

HUR Australia Webinar April 2021



MUSCLES AND MOBILITY MATTER

LESSONS FROM RESEARCH TO INFORM PRACTICE



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FACT #1

Skeletal muscle tissue has a turnover rate of 1–2% per day implying full renewal of muscle tissue every **2–3 months**

Smeets et al Brain 141(4), 112-29, 2018

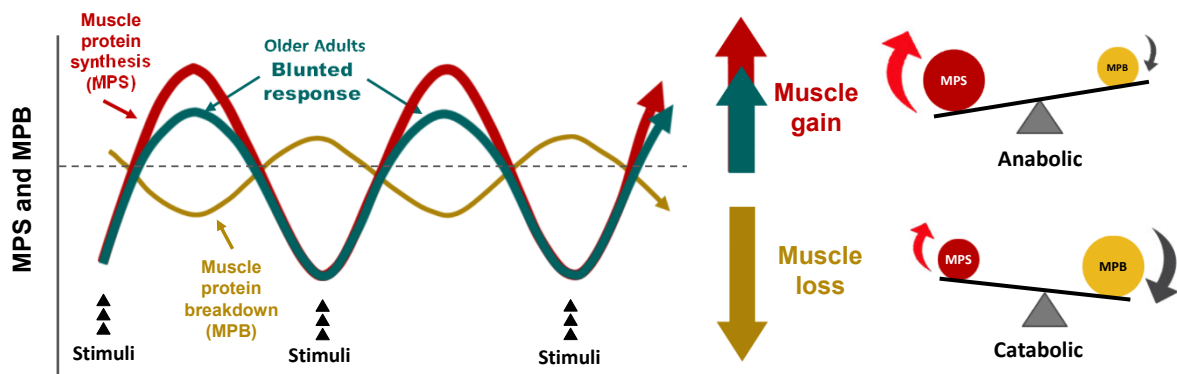
Photo by engin akurt on Unsplash



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Understanding Muscle Loss

Why We Lose Muscle – ‘Anabolic Resistance’ of Muscle to Ageing



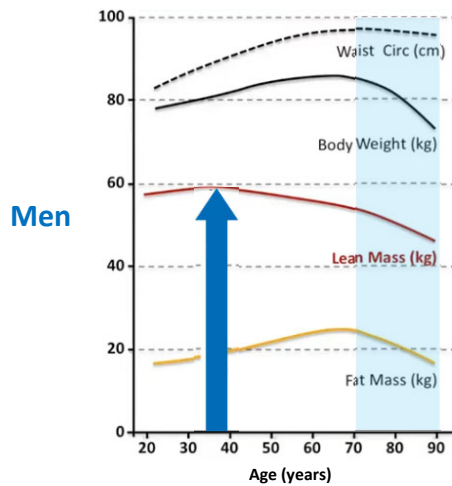
Adapted from Breen and Phillips. Nutrition Metab 2011;8:68

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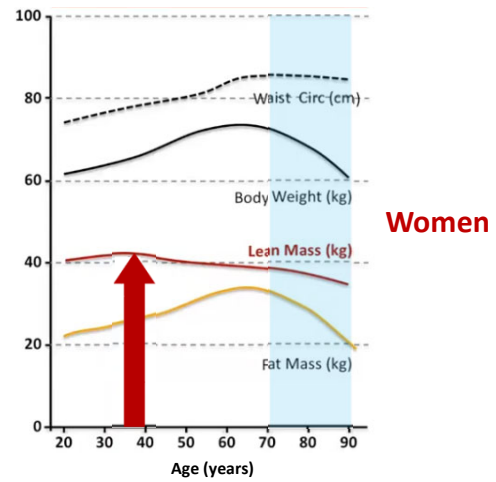
Age-related Skeletal Muscle Loss

Is skeletal muscle loss an early marker of ageing?



Muscle loss starts around 40-45 years

...but muscle mass, muscle strength or function is not routinely assessed in mid-life?



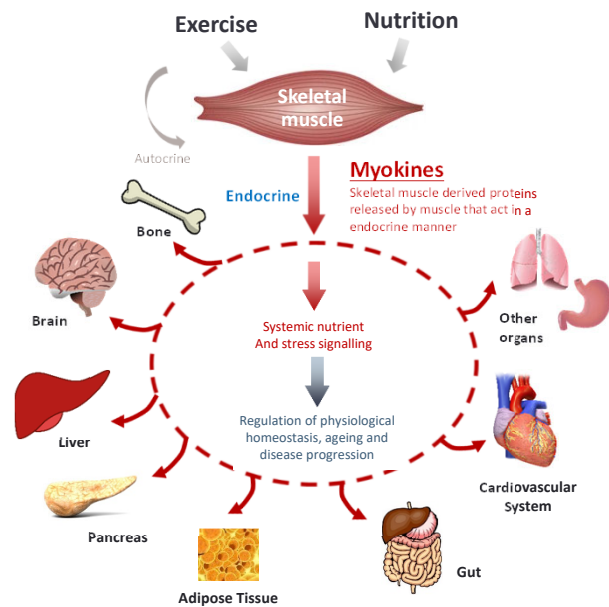
Ferrucci L and Studenski S. Clinical Problems of Aging. In Harrison's Principles of Internal Medicine, 18th Ed. Longo, Fauci, Hauser, Jameson and Loscalzo, McGraw Hill 2011

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Underappreciated Role of Muscle

- Skeletal muscle is a **secretory** organ with endocrine functions.
- Skeletal muscle responds to dietary and exercise challenges (and metabolic dysfunction) by secreting myokines (and other factors) that regulate systemic nutrient and stress signalling.
- **Myokines** mediate the communication of the muscle with other target tissues in an autocrine, paracrine and endocrine manner.

Rai M and Demontis F Ann Rev Physiol 78:85-107, 2016

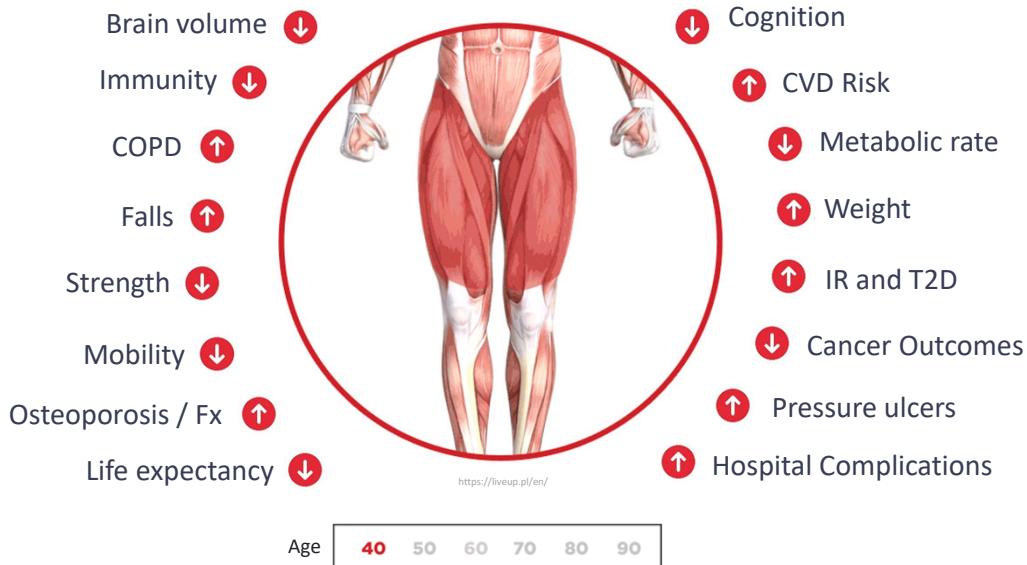


Adapted from Rai M and Demontis F Ann Rev Physiol 78:85-107, 2016

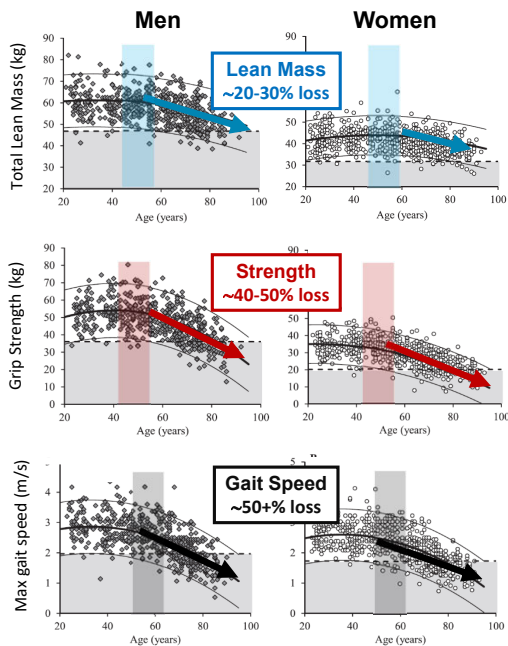
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Consequences of Muscle Loss



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Heterogeneity in the *Timing and Rate of Muscle Loss*

Muscle Mass

Starts ~50-60 years of age | loss of ~20-30% throughout life (~10 kg in men and ~6 kg in women)

Muscle Strength

Starts ~50 years of age | loss of ~40-50% throughout life. Accelerated loss after the age of 70-75.

Physical Performance

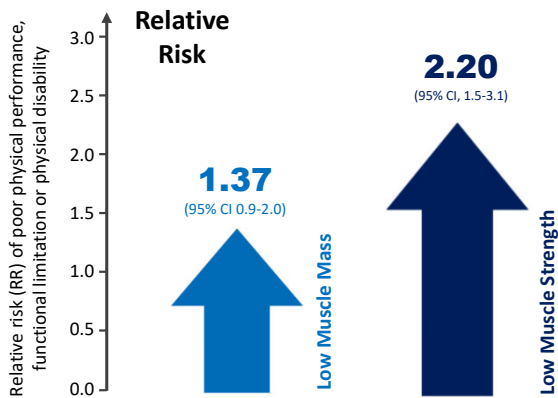
Starts ~55-60 years for men and ~50 years for women; accelerates after 70-75; 45% decrease with age.

Suetta C et al. J Cachexia, Sarcopenia and Muscle 2019

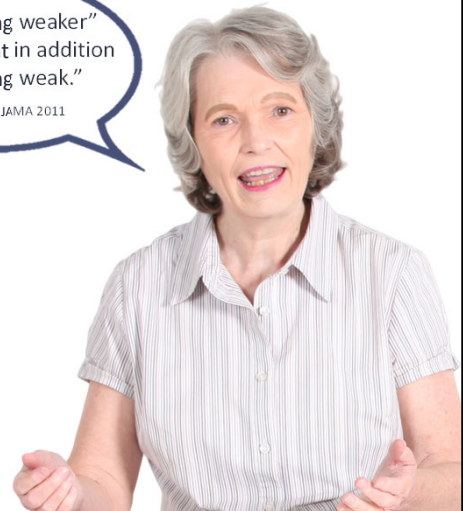
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FACT #2

Low Strength Linked to Disability



“becoming weaker” is important in addition to “being weak.”
Xue et al. JAMA 2011



Risk of Poor Physical Performance or Disability in Adults with Low Muscle Strength or Muscle Mass

Manini TM and Clark DC. J Gerontol A Biol Sci Med Sci 2011

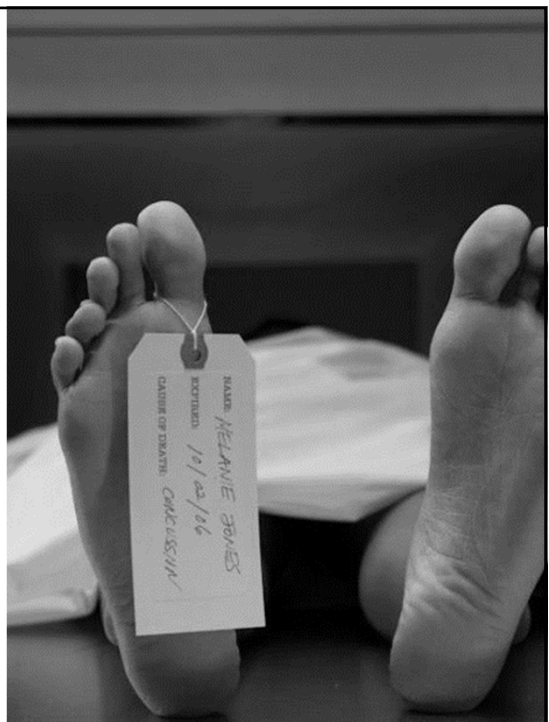
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Strength | Function Linked to Falls and Mortality

Hazard ratios (95% CI) per SD for components of sarcopenia and their association with all-cause mortality and falls-related hospitalization over 9.5 years in 903 older women (80 y)

	HR All-cause mortality ¹	HR Falls-related hospitalization ²
ALM/height ² (per SD)	1.00 (0.89-1.14)	0.94 (0.83-1.06)
ALM/BMI (per SD)	0.95 (0.83-1.07)	1.14 (0.96-1.35)
Grip strength (per SD)	0.83 (0.73-0.93)	0.79 (0.70-0.88)
TUG (per SD)	1.40 (1.29-1.51)	1.31 (1.20-1.42)

ALM = appendicular lean mass; HR = Hazard ratio; ¹ Sim M et al. J Am Med Dir Assoc. 2019 Jan;20(1):76-82; ² Sim M et al. Osteoporosis Int 2019; 30(1): 167-76



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ICD-10-AM
Code (2019)

Sarcopenia: A Clinical Condition

Sarcopenia is a syndrome characterized by progressive and generalized loss of skeletal **muscle mass, strength** and/or **function**.

Low Muscle Strength

- Grip Strength: *Men* < 27 kg; *Women* < 16 kg *or*
- Sit to stand > 15 sec for five rises

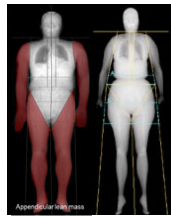
Low Muscle Mass

- Men: ALM < 20 kg; ALM/height² < 7.0 kg/m²
- Women: ALM < 15 kg; ALM/height² < 5.5 kg/m²

* ALM = Appendicular Lean Mass

Impaired Physical Function

- Gait speed < 0.8 m/s (or <1.0 m/s) *or*
- SPPB (Short Physical Performance Battery) * ≤ 8 points



DXA Appendicular lean (muscle) mass



Grip strength

www.youtube.com/watch?v=rBPIEDbLW7lw



Sit to stand

Video © Prof. Robin Daly



Gait Speed (4-6 m walking)

<https://www.youtube.com/watch?v=-JtTxdGFOY>

No universal consensus on the definition/criteria for sarcopenia

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Screening Tool for Sarcopenia

SARC-F: Self-administered simplified screening questionnaire for assessing sarcopenia risk

Component	Question	Scoring
S trength	How much difficulty do you have in lifting and carrying 10 pounds (~4 kg)	0 = None 1 = Some 2 = A lot or unable
A ssistance in walking	How much difficulty do you have walking across a room?	0 = None 1 = Some 2 = A lot, use of aids, or unable
R ising from a chair	How much difficulty do you have transferring from a chair or bed?	0 = None 1 = Some 2 = A lot or unable without help
C limb stairs	How much difficulty do you have climbing a flight of 10 stairs?	0 = None 1 = Some 2 = A lot or unable
F alls	How many times have you fallen in the past year?	0 = None 1 = 1-3 falls 2 = 4 or more falls

SARC-F score of ≥ 4 is predictive of sarcopenia

- **SARC-F** has a high specificity: effective tool for identifying those without sarcopenia.

Bahat G et al. JNHA 22(9):1034-38, 2018; Ida S et al. JAMDA 19:685-89, 2018; Hajaoui et al. JAMDA 20L1178-89, 2019

- **SARC-F + calf girth** improved the sensitivity of SARC-F, but calf girth alone was best screening tool for sarcopenia.

Cut points (moderate/severe):

 **Males <34 cm; <32 cm**

 **Females <33 cm; <31 cm**




Malmstrom TK and Morley JE JAMDA 14:531-32, 2013

Gonzalez et al. AJCN 2021; Mo et al. JAMDA 21(2):288-89, 2020

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Muscle Fibre Loss

Around **40,000 individual muscle fibres** are lost from the quadriceps per year beyond the age of 30 years.

McPhee JS et al. J Gerontol A Biol Sci Med Sci, 2018, 73(10), 1287-1294

Neural Changes

Around **40-50% of motor units** (thus motor neurons) supplying the muscles to the thigh are lost by the age of 70 years


A motor unit consists of the alpha motor neuron and the muscle fibers it innervates.

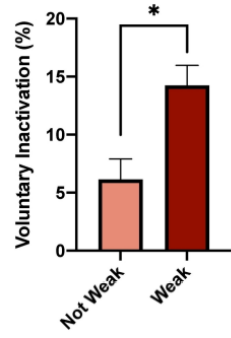
Plasecki et al. J Physiol 594(16): 4525-36, 2016

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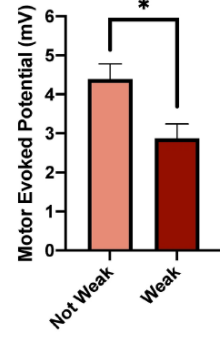
Is There a Failure to Communicate?

Relative contribution of neural excitability and thigh lean mass to clinically meaningful weakness in older adults





Group	Voluntary Inactivation (%)
Not Weak	~6
Weak	~14



Group	Motor Evoked Potential (mV)
Not Weak	~4.5
Weak	~3.0

Weak older adults had 84% lower strength compared to not weak (**Thigh lean mass was similar*).

Weak = decreased ability to fully activate leg extensor muscles.

Weak = decreased corticospinal excitability (↓ neural drive to muscles).

Neural excitability explained ~33% and thigh lean mass ~29% of the variability in strength

Clark et al. J Gerontol A Biol Sci Med Sci 2020; Clark and Carson J Gerontol A Biol Sci Med Sci 2021

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Loss in Muscle Power

Greater than Strength or Mass

Muscle Power (Maximal)

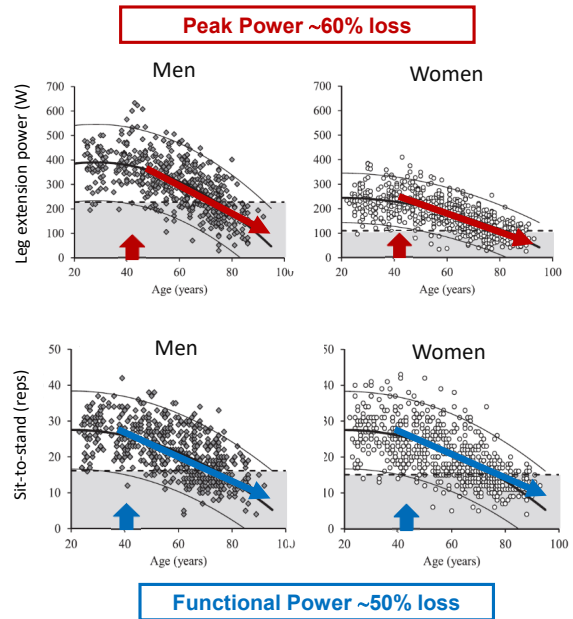
Starts ~40-50 years of age and is greater than rate of loss in muscle mass or strength; ~60% decrease with age.

Preferential loss in type II (fast-twitch) muscle fibres
Movement speed is a key determinant of power loss

Muscle Power (Functional)

Starts from ~45-50 years; no difference between men and women; ~50% decrease with age.

Suetta C et al. J Cachexia, Sarcopenia and Muscle 2019



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FACT #3

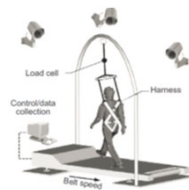
Power More Critical Than Strength

Muscle power more important than strength to prevent slip-related falls

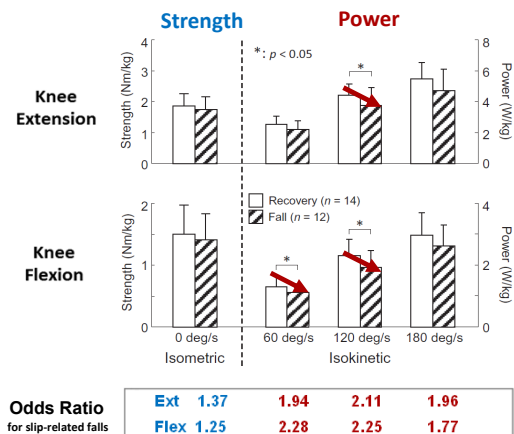
Set-up
26 young adults

Perturbation
while walking
on treadmill

Slip-related
Fall



Video from <https://blog.neura.edu.au/2019/05/exercise-right-week-qa-dr-kim-van-schooten-dr-yoshi-okubo/>



Han and Yang. Hum Movement Sci 44:192-200, 2015

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Resistance Training is Effective

Meta-analyses of RCTs in Older Adults and the Very Elderly (aged ≥ 75 years)



Typically **<10% of older adults participate** in muscle strengthening activities.

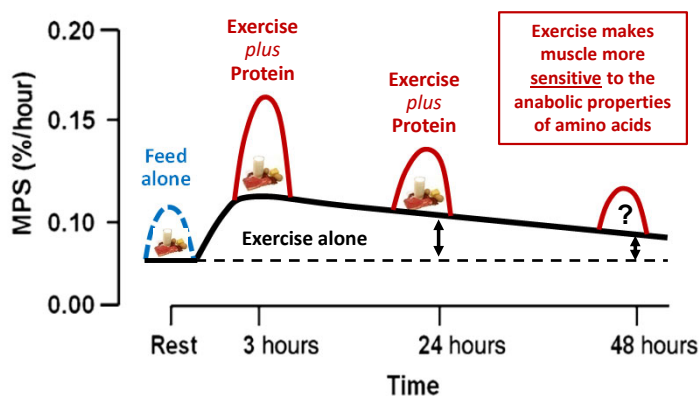
- ✓ Progressive resistance training (PRT) is the most effective modality for improving muscle strength and muscle mass.
- ✓ At least 2 sessions/week, progressive and periodized programs with 2-3 sets of 7-9 reps at moderate/high intensity (70-79% of max).
- ⊗ Heterogeneity with regard to the effects of PRT on physical function.

Borde et al. Sports Med 2015; Peterson et al. Med Sci Sports Exer 2011; Peterson et al Ageing Res Rev 2010; Grgic et al. Sports Med 2020; Kneffel Z et al. J Sports Sci 2021

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Protein and Exercise Interaction

Exercise (PRT) stimulates an increase in MPS that can remain elevated for at least 48 hrs

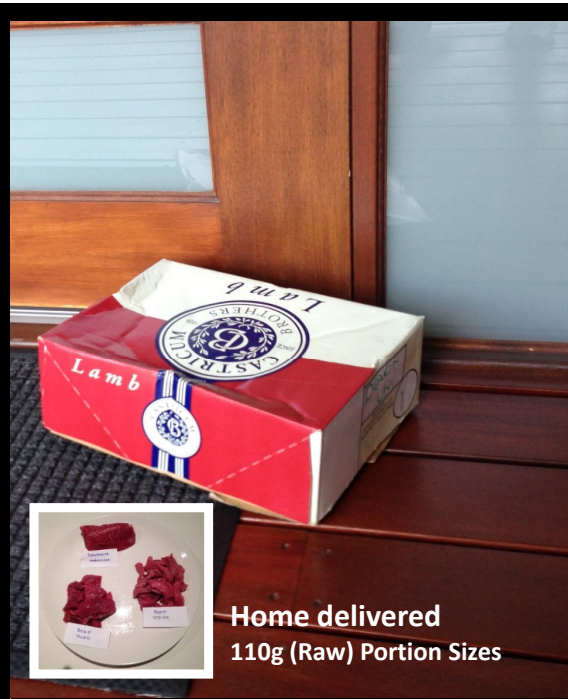


Churchward-Venne TA, et al. Nutr Metab. 2012;9:40; Phillips et al Am J Physiol 1997 273:E99-107.

- When undertaking exercise (resistance training), older adults should consume **$\geq 1.2\text{g/kg}$ body weight/d** of high quality protein.
- At least **20-30g of high quality protein** (e.g. whey or milk protein or lean red meat) post exercise has been shown to effectively stimulate an increase in muscle protein synthesis.

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Exercise + Lean Red Meat

Group	Change relative to baseline (%)
RT+CHO	~0.1
RT+Meat	~1.5

*** p<0.001 vs baseline

High quality Lean Red Meat
160 g cooked (~45 g protein) of lean red meat for 6 days per week

Protein intake increased from ~1.1 to 1.3 g/kg

Group	Change relative to baseline (%)
RT+CHO	~19
RT+Meat	~40

** p<0.01, *** p<0.001 vs baseline

Carbohydrate (pasta, rice, bread)

Protein intake stable at ~1.1 g/kg

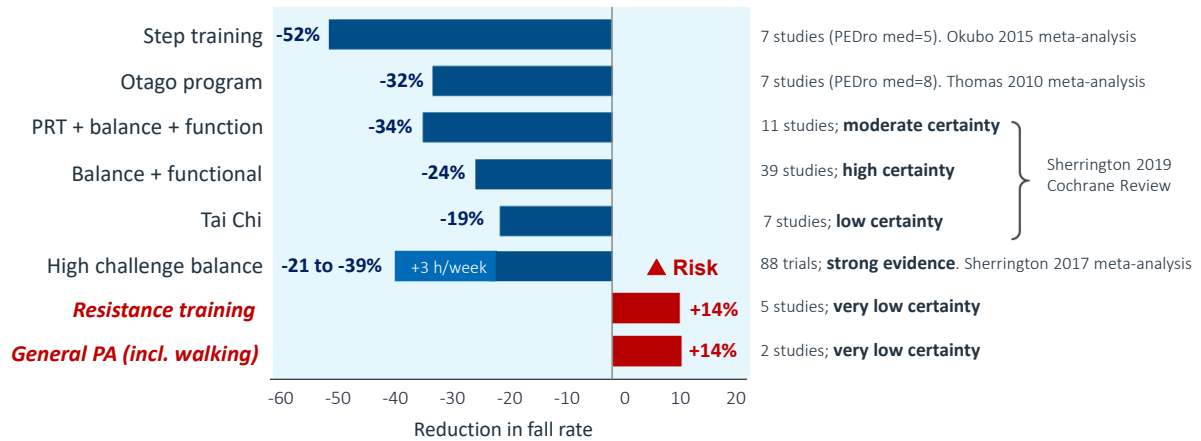
Daly R et al. Am J Clin Nutr 2014;99(4):899-910

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Exercise for Falls Prevention

Fall rate reduction varies by the type of exercise: Cochrane Review

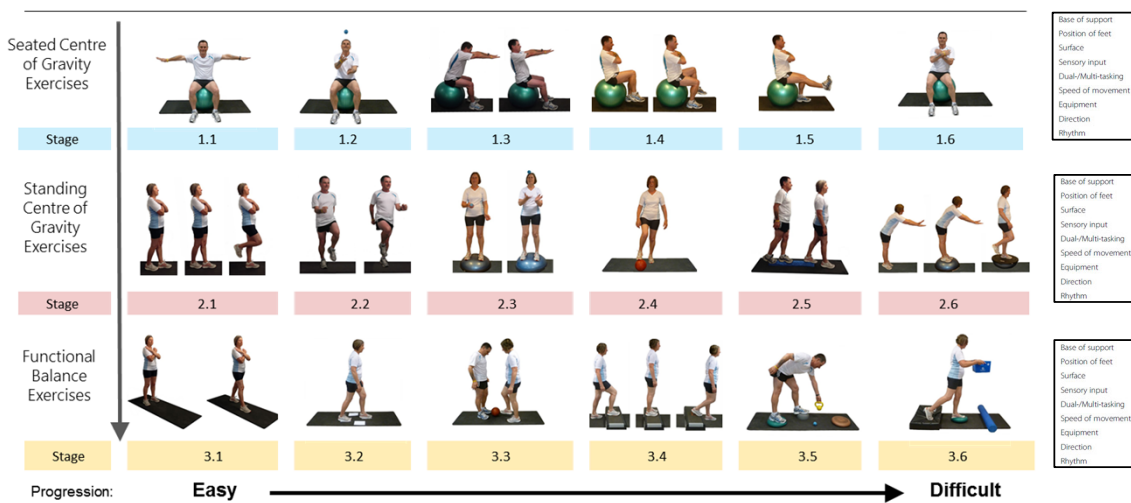


Sherrington C, et al. Cochrane Database of Systematic Reviews 2019; Okubo et al. Br J Sports Med 2017; Thomas et al. Age Ageing 2010; Sherrington et al. Br J Sports Med 2017

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Challenging Balance and Stepping

PACE-IT Trial: Progressively challenging balance / stepping / dual task exercises



Daly et al. Trials 2015 Mar 27;16:120

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Frequent Cause(s) of Falls in Older Adults

1 in 3

community dwelling older adults fall each year
(1 in 5 suffer multiple falls)



Community-dwelling Elderly

Activities associated with the highest proportion of falls:

- **Walking** (61%) at home (61%) during day (88%)
- Displacement from quiet position (13%)
- Gardening (9%) & getting out of bed/chair (5%)

The most frequent cause of falling:

- **Lost balance** (32%)
- **Slipped on an uneven/slippery surface** (29%)
- **Tripped over obstacle** (27%)
- High risk activity (9%)

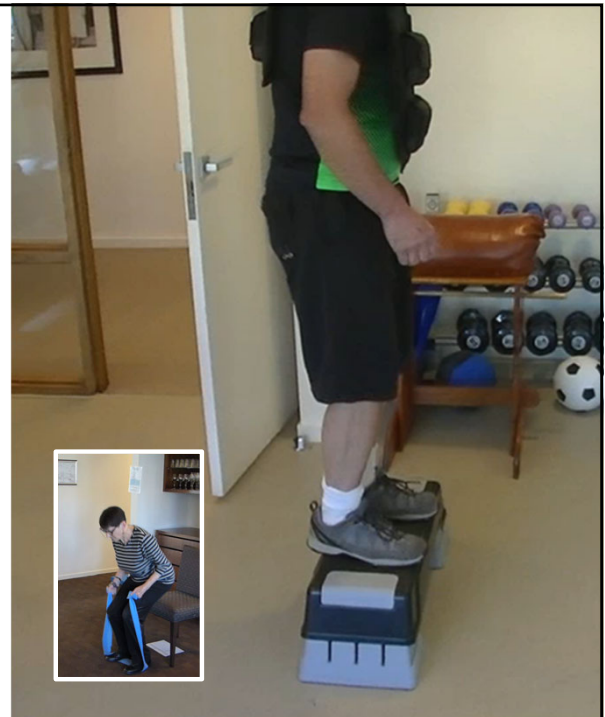
Sanders et al. Osteoporosis Int 2017 Published Online 19 July

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Train to Improve Movement Speed

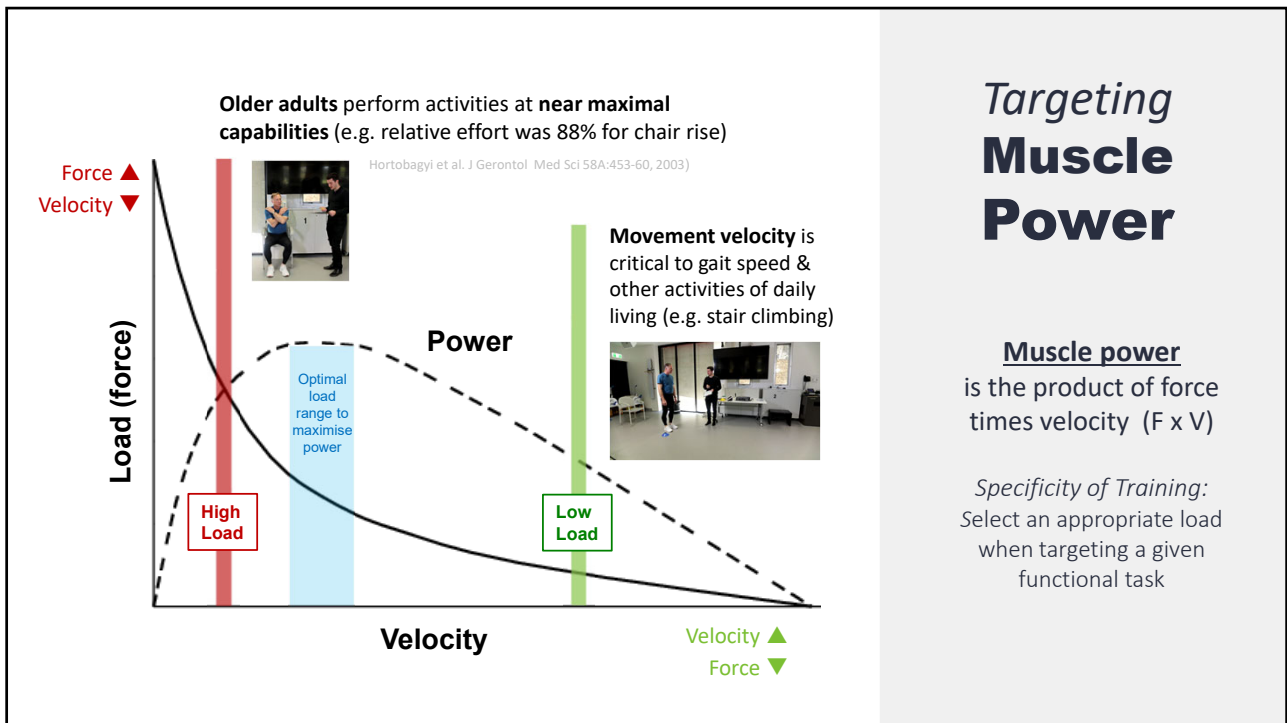
- ✓ **High-velocity (power) training:** perform the lifting phase as rapidly as possible.
- ✓ **Greater benefits** to functional capacity or mobility with HV-PRT training compared to traditional PRT.
- ✓ Meaningful benefits to **'real-life' everyday activities** (e.g. stair climbing ability, car braking speed).

Steib et al. Med Sci Sports Exerc. 2010; 42:902-14; Glenn JM et al Age Ageing 1-6, 2015; Reid et al. J Gerontol A Biol Sci Med Sci 2015

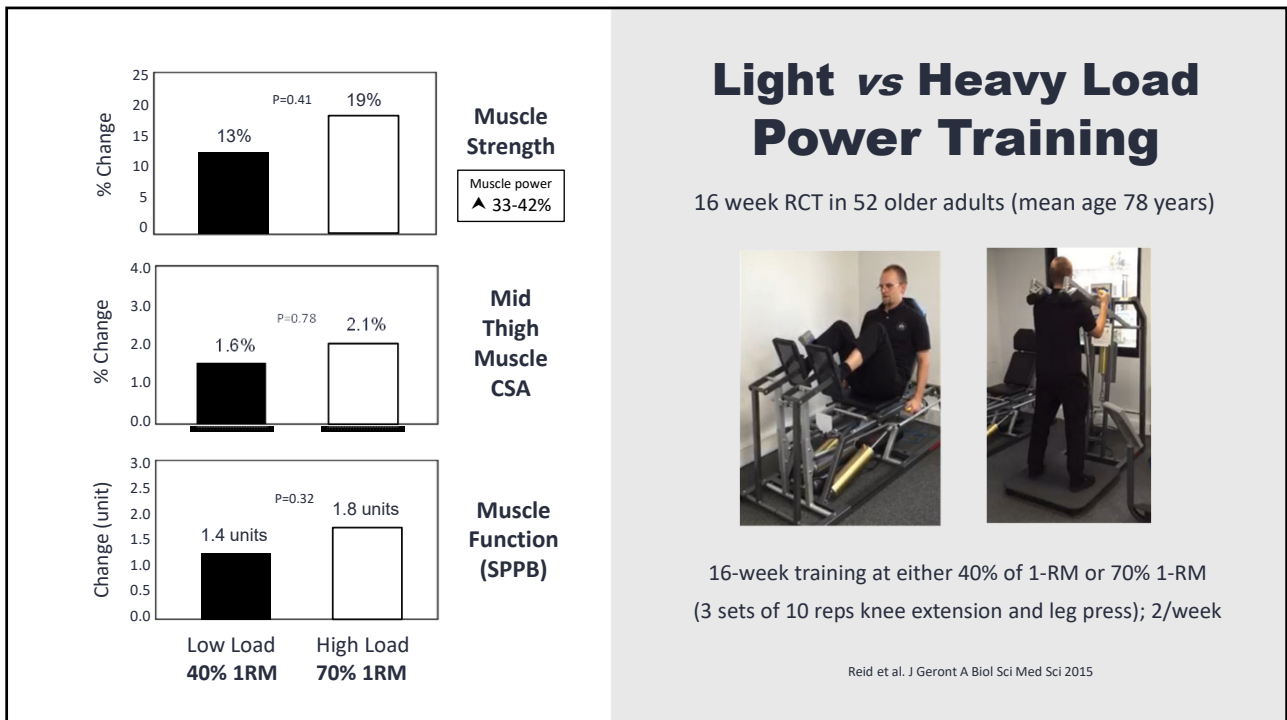


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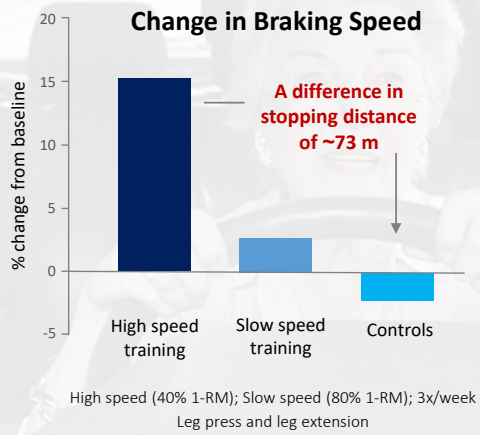
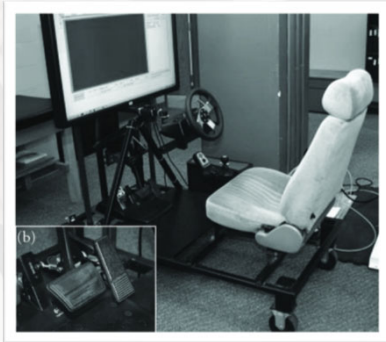
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Improved Braking Speed

12 week RCT in 72 older adults: High speed vs slow speed training

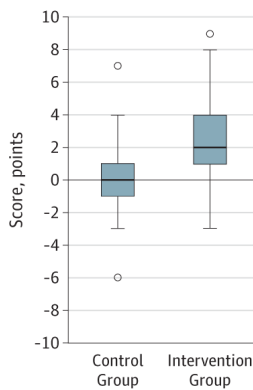


Sayers and Gibson J Aging Res 2012

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Multi-modal Exercise Acute Hospitalization

370 very elderly patients (mean age 87 years); median 5 days



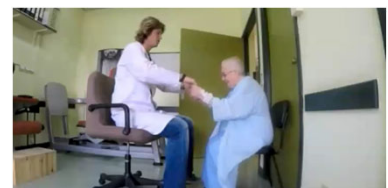
Martinez-Velilla et al. JAMA Int Med 2018

Mean 2.4 point increase in SPPB score
[1.0 point increase is clinically significant]

Exercise Intervention

- 2 daily sessions (morning & evening); 20 min duration (morning session only supervised)
- Resistance training (2-3 sets, 8-10 reps); 3 exercises performed at high speed.
- Challenging balance and gait retraining

No effect on length of hospital stay, falls during hospitalization or 3-month readmission



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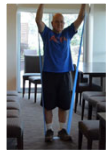
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Dual Task Functional Power Training

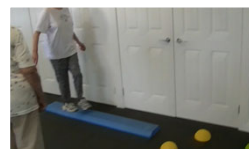
PACE-IT Trial
12-month cluster RCT

Residents (n=300) at fall risk in 22 independent living retirement villages randomised to DT-FPT or usual care control group

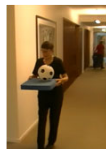
Study protocol available at Daly R et al. *Trials*. 2015 Mar 27;16:120.



Functional power and strength training



Challenging balance, stepping and mobility



Dual-task motor and cognitive exercises

Included dual task cognitive and/or motor activities

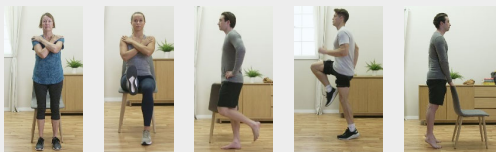
Included dual task cognitive and/or motor activities

Phase 1: 6 months supervised training (45-60 min, 2/week); Phase 2: step-down maintenance program

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Strength | Power Exercise Snacking

Twice-daily, 10-minute resistance 'exercise snacks' for 28 days

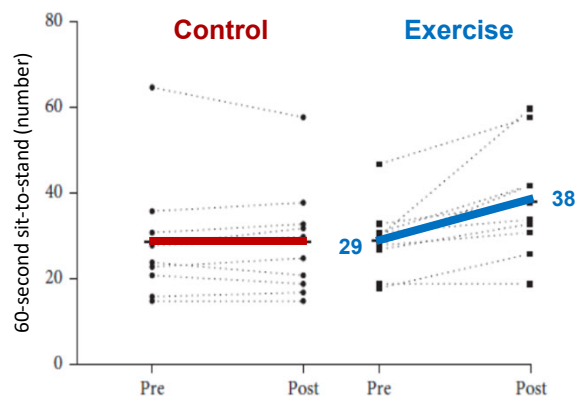


Chair Sit to stand Seated leg extension Standing leg curls Marching on the spot Standing Calf raises

All 5 exercises performed continuously for 60-seconds (60 sec recovery between exercises)

Mean adherence 98%; no adverse events

Healthy older adults aged 65-80 years
28-day unsupervised, home-based exercise program



Perkin et al. *J Ageing Res* 2019

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Take Home Message



Recognize sarcopenia ('muscle failure') as a disease

- Assessment of muscle mass, strength and function provides a wealth of information about health and disease risk. Include them in your assessment battery



Targeted and personalized programs

- Muscle loss is heterogenous; speed and power are critical for older people
- Program must be targeted, progressive and functionally relevant to everyday activities



Multifactorial interventions with exercise are needed

- Falls and fracture (and most chronic diseases) are multifactorial in nature and so a multifaceted approach is needed targeting specific areas of concern/weakness.

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