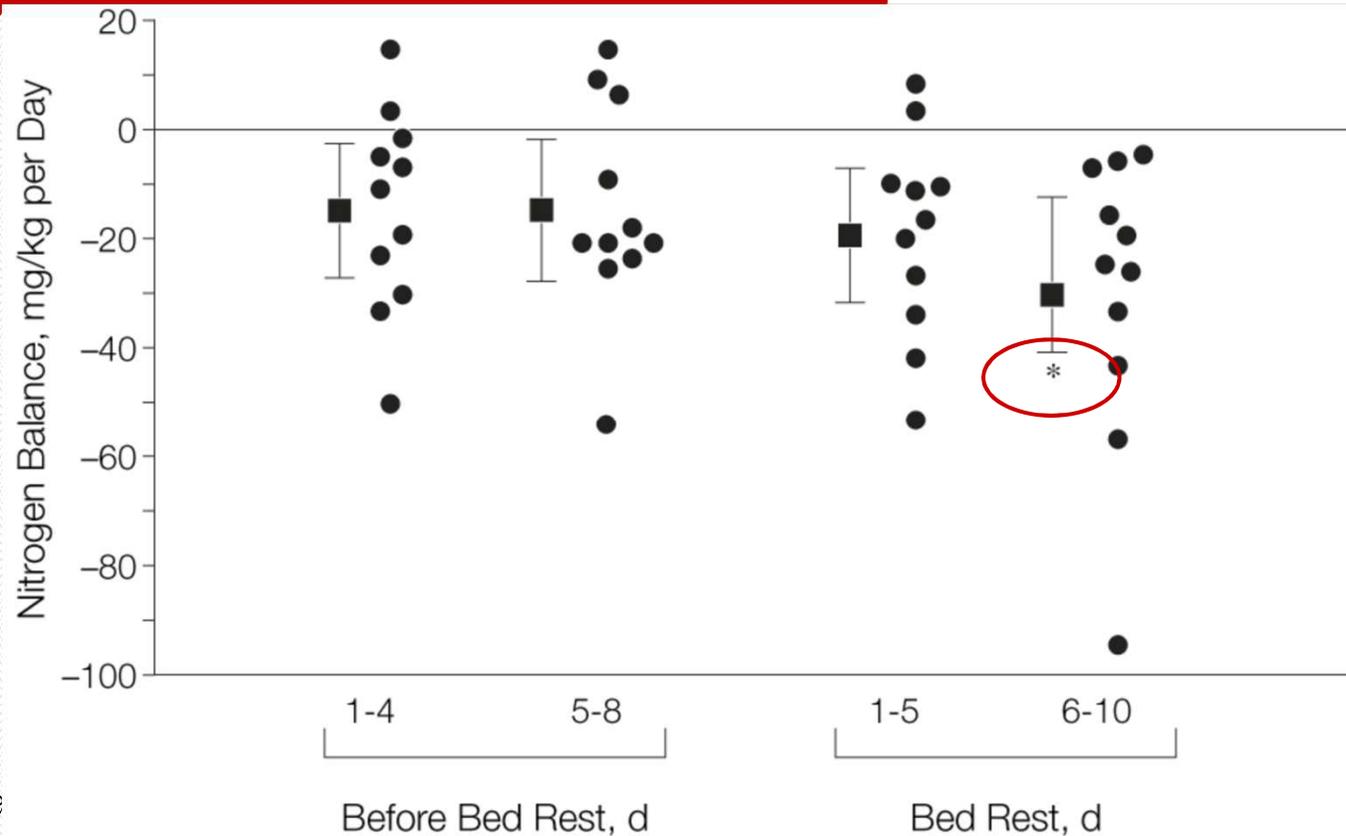


Figure Legend

Squares with error bars represent means and 95% confidence intervals for the time period. *Significantly different from days 1 through 4 and days 5 through 8 before bed rest (repeated measures analysis of variance with post hoc Tukey test, $P = .02$).

From: **Effect of 10 Days of Bed Rest on Skeletal Muscle in Healthy Older Adults**

JAMA. 2007;297(16):1769-1774. doi:10.1001/jama.297.16.1772-b

Table. Effects of 10 Days of Bed Rest in Older Adults

	No. of Participants (N = 12)*	Mean (95% Confidence Interval)			P Value
		Bed Rest		Change	
		Before	After		
Muscle fractional synthetic rate, % per h†	10	0.077 (0.059 to 0.095)	0.051 (0.035 to 0.067)	-0.027 (-0.007 to -0.047)	.02
% Change				-30.0 (-7.0 to -54.0)	
DEXA lean mass, kg‡	10				
Whole body		48.05 (40.61 to 55.49)	46.51 (39.57 to 53.45)	-1.50 (-0.62 to -2.48)	.004
% Change				-3.2 (-1.4 to -5.0)	
Lower Extremity		15.01 (12.41 to 17.61)	14.06 (11.85 to 16.27)	-0.95 (-0.42 to -1.48)	.003
% Change				-6.3 (-3.1 to -9.5)	
Isokinetic muscle strength, Nm per s§	11	120 (96 to 145)	101 (81 to 121)	-19 (-11 to -30)	.001
% Change				-15.6 (-8.0 to -23.1)	

Abbreviation: DEXA, dual-energy x-ray absorptiometry; Nm, Newton meter.

*One participant was excluded from all analyses because of insufficient protein intake.

†Because of a technical error, the muscle fractional synthesis rate measurement was excluded for 1 participant.

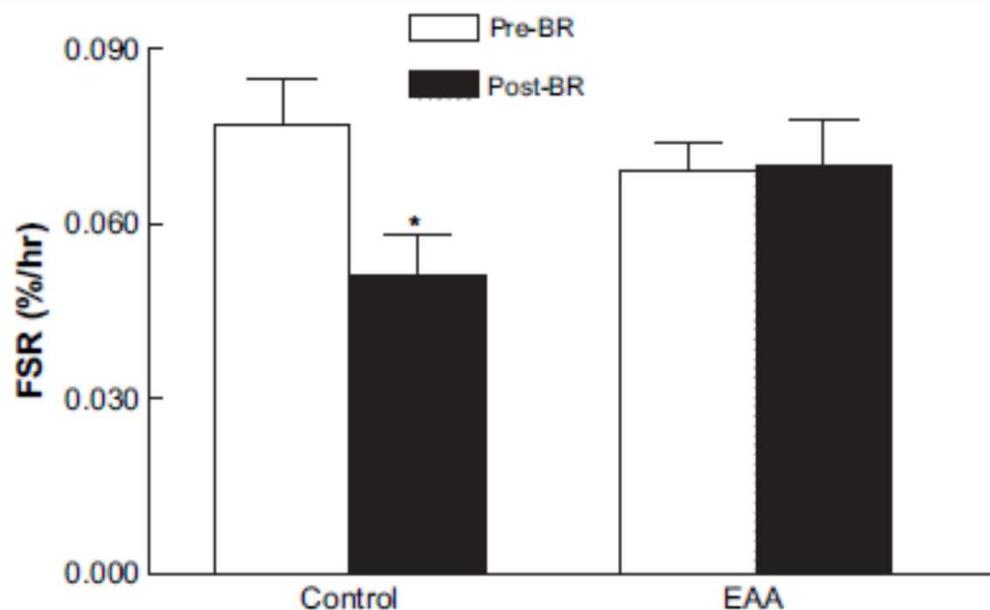
‡One participant was excluded from the DEXA analysis because the scan before bed rest was not administered.

§Isokinetic knee extension at 60° per second.

Effet d'une supplémentation AAE durant le repos au lit de 10 jours

Body mass and muscle function outcomes.

Parameter/group	Pre-bed rest		Post-bed rest		% Change		p ¹
	Control	EAA	Control	EAA	Control	EAA	
Total lean mass (kg)	46.8 (0.3), N = 9	43.0 (0.2), N = 11	45.3 (0.3)	42.1 (0.2)	-2.9 (0.9)	-2.0 (0.7)	0.29
Leg lean mass (kg)	14.5 (0.1), N = 9	13.3 (0.07), N = 11	13.6 (0.1)	12.5 (0.07)	-5.9 (1.6)	-6.3 (1.8)	0.77
Total fat mass (kg)	27.0 (0.3), N = 9	27.3 (0.1), N = 11	27.0 (0.3)	27.3 (0.3)	0	0	0.86
Standing plantar flexion (rep/30 s)	21.8 (3.4), N = 7	21.8 (1.4), N = 12	20.9 (2.8)	21.4 (2.5)	-0.19 (0.04)	-0.019 (0.10)	0.054
Stair ascent power (Nm/s)	407.2 (69.9), N = 8	293.5 (37.2), N = 11	336.5 (47.6)	284.2 (43.0)	-14 (4)	-2 (5)	0.029
Stair descent power (Nm/s)	440.5 (75.7), N = 8	320.6 (43.1), N = 12	339.6 (50.6)	290.3 (45.9)	-21 (4)	-8 (4)	0.026
Floor transfer (s)	8.3 (1.1), N = 11	9.6 (1.6), N = 12	12.8 (2.3)	10.5 (1.7)	51 (17)	13 (7)	0.027



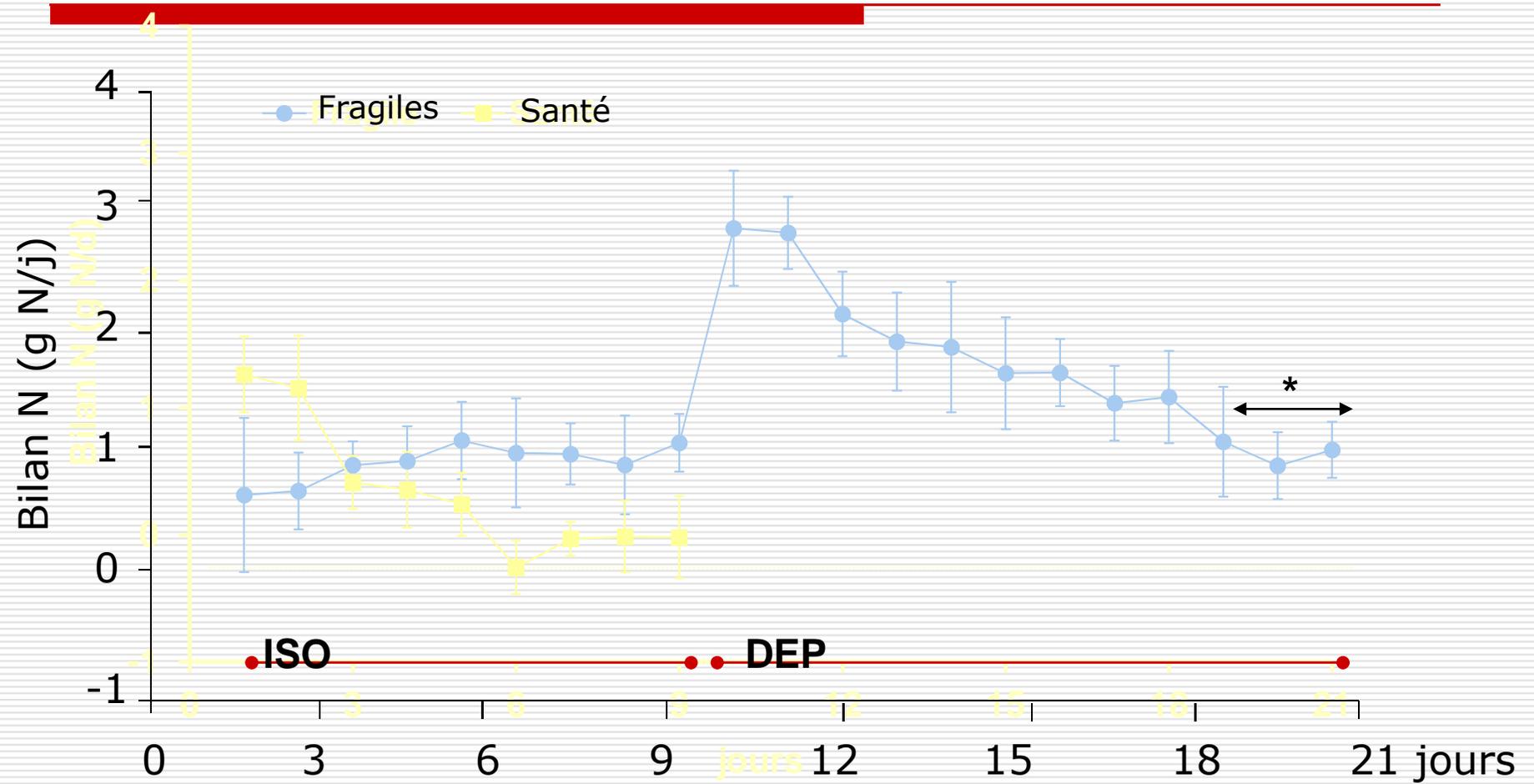
15g AAE 3x/j

Apport protéique

	Fragile ISO	Fragile DEP	Santé ISO
Apport protéique			
(g/d)	38 ± 3 *	54 ± 5 * †	68 ± 3
(g/kg PC.d)	0.87 ± 0.03 *	1.23 ± 0.02 †	1.25 ± 0.08
(g/kg MM.d)	1.25 ± 0.09 *	1.78 ± 0.12 †	1.95 ± 0.10
(% de l'énergie)	12 ± 1 *	17 ± 1 †	15 ± 1

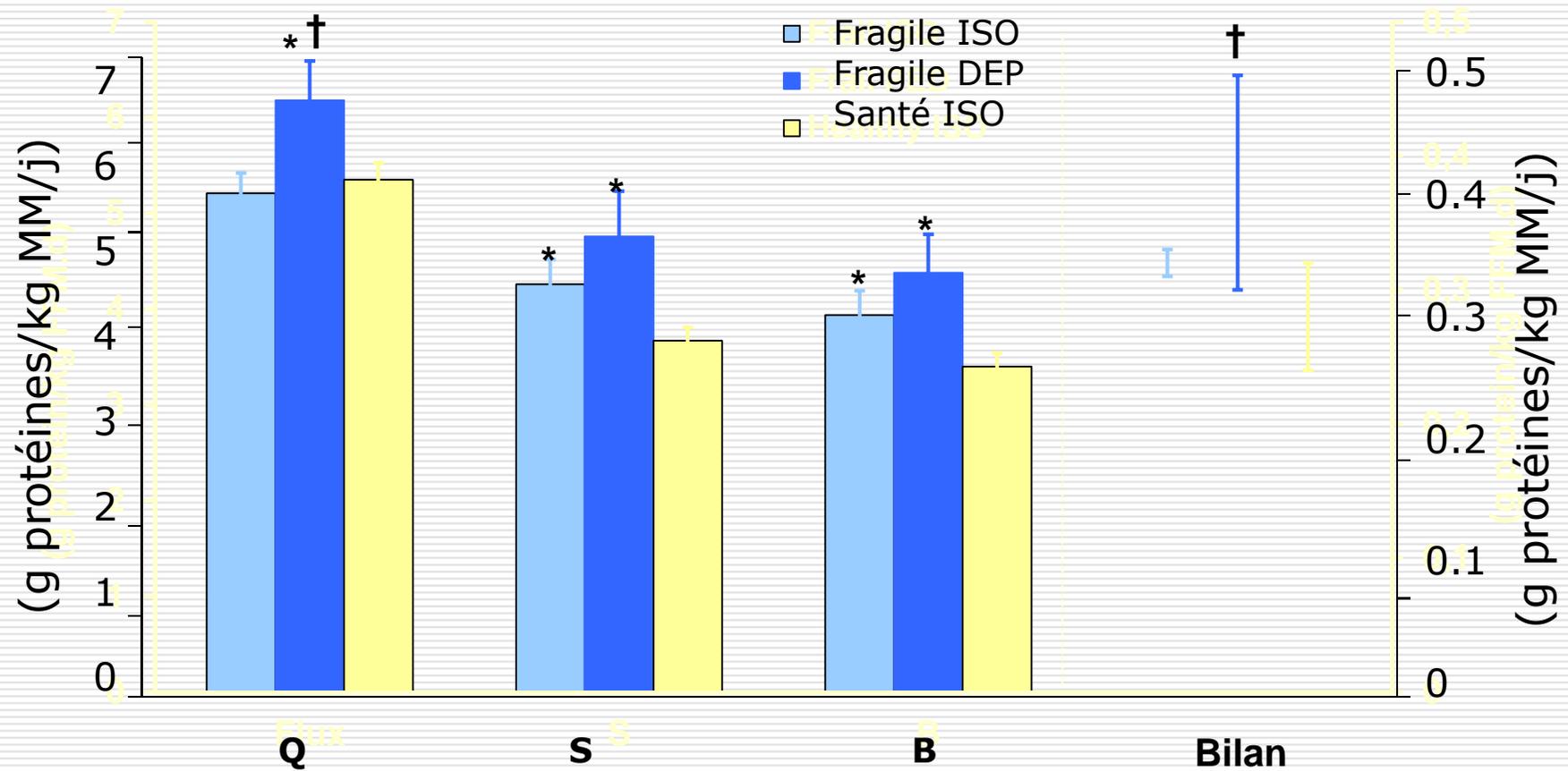
Moyenne ± ETM. *P<0.005 vs Santé ISO †P<0.05 vs Fragile ISO

Bilan azoté



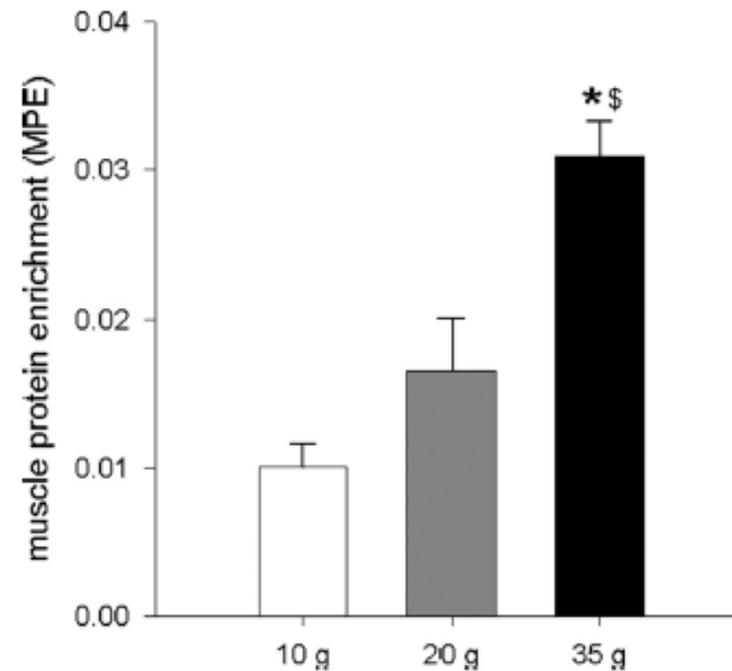
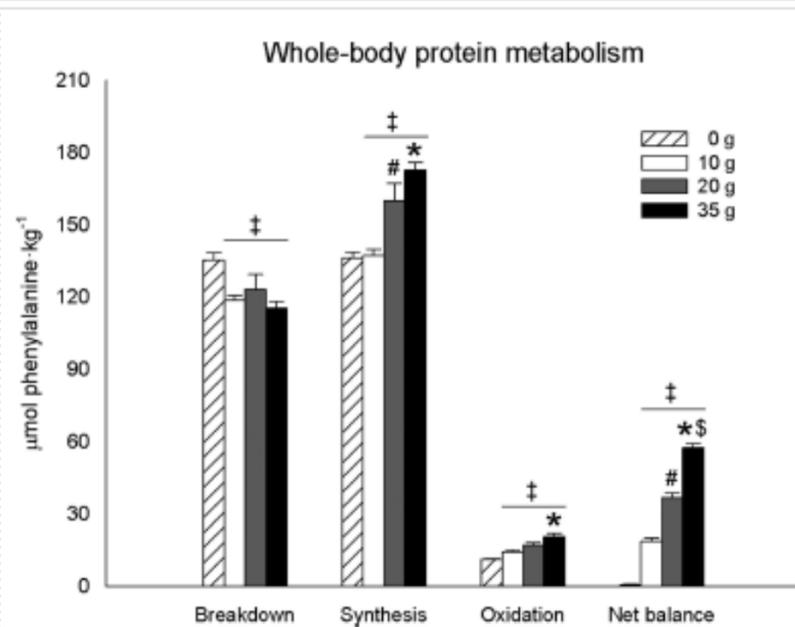
* différent de 0, $P < 0.05$

Cinétique des protéines corporelles



Mean \pm SEM. * $P < 0.05$ vs Healthy ISO † $P < 0.05$ vs Frail ISO

Effets des apports protéiques de petit lait



Effets aigus d'un repas

N = 11/groupe

Synthèse protéique est un système saturable

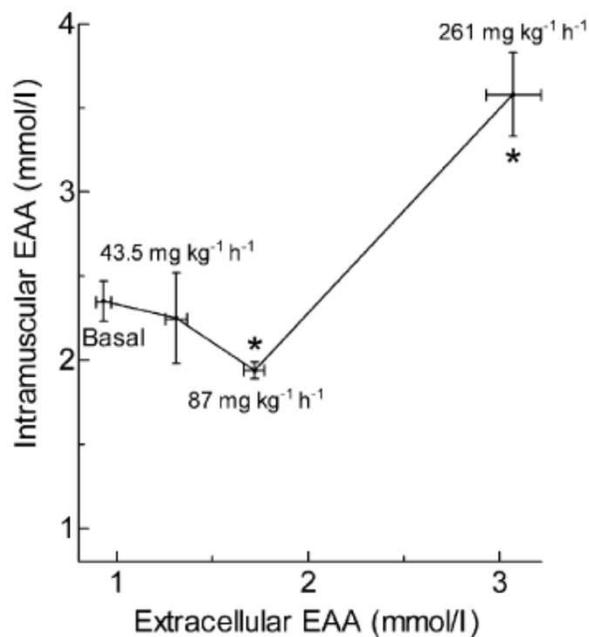
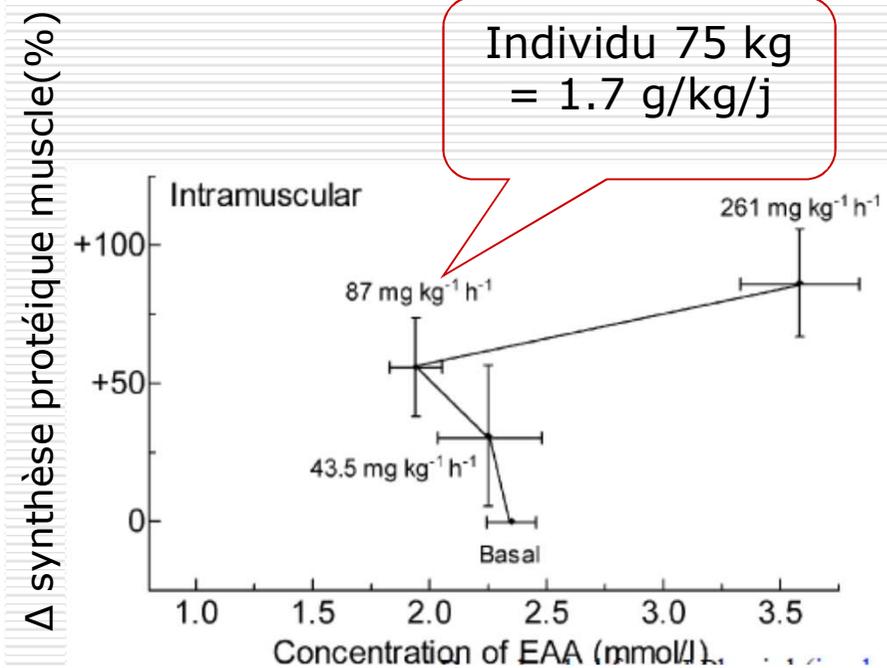
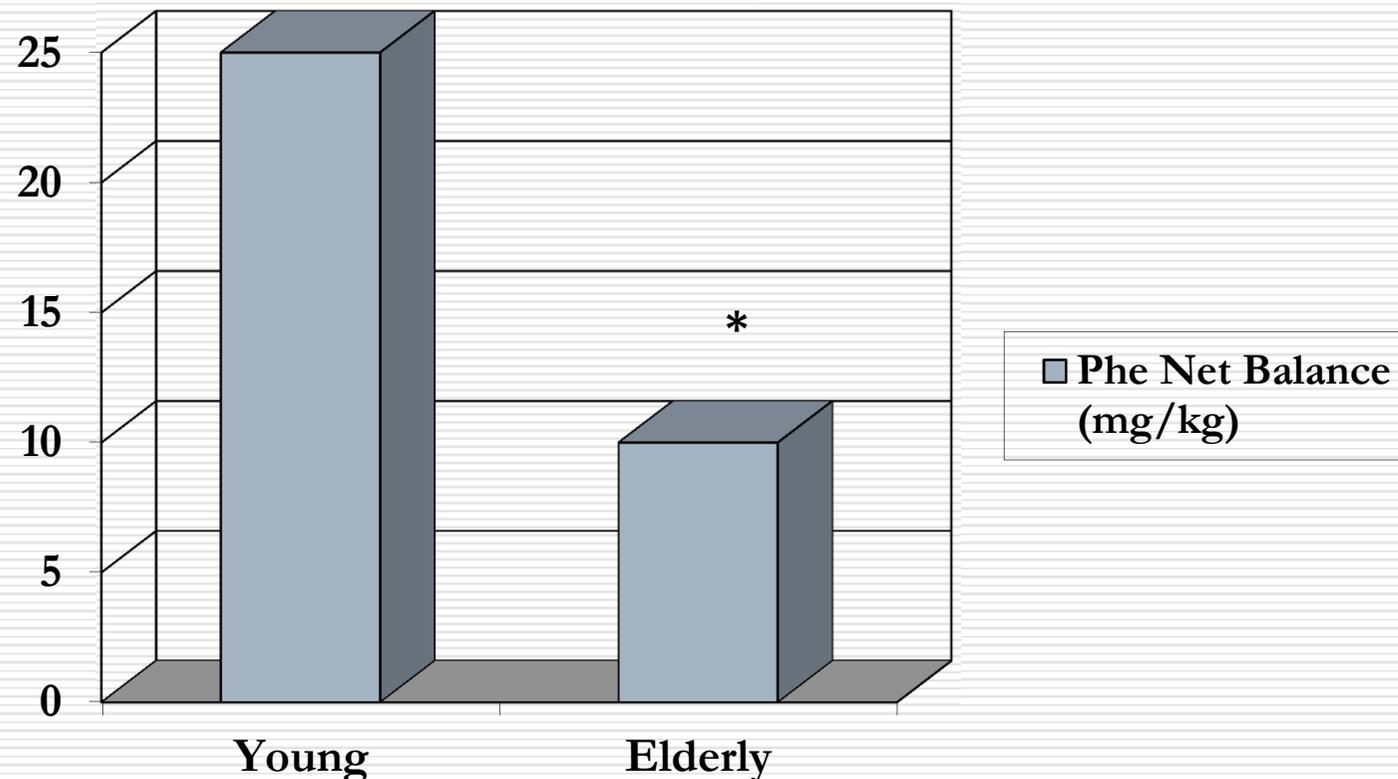


Figure 3. Relationship between extracellular and intramuscular concentrations of EAAs during infusion of mixed amino acids at different rates

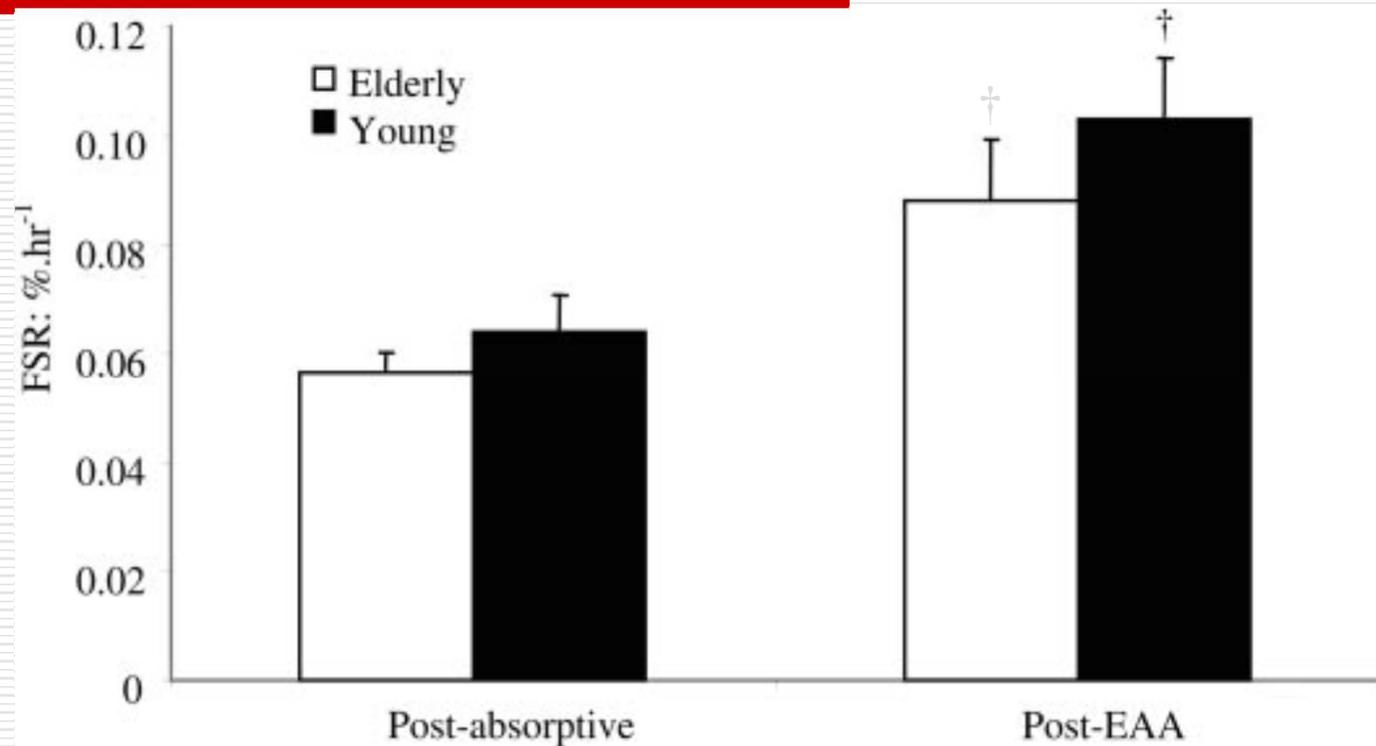


Réponse musculaire moindre avec un petit bolus AAE



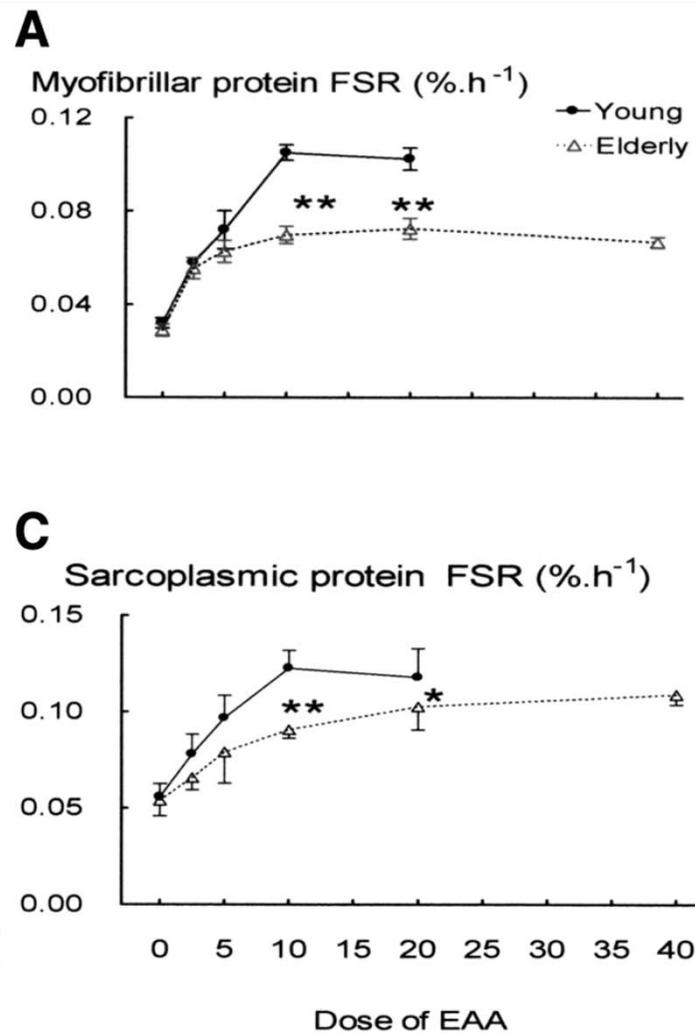
Mean (SEM) leg phenylalanine net balance 3.5 h after the ingestion of 7g EAA calculated by measuring the area under the phenylalanine net balance response curve in the elderly (*n*11) and the young (*n*8). Data were analyzed with a *t* test. *Significantly different from the young, *P* 0.010.

Une quantité suffisante d'AAE améliore la synthèse protéique musculaire

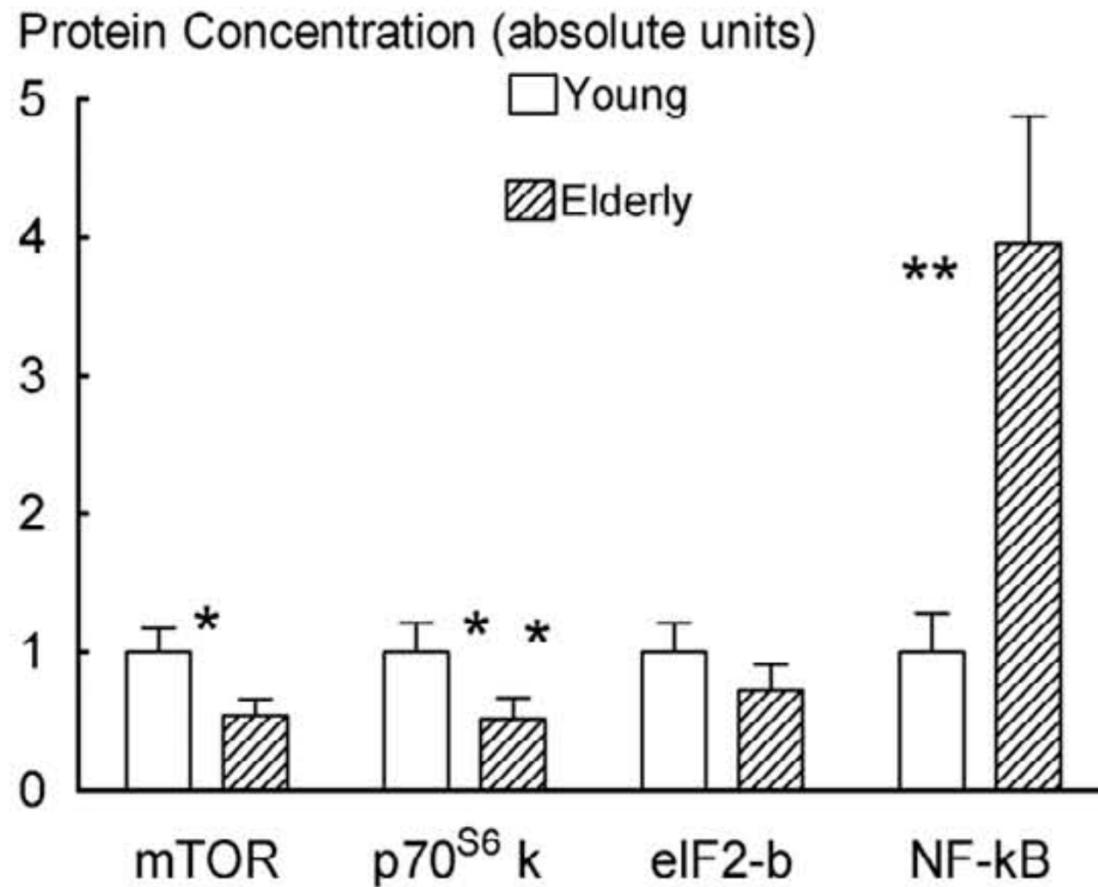


Mixed muscle fractional synthetic rate (FSR) in young and elderly before and after ingestion of **15 g of EAA**. †Significant difference from corresponding postabsorptive values: young, P 0.012; elderly, P 0.029.

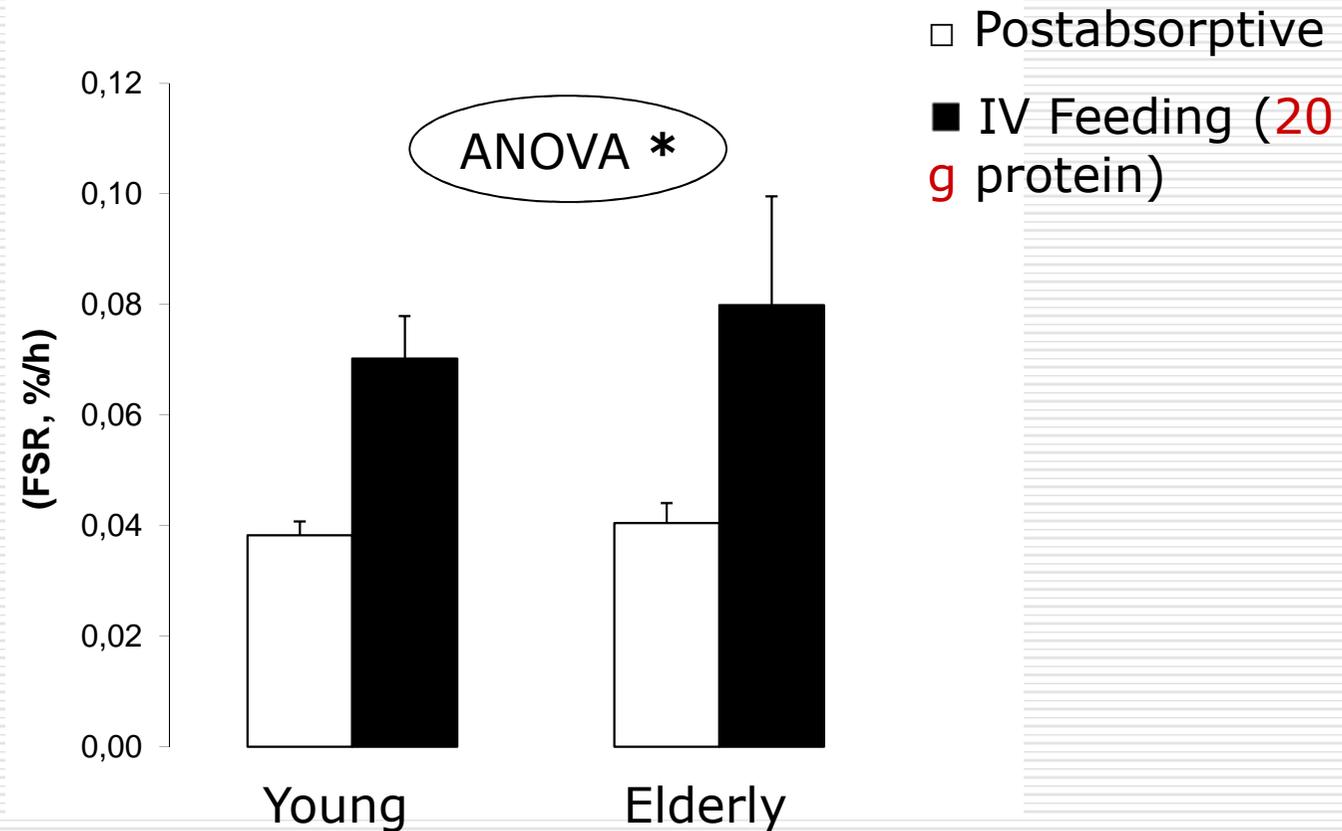
Résistance anabolique



Causes de résistance anabolique



Réponse musculaire au repas intraveineux chez la femme âgée



*Feeding effect, $P < 0.01$ Chevalier S, Morais et al J Gerontol A Biol Sci Med Sci, 2011

Distribution étalée et pulsée des protéines

N=8; dévis chassé-croisé
Étalé: 30-30-30 g
Pulsé: 10-16-64 g

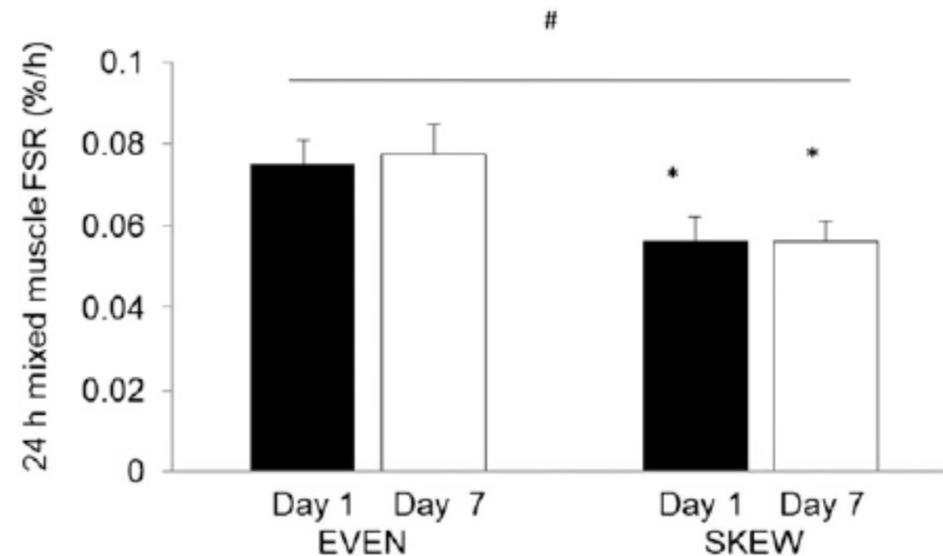
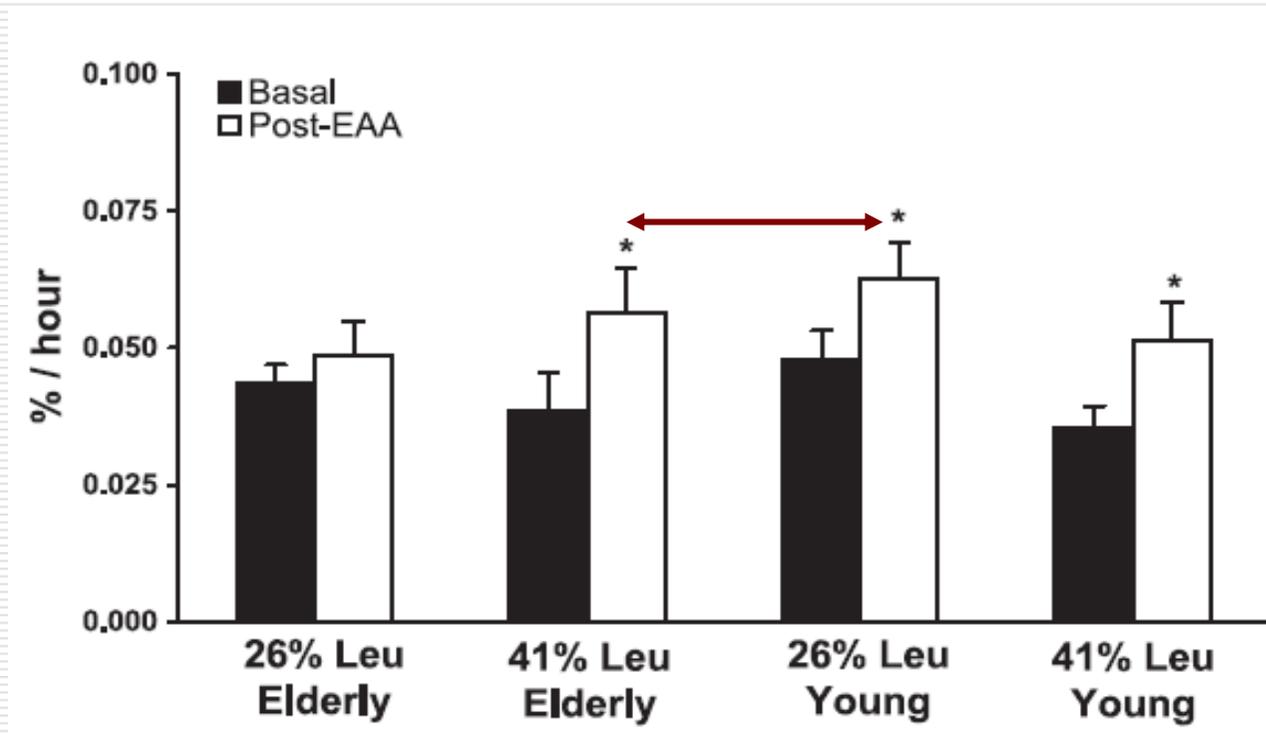


FIGURE 3 Twenty-four-hour mixed muscle protein FSRs in healthy adults on days 1 and 7 after the ingestion of diets with an EVEN or SKEW protein distribution. Values are means \pm SEMs; $n = 8$. *Different from EVEN at that time point, $P < 0.05$. #Main effect of group between EVEN and SKEW, $P < 0.05$. EVEN, even daily protein distribution; FSR, fractional synthesis rate; SKEW, skewed daily protein distribution.

Synthèse protéique musculaire: hommes âgés vs. jeunes

En réponse à l'ingestion d'une boisson d'acides aminés essentiels +/- enrichie en leucine



Effet à long terme de la supplémentation en leucine

Body composition¹ Intervention: 2.5 g leucine/repas x 3 mois

	Placebo (<i>n</i> = 14)		Leucine (<i>n</i> = 15)	
	Before	After	Before	After
Lean mass (kg)	55.8 ± 0.9	56.2 ± 1.1	54.6 ± 1.0	55.0 ± 1.5
Fat mass (kg)	19.8 ± 1.7	19.2 ± 2.0	20.0 ± 1.4	20.0 ± 1.3
Body fat (%)	24.5 ± 1.7	23.9 ± 1.9	25.3 ± 1.2	25.4 ± 1.2
Leg lean mass (kg)	17.6 ± 0.4	18.0 ± 0.4	17.1 ± 0.5	17.6 ± 0.4
Leg fat (%)	18.9 ± 1.5	19.4 ± 1.6	19.6 ± 1.2	19.8 ± 1.2
CSA (cm ²)	71 ± 3	71 ± 3	71 ± 2	71 ± 2
Leg volume (L)	7.5 ± 1.9	7.5 ± 1.7	8.1 ± 3.0	7.8 ± 4.1

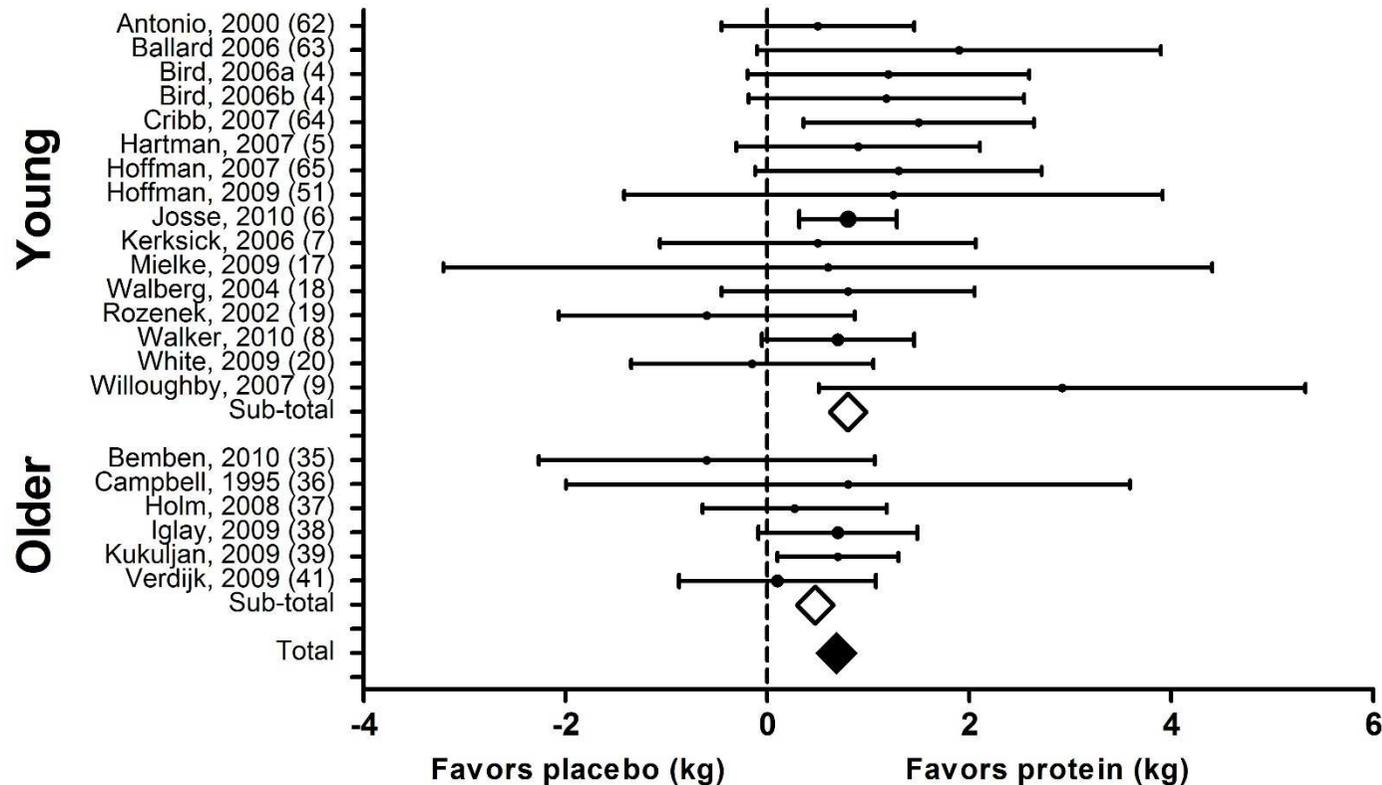
¹ All values are means ± SEMs. CSA, cross-sectional area. Data were analyzed by using repeated-measures ANOVA. No significant differences were observed between groups or over time.

Measures contrer la perte de la masse et de la force musculaire



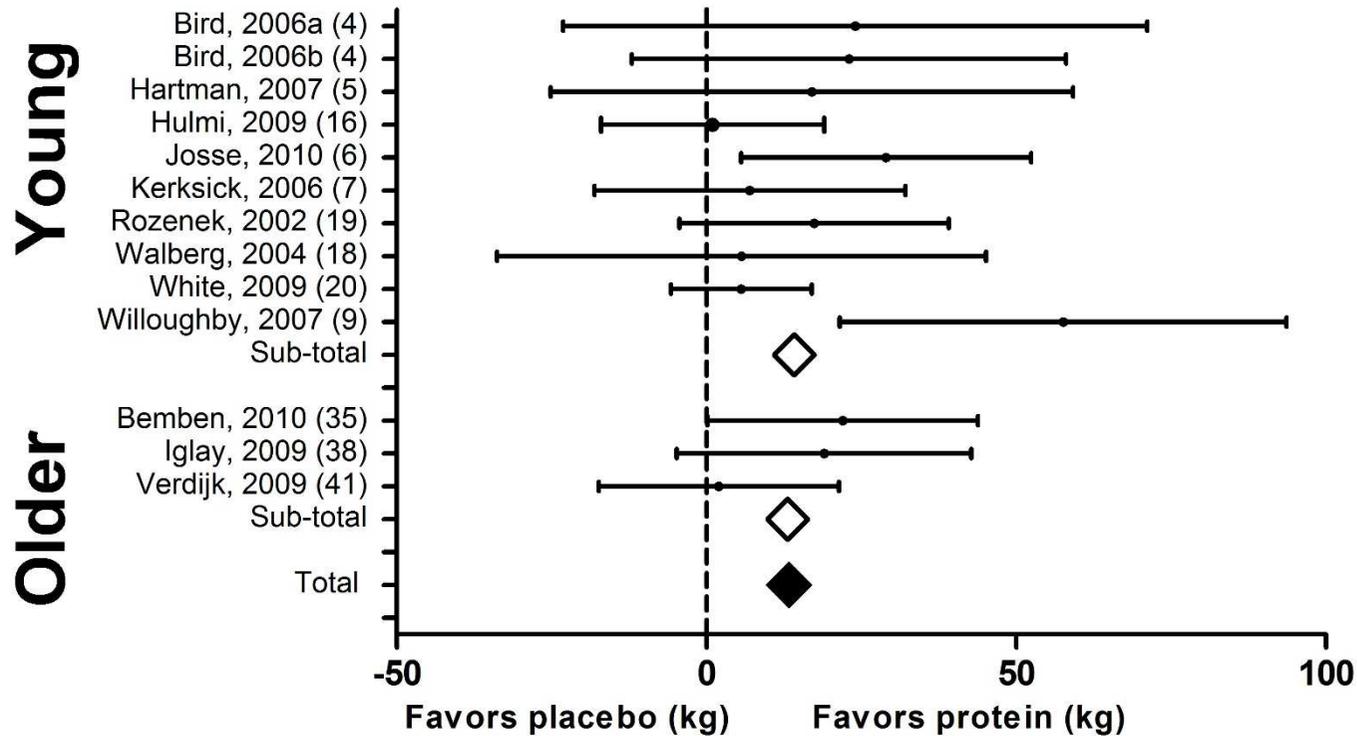
Olga Kotelko, 93 y.o

Programme d'exercice et supplémentation protéique - effet sur la masse maigre



Weighted mean difference: 0.69 kg; 95% CI: 0.47, 0.91 kg; P < 0.00001

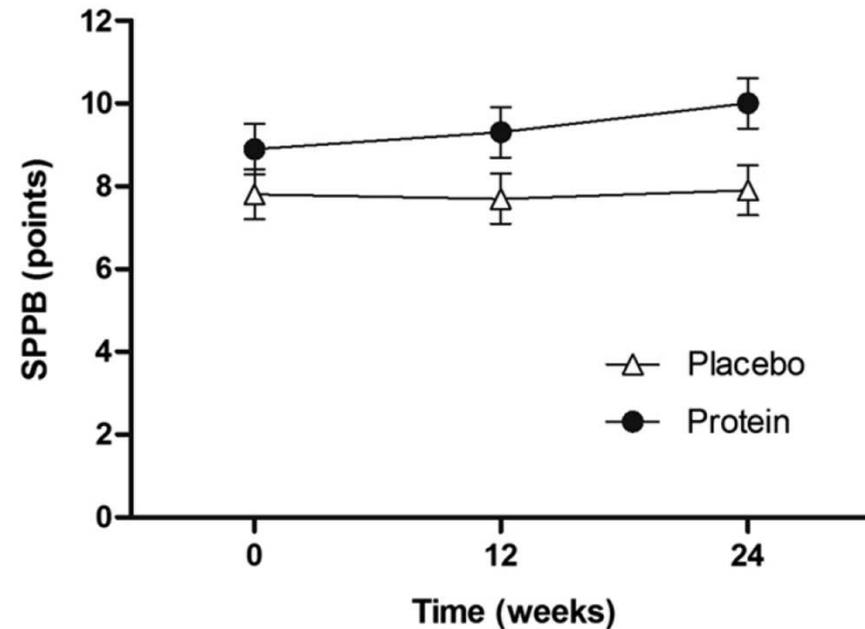
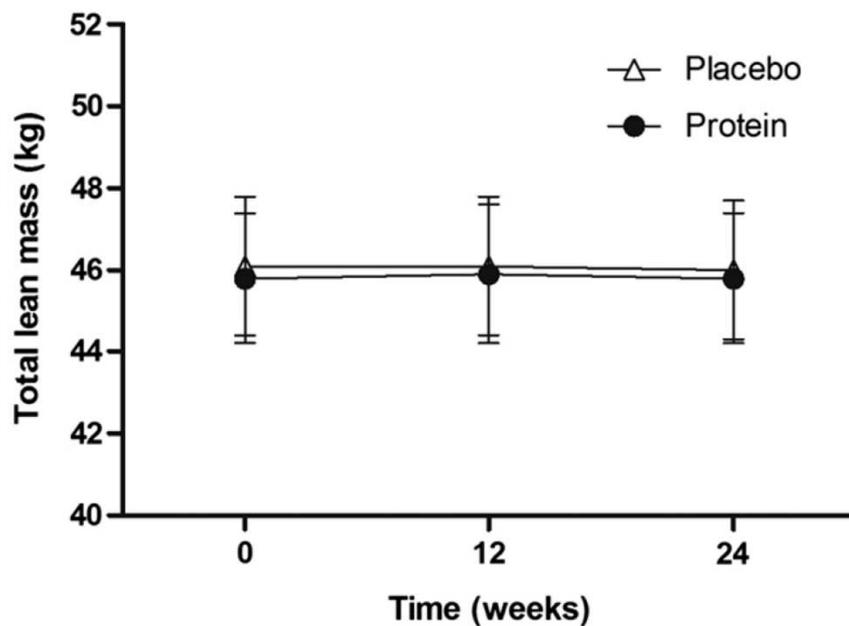
Programme d'exercice et supplémentation protéique - effet sur 1-RM leg press



Weighted mean difference: 13.5 kg; 95% CI: 6.4, 20.7 kg; $P < 0.005$

Supplémentation protéique chez la PA fragilisée

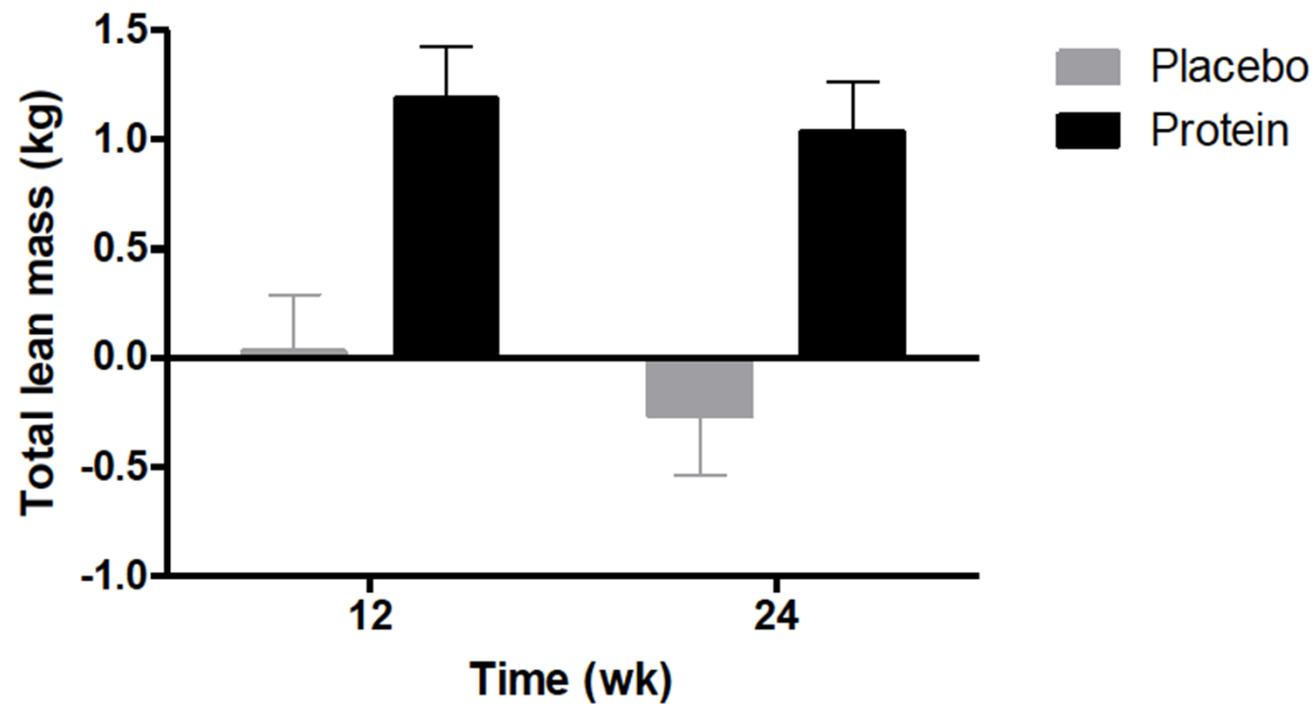
RCT of 15 g of protein supplementation BID, n=65



Treatment x Time interaction effect $p < 0.02$

Supplémentation protéique chez la PA fragilisée

RCT of Resistance EX 2 sessions / week \pm 15 g of protein BID



Risques de la diète hyperprotéique



Y-a-t-il des risques?

- Il y a potentiellement 3
 - Effet suppresseur de l'appétit (satiété)
 - Diète hyperprotéique: réputée efficace pour faire maigrir ([Whycherley TP AJCN 2012](#))
 - Bilan phospho-calcique
 - Protéines en moyenne sont acides et augmentent l'excrétion de Ca^+ ([Kurtz I Kidney Intern 1983](#))
 - Atteinte rénale
 - Protéines augmentent le flow sanguin causant des blessures glomérulaires considérées comme pathogénèse de la sclérose glomérulaire progressive ([Brenner BM NEJM 1982](#))
-

Protéines et bilan énergétique

	Percentage of energy from protein		
	5%	15%	30%
Energy intake (MJ/d)			
Total	9.33 ± 3.52 ^{2,a}	9.62 ± 3.51 ^a	7.21 ± 3.08 ^b
Snacks	2.15 ± 2.09 ^a	2.44 ± 2.35 ^a	1.16 ± 1.13 ^a
Meals	7.18 ± 2.38 ^a	7.13 ± 2.39 ^a	6.04 ± 2.12 ^b
Breakfast	2.56 ± 0.91 ^a	2.38 ± 0.91 ^b	2.19 ± 0.93 ^c
Lunch	2.10 ± 0.90 ^a	2.02 ± 0.88 ^a	1.56 ± 0.68 ^b
Dinner	2.52 ± 0.99 ^a	2.73 ± 1.03 ^b	2.29 ± 0.94 ^c
Protein intake, meals			
(MJ/d)	0.36 ± 0.12 ^a	1.07 ± 0.36 ^b	1.81 ± 0.64 ^c
(g · kg BW ⁻¹ · d ⁻¹)	0.3 ± 0.1 ^a	0.9 ± 0.3 ^b	1.6 ± 0.5 ^c
Carbohydrate intake, meals			
(MJ/d)	4.31 ± 1.43 ^a	3.57 ± 1.20 ^b	2.12 ± 0.74 ^c
(g · kg BW ⁻¹ · d ⁻¹)	3.7 ± 1.2 ^a	3.1 ± 1.1 ^b	1.8 ± 0.6 ^c
Fat intake, meals			
(MJ/d)	2.51 ± 0.83 ^a	2.50 ± 0.84 ^a	2.12 ± 0.74 ^b
(g · kg BW ⁻¹ · d ⁻¹)	1.0 ± 0.3 ^a	1.0 ± 0.3 ^a	0.8 ± 0.3 ^b
Protein:carbohydrate + fat			
Meals (% of energy)	5:95 ³	15:85	30:70
Total (% of energy)	5:95	12:88	23:77
$\Delta BW_{\text{day12}} - \text{day1}$ (kg)	-0.9 ± 1.1	-0.8 ± 1.0	-1.0 ± 1.0
Energy balance (MJ/d)	-2.48 ± 2.92 ^a	-2.20 ± 2.84 ^a	-4.61 ± 4.64 ^b

N = 79

Durée: 12 j

Age: 34 ans

VAS: même

Apports protéiques et perte pondérale dans l'étude de cohorte NuAge

TABLE 3 Unadjusted and fully adjusted conditional regression models for weight loss by protein-intake category in community-dwelling older adults

Protein-intake category	Unadjusted model		Fully adjusted model ¹	
	OR (95% CI)	Parameter estimate <i>P</i> value	OR (95% CI)	Parameter estimate <i>P</i> value
Low: <0.8 g/(kg · d)	1.92 (1.12, 3.31)	0.018	2.56 (1.01, 6.50)	0.048
Moderate: 0.8–<1.0 g/(kg · d)	1.71 (1.04, 2.80)	0.039	2.15 (1.02, 4.56)	0.045
High: 1.0–<1.2 g/(kg · d)	1.33 (0.77, 2.28)	0.31	1.60 (0.81, 3.15)	0.17
Very high: ≥1.2 g/(kg · d)	Reference		Reference	

¹ Adjusted for energy intake, BMI, smoking, physical activity, physical function, dieting to lose weight, appetite, number of medications, number of chronic diseases, depressive symptoms, and serum albumin concentrations. All covariates were simultaneously forced in this multivariable conditional logistic regression model. In this model, except for protein intake, categorical variables were treated as continuous. Model likelihood ratio test: chi-square = 31.797, df = 14, *P* = 0.004, *n* = 183 pairs.

Effet de la diète hyperprotéique sur le métabolisme du Ca⁺

Excrétion urinaire

	wk 0 ²	wk 1	wk 2	wk 3	wk 5	wk 7	Pooled SD	Diet
Calcium, <i>mmol/d</i>							1.7	0.005
LPLP	4.2	3.9	3.6	3.9	4.2	4.0		
HPHP	5.3	5.0	5.2	4.9	5.3	5.0		
Magnesium, <i>mmol/d</i>							0.9	0.03
LPLP	2.9	4.1	3.8	3.8	4.0	3.9		
HPHP	3.6	3.3	3.5	3.1	3.5	3.6		
Phosphorus, <i>mmol/d</i>							5	<0.0001
LPLP	23	19	17	17	19	17		
HPHP	26	30	30	27	30	32		

	LPLP	HPHP	Pooled SD	P-value
⁴⁷ Ca retention at d 21, ² %	16.9	19.7	3.8	0.05
Absorption, ³ %	22.3	26.5	5.4	0.05
Ca absorbed, <i>mg/d</i>	200	227	46	0.12

Devis: croisé

N=16

Sujets: ♀ en postménopause

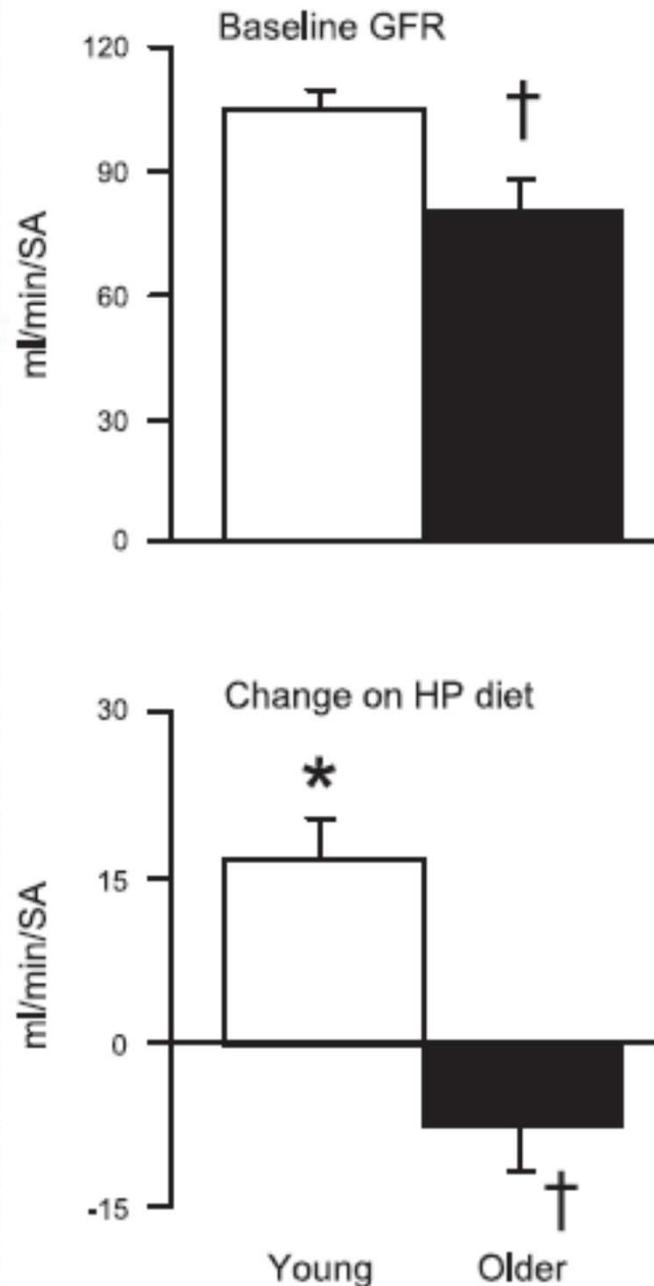
Apport protéique et changement DMO (Framingham Offspring Study)

Variable	Femoral Neck BMD				Trochanter BMD				Lumbar Spine BMD			
	<i>n</i>	β	<i>SE</i>	<i>p-value</i>	<i>n</i>	β	<i>SE</i>	<i>p-value</i>	<i>n</i>	β	<i>SE</i>	<i>p-value</i>
Men												
Model 1 ^b	497	-0.0216	0.017	0.19	497	-0.0359	0.018	0.04*	489	0.00007	0.017	0.99
Model 2 ^c	493	-0.0052	0.019	0.78	493	-0.0498	0.020	0.01*	485	-0.0062	0.019	0.75
Women												
Model 1	680	-0.0001	0.016	0.99	680	-0.0090	0.021	0.68	678	0.01439	0.016	0.38
Model 2	673	-0.0131	0.017	0.44	673	-0.0288	0.023	0.21	671	0.0042	0.018	0.81

Chez la femme, apport protéique élevé est protecteur pour celles qui ont des apports bas en Ca⁺.

Apport protéique et Fct rénale

Devis croisé
N=10 jeunes vs 9 âgés
0.9 vs 1.8 g/kg/j
Durée: 10 j



Conclusions

- Les évidences s'accumulent à l'effet que les personnes âgées requièrent plus de protéines dans leur diète par kg de PC que les jeunes gens
 - Nouvelles techniques le confirment (IAAO)
 - Démonstration de la résistance anabolique aux AA
 - Études épidémiologiques à l'appui
 - Les apports recommandés se situent entre 1-1.2 g/kg/j et devraient même être de 1.3-1.5 g/kg/j si malnutrition et mal. chroniques
 - Endossé par la **EUGMS et PROT-AGE Study group** (Bauer J JAMDA 2013), **ESPEN Expert Group** (Deutz N E P Clin Nutr 2014)
-

Conclusions

- ❑ Le but est de prévenir la perte de la masse et de la force musculaire avec l'âge
 - ❑ L'exercice physique demeure un atout et devrait être encouragé
 - Un peu activité vaut mieux que rien!
 - ❑ Les effets néfastes, i.e., diminution des apports énergétiques, perte de la masse osseuse, et atteinte de la fct. rénale, ne se sont avérés vrais que pour la fct. rénale
 - On devrait se restreindre à 1 g/kg/j si TFG <30 mL/min.
-

Merci

