Human Research Program Informed Consent Briefing

Lori Ploutz-Snyder, Ph.D.



# Integrated Resistance and Aerobic Training Study- Sprint



#### **Exercise Physiology Laboratory/SK**

Lori Ploutz-Snyder, Ph.D. Alan Moore, Ph.D. Jeffrey Ryder, Ph.D. Meg Everett, M.S.

**Neuroscience Laboratory/SK** Jacob Bloomberg, Ph.D.

**Bone Laboratory/SK** Jean Sibonga, Ph.D. Linda Shackelford, MD **Cardiovascular Laboratory/SK** Steven Platts, Ph.D. David Martin, B.A.

**Research Statistician/SK** Robert Ploutz-Snyder, Ph.D.

ASCR/SD Mark Guilliams, M.A., CSCS

University Partners Scott Trappe, Ph.D. – Ball State Univ Todd Trappe, Ph.D - Ball State Univ Wendy Kohrt, Ph.D. – Univ of Colorado Ed Coyle, Ph.D. – Univ of Texas

## Background

Space flight causes reductions in fitness/health:

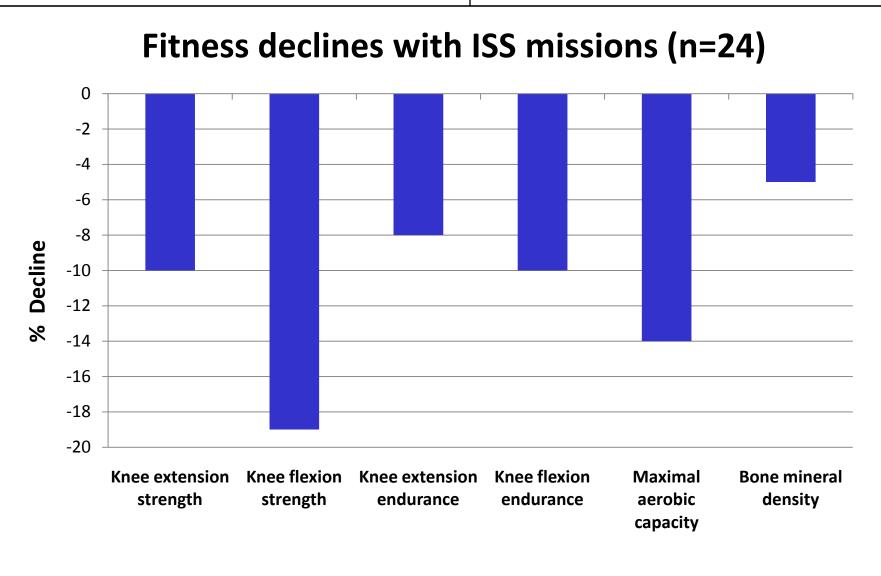
- •Cardiovascular reduced VO<sub>2</sub>max, cardiac output
- •Bone reduced bone mineral density
- •Muscle reduced mass, strength and endurance

Exercise is the primary countermeasure to protect against these changes and was made operational before completely mature; research continues to identify most effective/efficient exercise programs.

Crew medical tests (cardio, muscle, bone) do not yield sufficient information to fine tune the effectiveness of exercise programs, thus there is a need for more detailed testing aimed at identifying the most effective training program.

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# Background

- NASA has spent millions of dollars over past decade to fund new exercise equipment and research on exercise effectiveness.
- Ground research clearly shows that intensity is the most important factor related to maintenance of fitness
  - Duration and frequency can be dramatically reduced only if intensity is kept high
- ARED and T2 allow for more variety and higher intensities



## Background

- June and October 2008 workshops identified the need for an optimized exercise prescription.
  - ASCR, ExPC, HRP management, flight surgeons, medical operations, external experts in muscle, bone and cardiovascular function.
- Major recommendations
  - Higher intensity, less frequent resistance exercise
  - More variety of resistance exercises
  - Alternate days of moderate intensity continuous aerobic exercise with higher intensity interval aerobic exercise
  - Monitor in-flight exercise performance using instrumented hardware
  - Include more robust physiological outcome measurements to document the efficacy of the exercise program.

# Objective

- Obtain detailed information about crew physical fitness pre- and post-flight.
  - Participation involves pre- and post-flight testing (modifications to standard medical & new tests).
- Evaluate new evidence based exercise prescription with higher intensity, lower duration and frequency.

# **Participation Options**

- Active subject Full participation of all pre-, in-, and postflight data collection and in-flight exercise program
- Control subject Pre- and post-flight testing only
- Data Sharing subject Agree to share medical pre, in, and post bone and exercise data
- Muscle biopsy

# In-flight exercise program

- Higher intensity, lower frequency and duration
- Muscle
  - Bedrest and unloading studies show 2-3 days/week of training is sufficient if the contractions during resistance exercise are maximal or nearly maximal
  - Aerobic intervals optimal for affecting muscle aerobic metabolism/endurance
- Bone
  - Evidence suggests multiple daily sessions required
  - High magnitude and rate of strain
  - Site specificity
- Cardiovascular
  - Need high intensity, best achieved with intervals



## Integration of Resistance and Aerobic

	Day 1	Day 2	Day 3	Day 4	Day 5	Day 6	Day 7
Resistance	35-60 min		35-60 min		35-60 min		
Aerobic Interval		32 min		15 min		35 min	
Aerobic Continuous	30 min		30 min		30 min		

Note: Time savings up to 3 hours/week compared to current exercise time At least 4 hrs, preferably 8 hrs separating exercise sessions

## **Sprint Aerobic Intervals**

- Short Sprint 10 minute warm up at 50% of HRmax, followed by 7-8 sets of near-maximal exercise for 30 seconds, followed by 15 seconds rest. Increase load after 9 sets (Burgomaster et al., 2008; Gibala & McGee, 2008; Gibala et al., 2008; Tabata et al., 1996)
  - Increases mitochondrial function, peripheral cardiovascular adaptations such as muscle enzymes and capillary density.
- 2 minute 5 minute warm up at 50% VO<sub>2</sub>max, followed by 6x2 minute stages at 70, 80, 90, 100, 90%, 80% VO<sub>2</sub>max. The first 5 stages are separated by 2 minute active rest stages at 50% VO<sub>2</sub> max. The final stage is a 5 min active rest at 40% VO<sub>2</sub>max. (Greenleaf et al., 1989)
  - Maintain maximal aerobic capacity, similar protocol well tolerated on ISS
- 4 minute 5 minute warm up at ~50% HRmax, followed by intervals of exercise at 90% HRmax. The exercise intervals will be 4x4 min bouts, with 3 min active rest periods. (Helgerud et al 2007)
  - Cardiac adaptations, cardiac output and stroke volume

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### **Sprint Resistance Exercise**

	Day 1	Day 2	Day 3
	Squat, Bench Press, Romanian Dead Lift, Upright Row, Heel Raise	Dead lift, Shoulder Press, Bent-over Row, Single Leg Squat, Heel Raise	Front Squat, Bent-over Row, Dead lift, Bench Press, Heel Raise
Week			
1	Light	Light	Light
2	Light	Light	Light
3	Moderate	Light	Heavy
4	Heavy	Moderate	Light
5	Light	Heavy	Moderate
6	Moderate	Light	Heavy
7	Heavy	Moderate	Light
8	Light	Heavy	Moderate
9	Moderate	Light	Heavy
10	Heavy	Moderate	Light
11	Light	Heavy	Moderate
12	Moderate	Light	Heavy

### **Sprint Resistance Exercise – session detail**

	Weel	ks 1-6	
	Light	Moderate	Heavy
	3	3	3
Reps	12	8	5
Rest (sec)	90	120	120
Total time (min)	35	40	40
	Week	s 7-12	
	Light	Moderate	Heavy
Sets	3	3	3
Reps	10	6	3
Rest (sec)	90	150	180
Total time (min)	35	60	40

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## **Exercise Logs**

- Complete in-flight exercise logs outlining the exercise performed
  - Aerobic
  - Treadmill & cycle speed, load, duration, HR
  - Resistance
    - Exercise, load, reps, sets & rest between sets



## In-flight data collection

- Muscle strength
  - Monitor training loads and adjust prescription as needed
- VO<sub>2</sub>max every 30 days
- Muscle size with ultrasound every 30 days

• Prescription can be adjusted accordingly based on in-flight measurements

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# Pre and post-flight data collection

- Muscle
  - Muscle Function
  - Imaging
  - Muscle Biopsy
- Cardiovascular
  - Pre-, in-, post-flight VO<sub>2</sub>max
  - Ventilatory threshold
  - Cardiac ultrasound
- Bone
  - DEXA, qCT

### **Testing – Muscle Function**

•Leg Press

- **Maximal Isometric Force**: Push against fixed footplate.
- •Power/Endurance: Push weight away as fast as possible (40% max. force, 21 repetitions). Leg extension push only; weight caught by a braking system.

### •Knee Extension

- Force Control: Match leg force with a reference force displayed on a computer screen during isometric leg extension (5% maximal force).
- **Neuromuscular Drive**: Brief, electrical muscle stimulus provided to thigh muscle during isometric leg extension.

16

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### Bench Press

- Maximal Isometric Force: Push against fixed bar.
- **Power/Endurance:** Push weight away as fast as possible (30% max force, 21 repetitions) Arm extension only; weight caught by a braking system.
- Muscle Force Control: Match isometric arm force with a reference force displayed on a computer screen (5% maximal force).



## Testing – mods to standard medical

- Cycle Test MEDB 4.1
- VO<sub>2</sub>max test for ventilatory threshold (L-270, L-80, L-50; R+1-3, 8-10, 30)
  - Peak: 3 min warmup, then 1 min25 W increments to max
  - Allows for evaluation of ventilatory threshold which has never before been assessed with spaceflight but is more functionally important than VO<sub>2</sub>max

## **Bone – density and architecture**

- Bone densitometry MEDB 1.11
  - DEXA scan, L-<365,</li>
     R+30, R+180, then
     yearly

- Add qCT for bone density and geometry of hip and spine
  - L-<365 and R+30</p>

# Imaging- MRI for muscle size

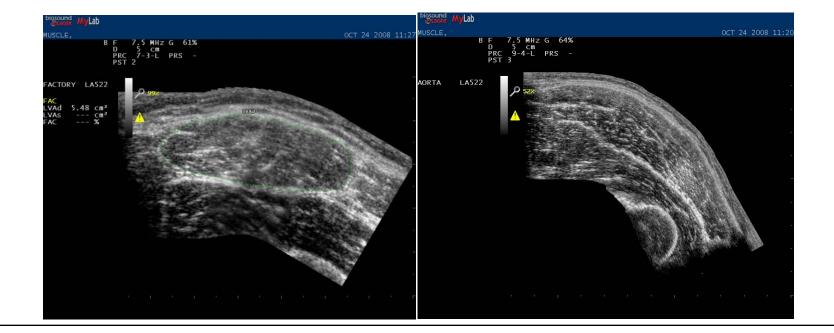
- MRI of the legs for muscle size/volume of thigh and calf (L-80, L-50; R+ 1-3)
  - 30 min supine rest
  - 8 min scan of calf
  - 8 min scan of thigh





## Imaging- Ultrasound

- Ultrasound of the leg muscles (L-80, 50; R+0-1)
- Cardiac ultrasound

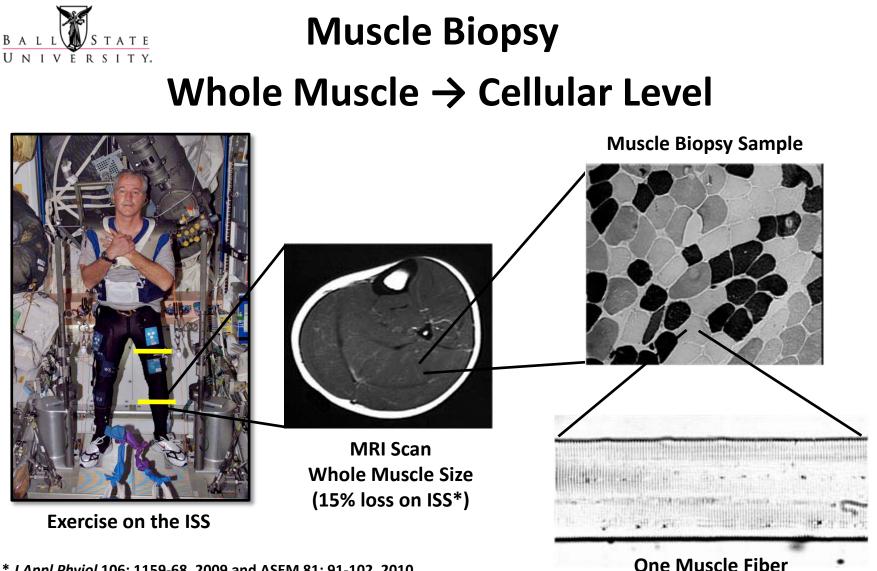


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Sprint		Lori Ploutz-Snyder, Ph.D.		
	Pre-flight		Post-flight	
	L<365: DXA*, QCT†		R+0/1: Ultrasound, Muscle Biopsy	
	L-270: Peak Cycle (VO2max)*		R+1: Muscle Performance <sup>+</sup> , MRI, VO2max**	
	L-180: Isokinetic Testing*, Muscle Performance†		R+5-7: Isokinetic Testing*	
	L-80: Isokinetic Testing*, Muscle Perform MRI, Ultrasound, VO2max**	ance†,	R+6: Muscle Performance <sup>+</sup>	
<ul> <li>* Nominal Medical testing requirement</li> </ul>	* Nominal Medical Isokinetic Testing, Muscle Performance <sup>+</sup> , V		R+8-10: VO2max <sup>+</sup>	
<ul> <li>** Testing replaces</li> <li>existing nominal</li> <li>medical testing</li> </ul>	existing nominal Muscle Biopsy (optional) medical testing		R+30: Isokinetic Testing*, Muscle Performance†, VO2max**	
requirement (either Functional Fit or Submax)			<r+30: DXA*, QCT+</r+30: 	
<ul> <li>Possible Data Share with existing experiments</li> </ul>	6.25 hrs shared medical 10 hrs study specific and possible experiment data sharing		6 hrs shared medical 6.5 hrs study specific and possible experiment data sharing	

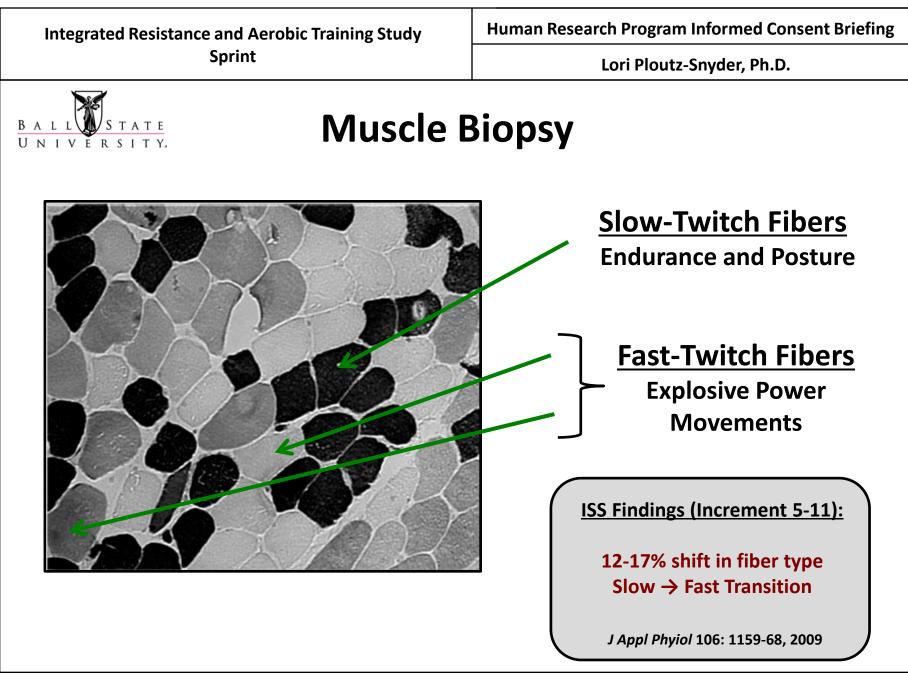
ntegrated Resistance and Aerobic Training Study	Human Research Program Informed Consent Brie Lori Ploutz-Snyder, Ph.D.		
Sprint			
Inflight Medical Requirements/Activities	Inflight Sprint		
<ul> <li>Periodic Fitness Evaluation (PFE):</li> <li>FD 14 and about every 30 days after (6 sessions/1.5 hrs per session)</li> <li>9 hrs total</li> </ul>	<ul> <li>VO2max (replaces PFE):</li> <li>FD 14 and about every 30 days after (6 sessions/3.6 hrs per session)</li> <li>21.5 hrs total</li> </ul>		
	<ul> <li>Muscle ultrasound:</li> <li>FD 14, 30 and about every 30 days after (7 sessions/1.75 hrs per session)</li> <li>12.25 hrs total</li> </ul>		
<ul><li>Throughout mission:</li><li>2.5 hrs per day of exercise training</li></ul>	<ul><li>Throughout mission:</li><li>15-90 min per day of exercise training*</li></ul>		
Total testing time (PFE) = 9 hrs	Total testing time = 33.75 hr		

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\* J Appl Phyiol 106: 1159-68, 2009 and ASEM 81: 91-102, 2010

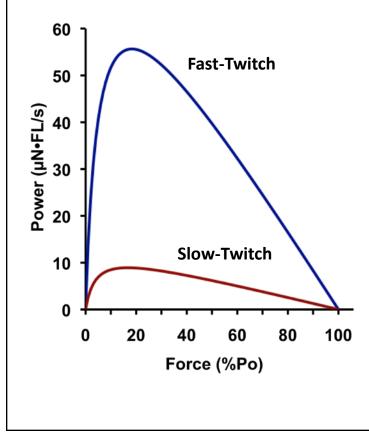


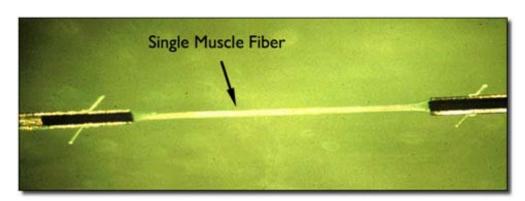
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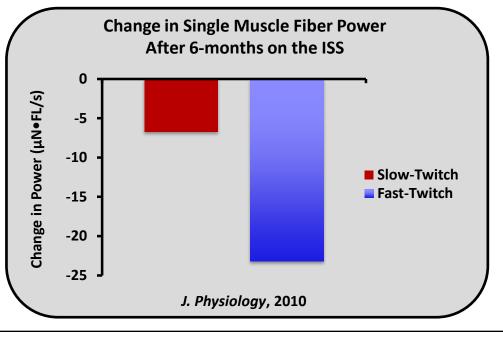


## **Muscle Biopsy**











**Thigh – Vastus Lateralis** 

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Muscle Biopsy Pre: L-50-55 Post: R+0



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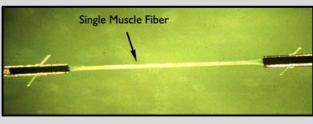
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## **Muscle Biopsy – Science**

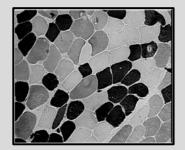
### **Function & Structure**

#### **1. Single Muscle Fiber Function**



To determine size, strength, speed and power in slow-and fast-twitch muscle fibers which impacts muscle performance

#### 2. Muscle Fiber Type Transformations



To determine changes in fiber type which impacts muscle performance, metabolic health, fuel use, and fatigue

### **Metabolic Capacity**

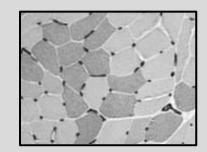
#### 3. Aerobic and Anaerobic Enzymes



**Muscle Biochemistry to** determine metabolic profile which impacts energy transfer and metabolic health

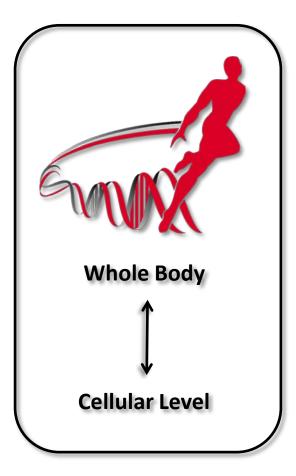
**Mitochondria** 

#### 4. Capillary Density



To determine capillary network which impacts blood flow, oxygen, and nutrition delivery

## **Muscle Biopsy – Science Application**



1. The cellular data will complement the whole muscle assessment to provide a detailed profile of skeletal muscle health and the effectiveness of the current ISS hardware and new exercise prescription.

2. Our recently published ISS muscle biopsy research from Increment 5-11 in combination with new whole muscle and cellular information will provide a strong scientific platform to help guide future countermeasure programs.



Recent ISS muscle publications: 1. Trappe et al. JAP 106: 1159-68, 2009 and 2. Fitts et al. J. Physiol. 2010.

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## **Muscle Biopsy Team Experience**



Space Flight: Shuttle & ISS 14 crewmembers 56 muscle biopsies total

Bed Rest: 17, 60 and 90-d

Aging: up to 93 years old

### Athletes: Recreational, Competitive, and Olympic

Drs. Scott and Todd Trappe have performed >3,000 muscle biopsies (Human Performance Laboratory > 10,000)



R+0

biopsy in Star City



## **Experiment Training**

- L-18-12 months
  - 1 hr overview familiarization
- L-12-6 months
  - 2 hr PPFS
  - 1.5 hrs VO2max nominal operations
  - 1 hr ultrasound
- L-18 months (as soon after consent as possible)
  - Sprint exercises during scheduled gym time

## **Possible Risks or Discomforts**

- Reasonable risk
- Imaging
  - MRI personal injury from magnetic objects, noise, claustrophobia
  - qCT radiation exposure
  - Ultrasound gel
  - Protections screen for metal implants, earplugs, feet first scanning, monitor crew radiation exposure, ask about allergy to ultrasound gel.
- Muscle biopsy
  - Discomfort, bleeding, infection, scarring of skin
- Protections use of standard sterile procedures, experienced investigator to obtain biopsy.

## **Possible Risks or Discomforts**

- Exercise testing typical risks associated with exercise
  - Muscle soreness, cramping, joint injury, strains, sprains, cardiovascular events, light-headedness after exercise, discomfort
  - Protections warm up, supervision, concentric only contractions, subjects already screened for heart disease, cool down

# **Participation Options**

- Active subject Full participation of all pre-, in-, and post-flight data collection and in-flight exercise program
- Control subject Pre- and post-flight testing only
- Muscle biopsy