



# Failure

## Out patient care

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# Present

1960-1980: Bed Rest; No exercise!

1980-1990: Exercise training in LV dysfunction

***2003: HF-Action***

***RCT Exercise study in HF [LVEF $\leq$ 35%]; 3000pts***

***Exercise Training (> 1x/wk)***

***at moderate to vigorous exercise***

***Mortality, Morbidity, Mechanisms***

***Long term FU: 4 years***

# All Guidelines

## Exercise based cardiac rehabilitation in chronic heart failure

Australia

People with chronic heart failure often present to their general practitioner with questions about their participation in cardiac rehabilitation programs. This article outlines the risk and benefits of such programs.

Working Group on Cardiac Rehabilitation & Exercise Physiology and Working Group on Heart Failure of the European Society of Cardiology

### AHA Scientific Statement

#### Exercise and Heart Failure

A Statement From the American Heart Association Committee on Exercise, Rehabilitation, and Prevention

# ACC/AHA Practice Guideline

## ACC/AHA 2005 Guideline Update for the Diagnosis and Management of Chronic Heart Failure in the Adult - Summary Article

A Report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Writing Committee to Update the 2001 Guidelines for the Evaluation and Management of Heart Failure)

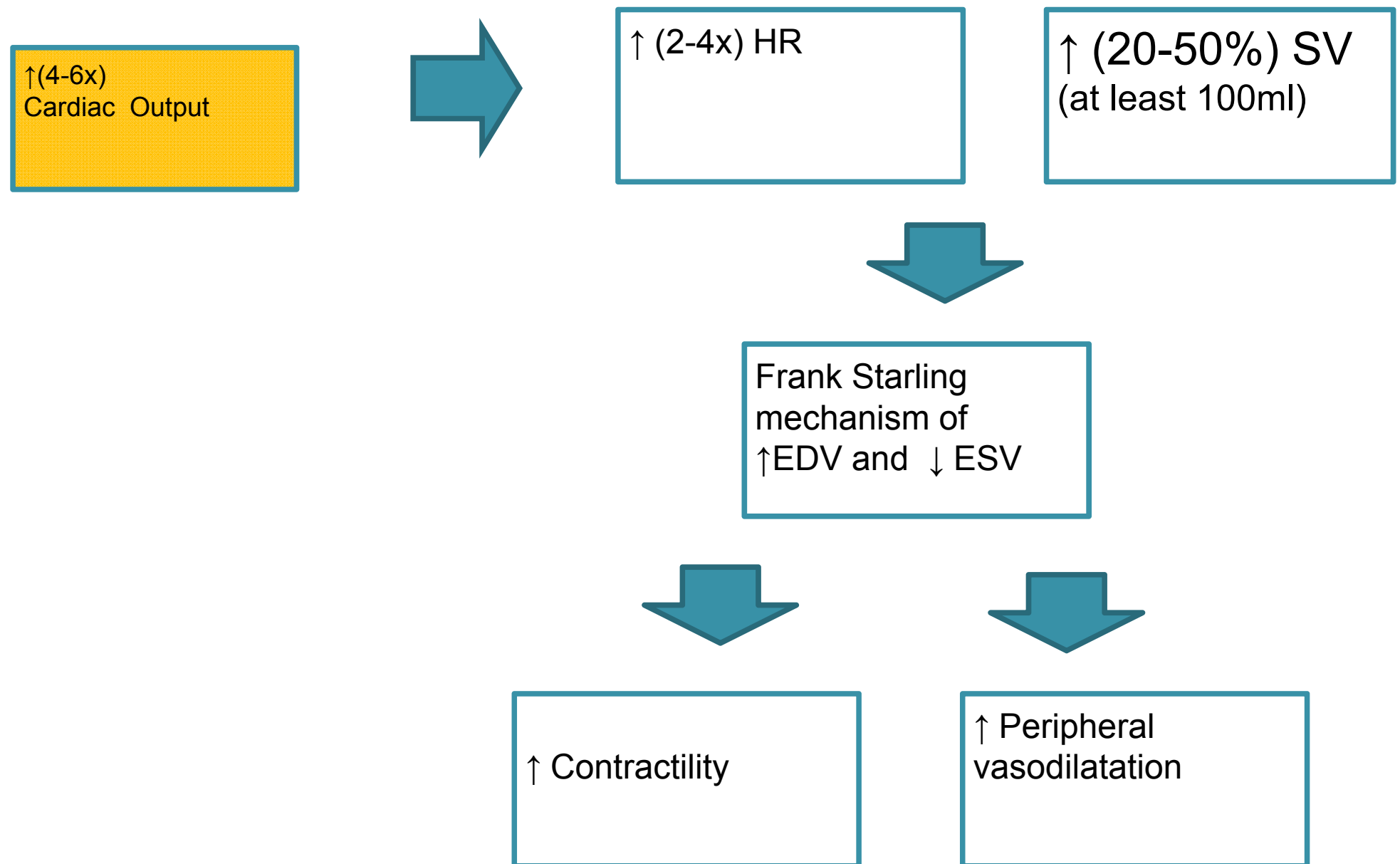
Stage C: Symptomatic HF  
Exercise Training is Class I  
Recommendation

# Chronic Heart Failure

Dyspnea & Fatigue



Exercise intolerance



< 50% of normals ↑  
Cardiac Output



↑ lower max  
HR (low workload)

↑ SV  
(limited to 50-65 mL)



Minimal preload  
reserve to ↑EDV and  
/or inability to ↓ ESV



↓ Contractility, ↓  $\beta$ -  
adrenergic  
responsiveness



↑ systemic vascular  
resistance

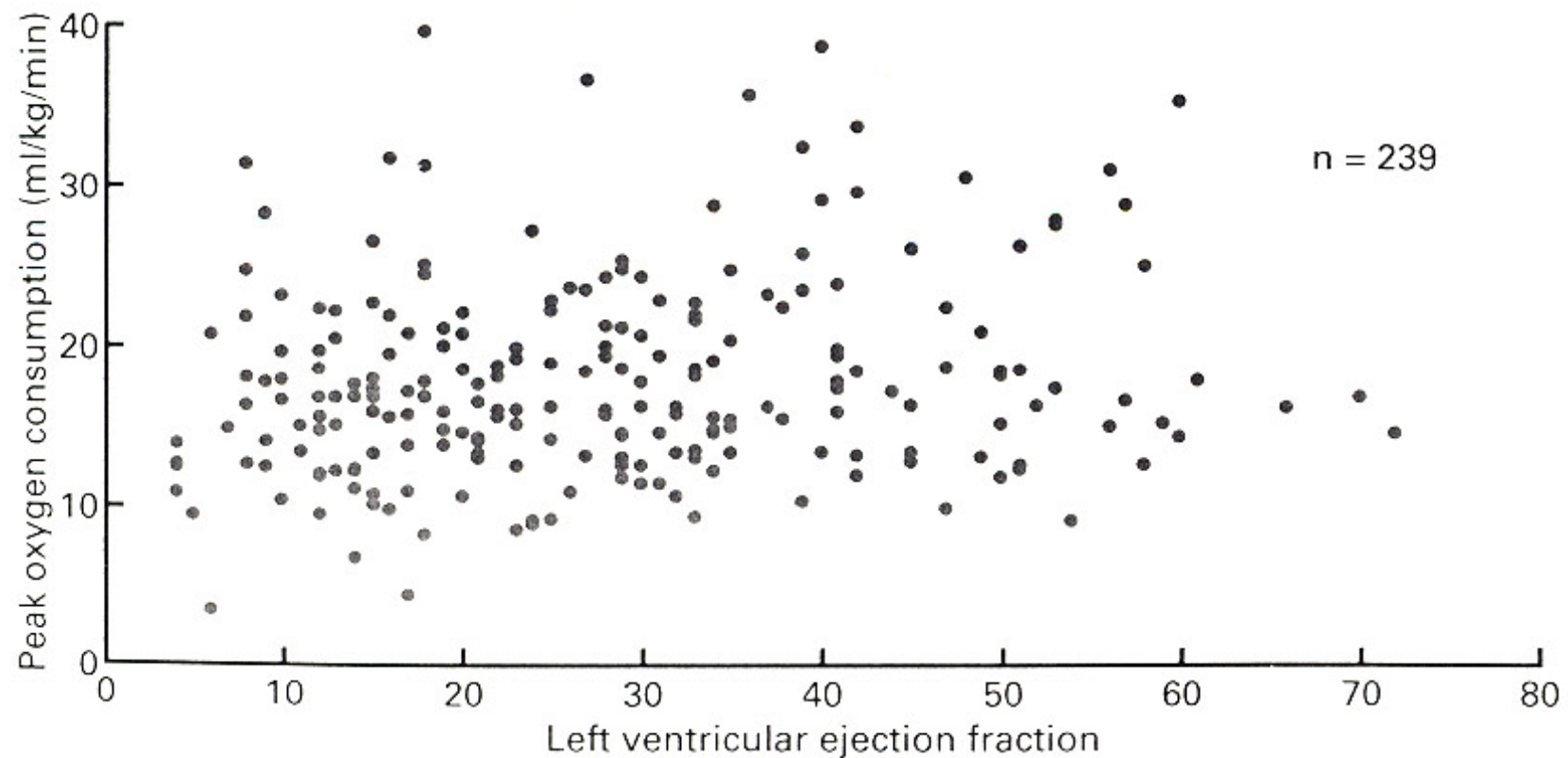


↑ Sympathetic &  
renin-angiotensin  
systems



↓ Arterial vasodilation  
response to exercise

# CHF



Lack of correlation between resting LVEF



# HF

- Abnormal skeletal muscle metabolism
- Skeletal muscle atrophy
- Decreased skeletal muscle endurance
- Type II skeletal muscle atrophy and decreased oxidative enzyme concentration
- Decreased mitochondrial volume
- Reduced respiratory muscle strength
- Reduced respiratory muscle endurance



# **BENEFITS OF EXERCIS TRAINING IN HEART FAILURE**

# Efficacy and Safety of Exercise Training in Patients With Chronic Heart Failure

## HF-ACTION Randomized Controlled Trial

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David J. Whellan, MD, MHS

Kerry L. Lee, PhD

Steven J. Keteyian, PhD

Lawton S. Cooper, MD, MPH

Stephen J. Ellis, PhD

Eric S. Leifer, PhD

William E. Kraus, MD

**Context** Guidelines recommend that exercise training be considered for medically stable outpatients with heart failure. Previous studies have not had adequate statistical power to measure the effects of exercise training on clinical outcomes.

**Objective** To test the efficacy and safety of exercise training among patients with heart failure.

**Design, Setting, and Patients** Multicenter, randomized controlled trial of 2331 medically stable outpatients with heart failure and reduced ejection fraction. Participants in Heart Failure: A Controlled Trial Investigating Outcomes of Exercise Training (HF-ACTION) were randomized from April 2003 through February 2007 at 82 centers within the United States, Canada, and France; median follow-up was 30 months.

# Summary of Clinical End-point Results from HF-ACTION

End point	Hazard ratio	95%	p
All-cause mortality/hospitalization	0.93	0.84–1.02	0.13
After adjustment	0.89	0.81–0.99	0.03
CV mortality/CV hospitalizations	0.92	0.83–1.03	0.14
After adjustment	0.91	0.82–1.01	0.09
CV mortality/HF hospitalizations	0.87	0.75–1.00	0.06
After adjustment	0.85	0.74–0.99	0.03

# Effects of Exercise Training on Health Status in Patients With Chronic Heart Failure

## F-ACTION Randomized Controlled Trial

Maryn E. Flynn, PhD

Anna L. Piña, MD

David J. Whellan, MD, MHS

John P. DiMarco, MS

**Context** Findings from previous studies of the effects of exercise training on patient-reported health status have been inconsistent.

**Objective** To test the effects of exercise training on health status among patients with heart failure.

**Conclusions** Exercise training conferred modest but statistically significant improvements in self-reported health status compared with usual care without training. Improvements occurred early and persisted over time.

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overall summary scale and key subscales at baseline, every 3 months for 12 months, and annually thereafter for up to 4 years. The KCCQ is scored from 0 to 100, with higher scores corresponding to better health status. Treatment group effects were estimated using linear mixed models according to the intention-to-treat principle.

**Results** Median follow-up was 2.5 years. At 3 months, usual care plus exercise training led to greater improvement in the KCCQ overall summary score (mean, 5.21; 95% confidence interval, 4.42 to 6.00) compared with usual care alone (3.28; 95% confidence interval, 2.48 to 4.09). The additional 1.93-point increase (95% confidence interval, 0.84 to 3.01) in the exercise training group was statistically significant ( $P < .001$ ). After 3 months, there were no further significant changes in KCCQ score for either group ( $P = .85$  for the difference between slopes), resulting in a sustained, greater improvement overall for the exercise group ( $P < .001$ ). Results were similar on the subscales, and no subgroup interactions were detected.

HEART FAILURE IS A SYNDROME characterized by dyspnea and fatigue; however, patients with heart failure often

## Abstract

Background: The large randomized controlled multicentre clinical trial, HF-ACT recently demonstrated that a programme of recommendation of regular exercise training at moderate intensity is safe, improves quality of life, and reduces the combined endpoint of all-cause death and hospitalization in patients with chronic heart failure. However, the size of beneficial effects was modest compared to results published in smaller single studies and meta-analyses. Objective: Base

**Hypothesis: programme comprising interval training at high relative intensity would yield significantly larger effects in terms of left ventricular remodelling compared to moderate continuous exercise training.**

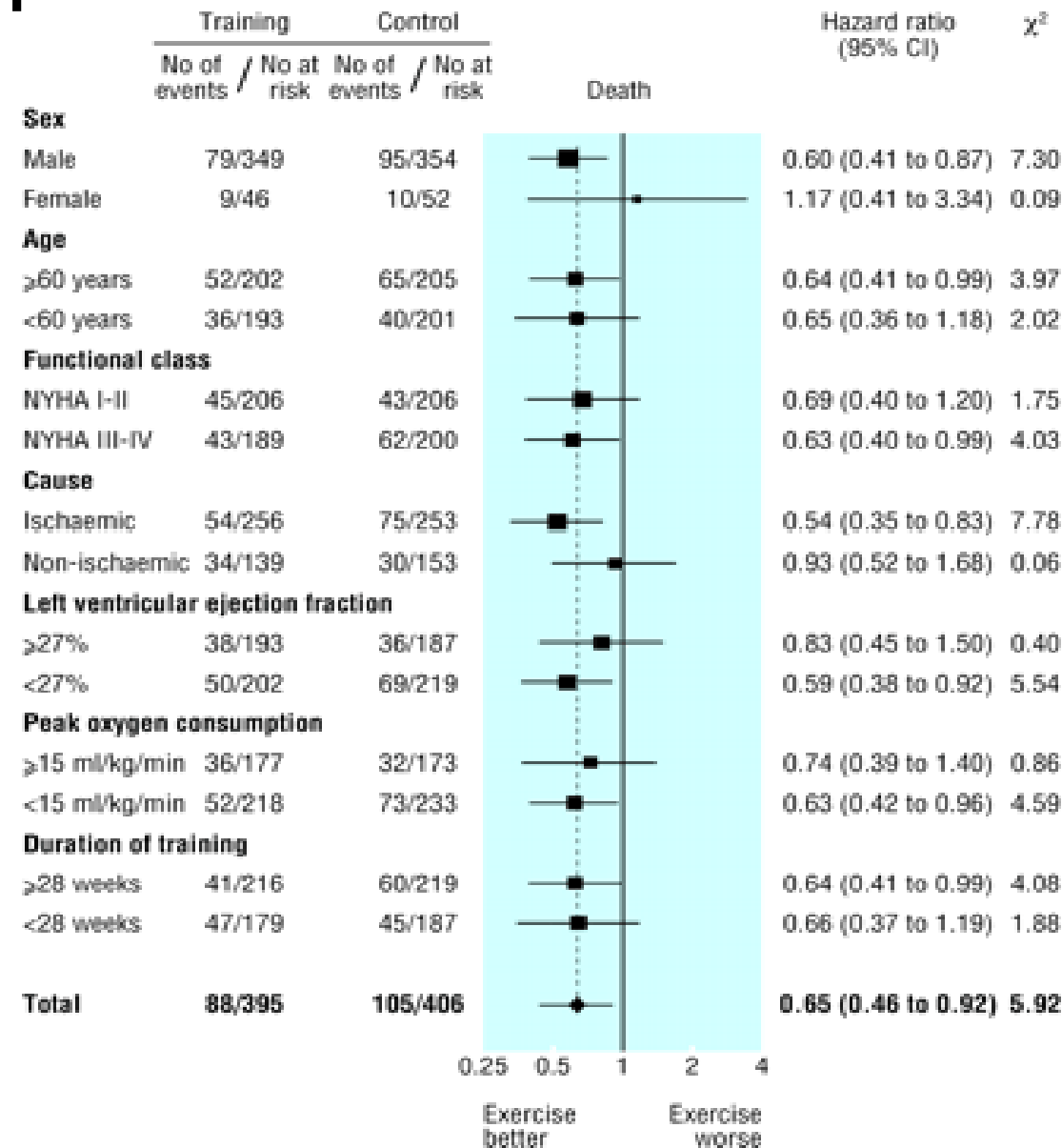
practice at the local centre. The primary endpoint is reverse remodelling, defined as change in left ventricular end-diastolic diameter assessed by echocardiography. Secondary endpoints include peak oxygen uptake ( $\text{VO}_{2\text{peak}}$ ), biomarkers, quality of life, and level of physical activity assessed by questionnaires. In addition, long-term maintenance of effects after the supervised training period will be determined. Assessments will be made at baseline, after the 12-week intervention programme, and at 1-year follow up. A total number of 200 patients on treatment per protocol, randomized to the three groups in a 1 : 1 : 1 manner, is estimated to detect clinically relevant differences in effect with HIT vs. MCT and RE ( $p < 0.05$ ; statistical power 0.90) for the primary endpoint. Inclusion of patients started May 2009 and will

# in patients with HF (ExTraMATCH)

- 9 RCT, 801 patients
- Follow up 2 years

↓ Overall mortality 35%

↓ Admission to the hospital 28%



# HF

- Exercise Capacity
- Myocardial function
- Ventilatory function
- Autonomic Nervous System & neuroendocrine
- Peripheral blood flow
- Skeletal muscle function
- Health-Related Quality of Life



# Exercise based rehabilitation for HF

- 29 RCT; 1,126 HF patients (NYHA I-III)
- 23 aerobic exercise; 6 resistance exercise
- FU 4-60 weeks
- Significant Improvement in
  - $\text{VO}_{2\text{max}}$  [2.16 ml/kg/min]
  - **Exercise duration [2.38 minutes]**
  - **Work capacity [15.1 watts]**

# Clinical & Biological Parameters other than V

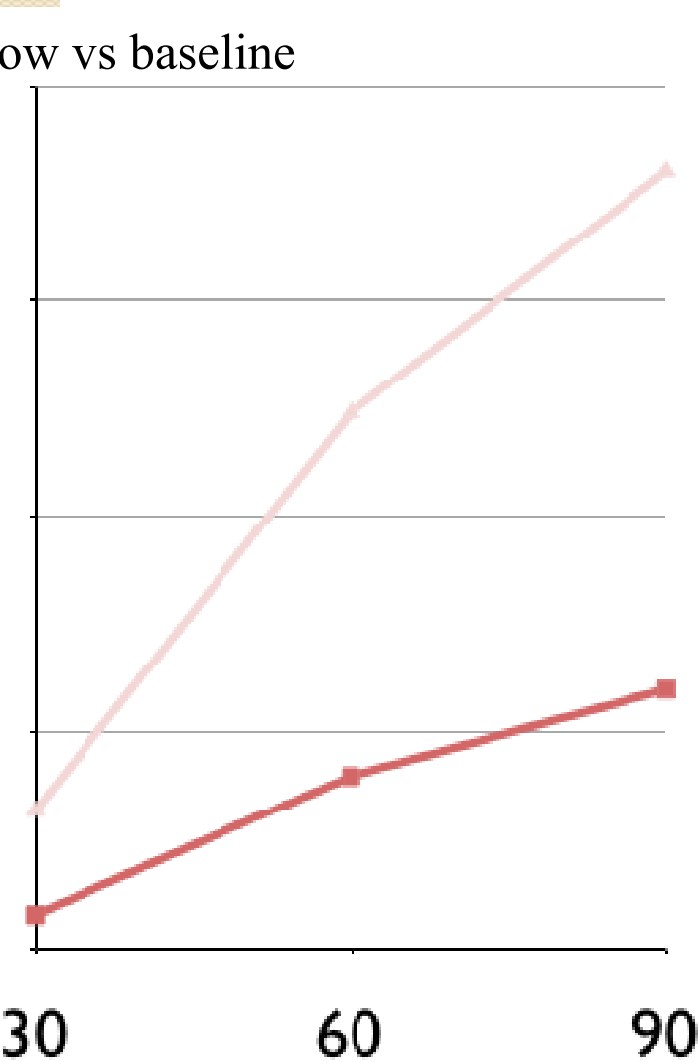
Authors /years	Exercise Program	Improvement
Guori (1999)	Cycle 30 min 3x/wk at 50%-60% $VO_{2max}$ for 8 mo	Plasma NE decreased at rest; no adverse effects on catabolic hormone
ts (1992)	Cycle 20 min 3x/wk at 60%-80% max HR for 8 wk	Improved HRV: decrease in ventilation
ardinelli (1995)	Cycle 40 min 3x/wk at 60% $VO_{2max}$ for 8 wk	Increase muscle mitochondria volume density
mbrecht (1995)	Walk 10 m $VO_{2max}$ in 6 x/d at 70% for 3 wk	Improved ejection fraction and stroke volume
aelli (1996)	Cycle 20 min 5 d/wk for 5 wk	Improved endothelial function
ardinelli (1999)	Cycle 3x/wk at 60% for 8 wk. then 2x/wk for 12 mo.	
alevo (2006)	Strength training 3x/wk for 8 wks	
eyian (1999)	Treadmill, cycle, arm ergometry 33 min 60% 80% MHR 3x/wk for 24 wk	

# function

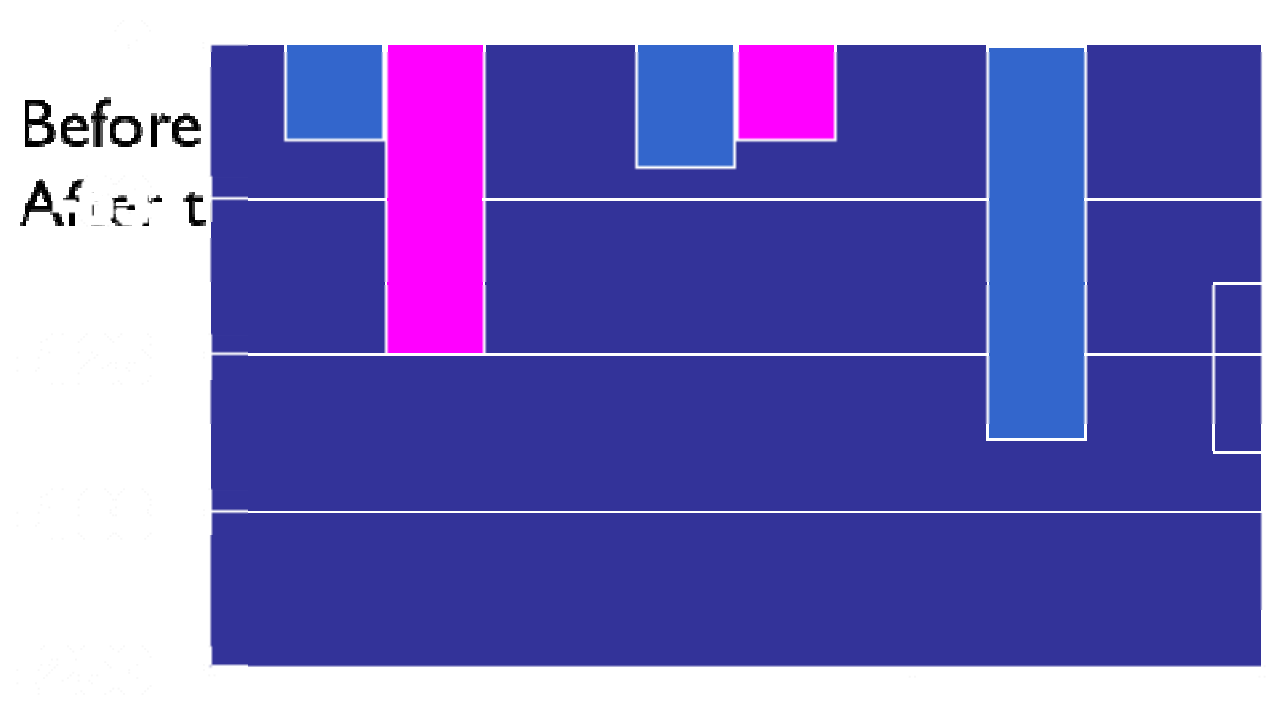
acetylcholine-induced increase  
of peripheral blood flow

Change in peripheral blood flow  
response to administration of L-NMMA

Flow vs baseline



Before  
After t



\* Augmented endothelium-mediated  
vasodilatation

\* Improve of Nitric oxide formation



# **HOW TO TRAIN THE FAILING HEART?**

# Pre-exercise evaluation

- Controlled heart failure
- Exercise test
- Pre-morbid status
- Associated symptoms or diseases
- Contraindication to exercise

# Absolute Contraindications

- Progressive worsening of exercise tolerance or dyspnea at rest or on exertion over 3-5 days
- Significant ischemia at low work rates (<2METs)
- Uncontrolled diabetes
- Acute systemic illness or fever
- Recent Embolism
- Thrombophlebitis
- Active myocarditis or pericarditis
- Moderate to severe aortic stenosis
- Regurgitant valvular heart disease required surgery
- Myocardial infarction within previous 3 weeks
- New onset atrial fibrillation

# Initiation of Aerobic exercise Program

- Ability to speak without signs or symptoms of dyspnea ( $RR < 30$  breaths/min)
- Patients is only modestly fatigue generally
- Crackles present in  $< \frac{1}{2}$  of the lungs
- Resting heart rate  $< 120$  bpm
- Cardiac index  $> 2$  L/min/m<sup>2</sup> (for invasively monitored patients)
- Central venous pressure  $< 12$  mmHg (for invasively monitored patients)

# Exercise Prescription

- **Type:** aerobic, resistance/strengthening
- **Mode:** cycling, walking, [less evidence: aqua exercise, yoga, tai chi, etc.]
- **Intensity:**
- **Duration:**
- **Frequency:**
- **Progression:**



# Exercise Test

- Maximum Exercise Test
- Sub-maximum Exercise Test
  - ▲ 6 minute walk test
  - ▲ Treadmill or bicycle test

# Six Minute Walk Test

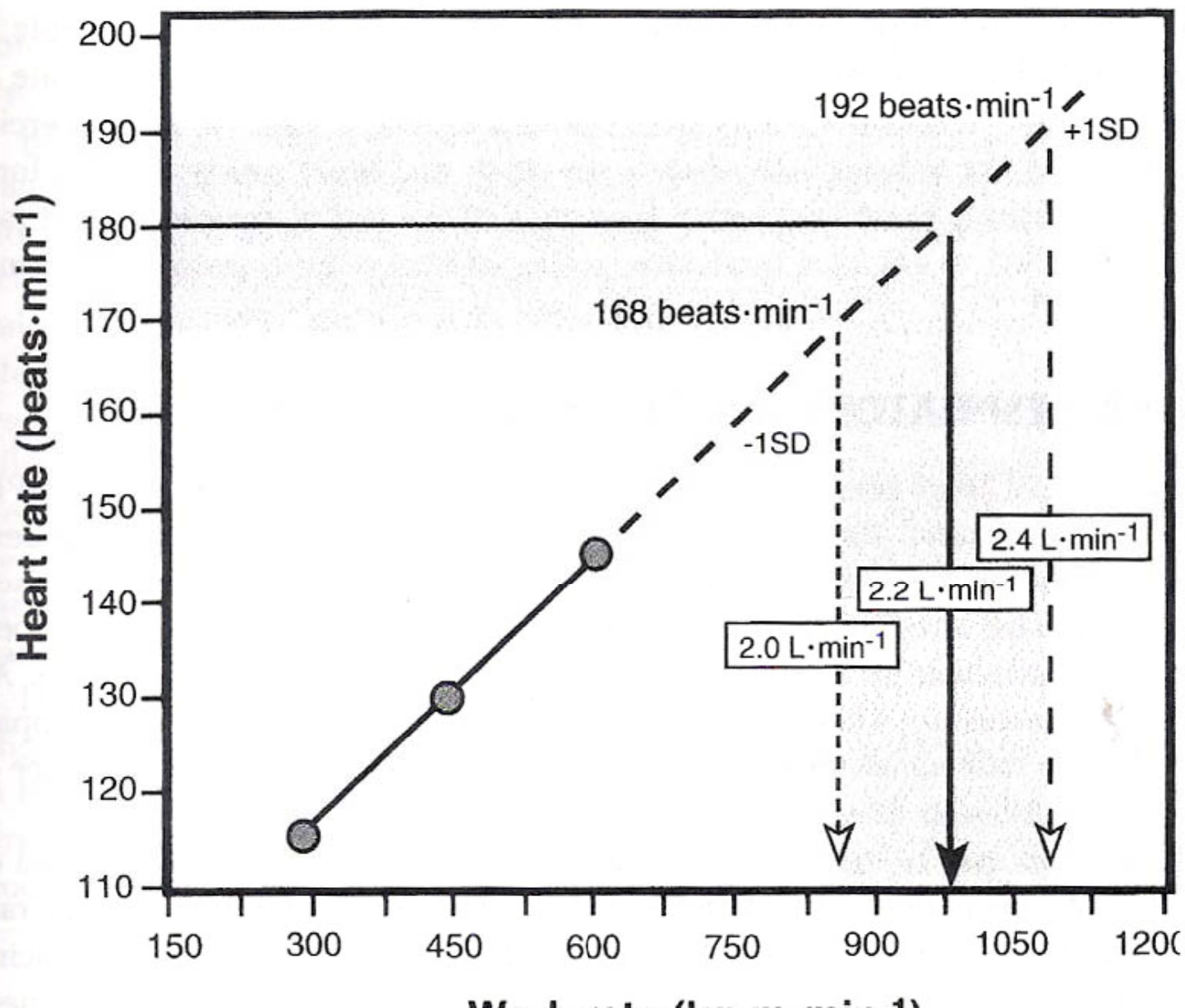


Peak  $\text{VO}_2$  = Distance

Peak  $\text{VO}_2$  max =  $0.03 \times \text{distance (m)} + 3.98$

# Estimated $\dot{V}O_2$ Max

## CA Protocol



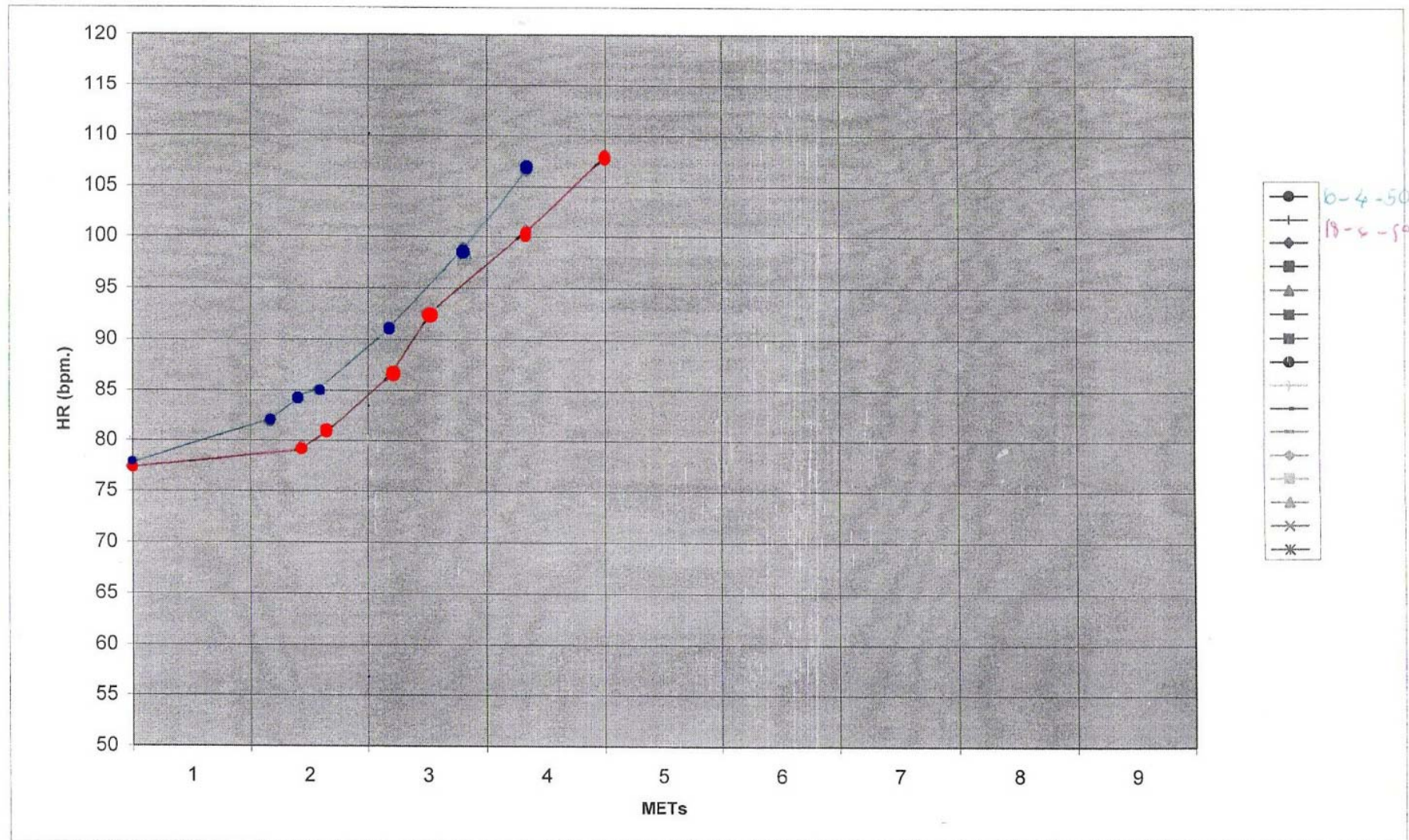
# Sub-maximum Exercise Test

## Treadmill Sub-maximum Exercise Test

Stage	Speed(m/h)	Grade(%)	METs	Time
<b>1</b>	<b>2</b>	<b>3.5</b>	<b>3</b>	<b>3</b>
<b>2</b>	<b>2</b>	<b>7</b>	<b>4</b>	<b>3</b>
<b>3</b>	<b>2</b>	<b>10.5</b>	<b>5</b>	<b>3</b>
<b>4</b>	<b>2</b>	<b>14</b>	<b>6</b>	<b>3</b>



# Training Effects













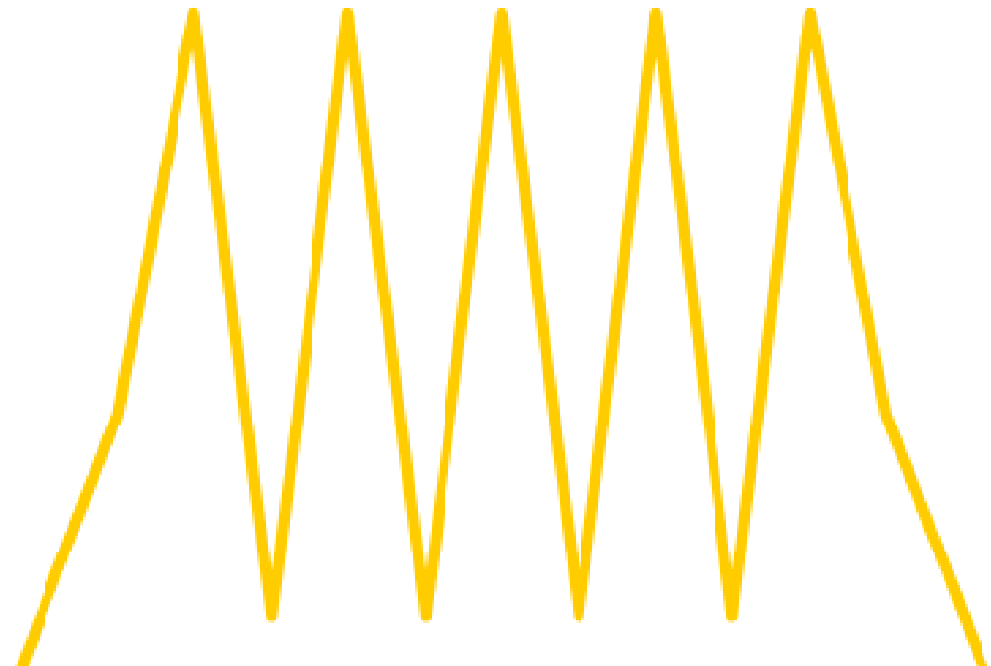


# Aerobic Exercise Prescription

## Steady State vs. High Intensity Interval Training

**Interval Training**

**Steady State Training**

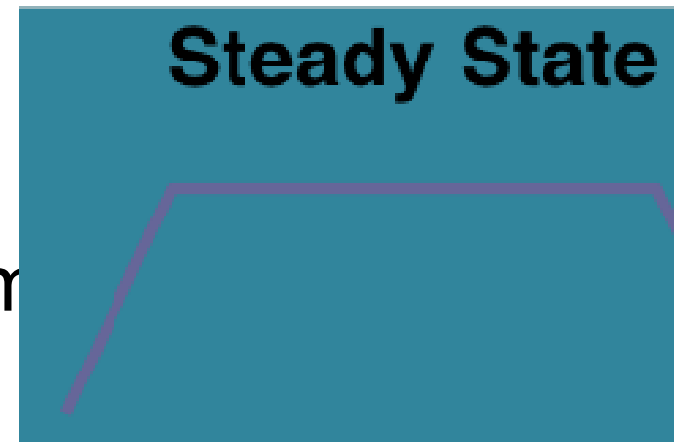


# Exercise Intensity

Intensity	Subject Measures		Physiological /Relative Measure		Absolute Measure
	<i>Talk Test</i>	<i>RPE</i>	<i>% HRR; <math>VO_2R</math></i>	<i>%Max HR</i>	<i>METs, <math>VO_2max</math></i>
<i>Light</i>	<i>Able to talk and/or sing</i>	<i>&lt; 3</i>	<i>&lt; 40</i>	<i>&lt; 64</i>	<i>&lt; 3</i>
<i>Moderate</i>	<i>Able to talk but not sing</i>	<i>3-4 (12-13)</i>	<i>40-60</i>	<i>64-76</i>	<i>3-6</i>
<i>Vigorous</i>	<i>Difficulty talking</i>	<i><math>\geq 5</math></i>	<i>&gt; 60</i>	<i>&gt; 76</i>	<i>&gt;6</i>

# Steady State training

- **Intensity:** 40 –80%  $\text{VO}_{2\text{m}}$
- **RPE = 12-13**
- **Duration; Frequency; Progression**
- Depend on functional capacity
  - < 3 METs: 5-10 min; multiple sessions
  - 3-5 METs: 15 min bid
  - > 5 METs: 20 – 30 min/ 3-5 times/week
- **Increase time before intensity**

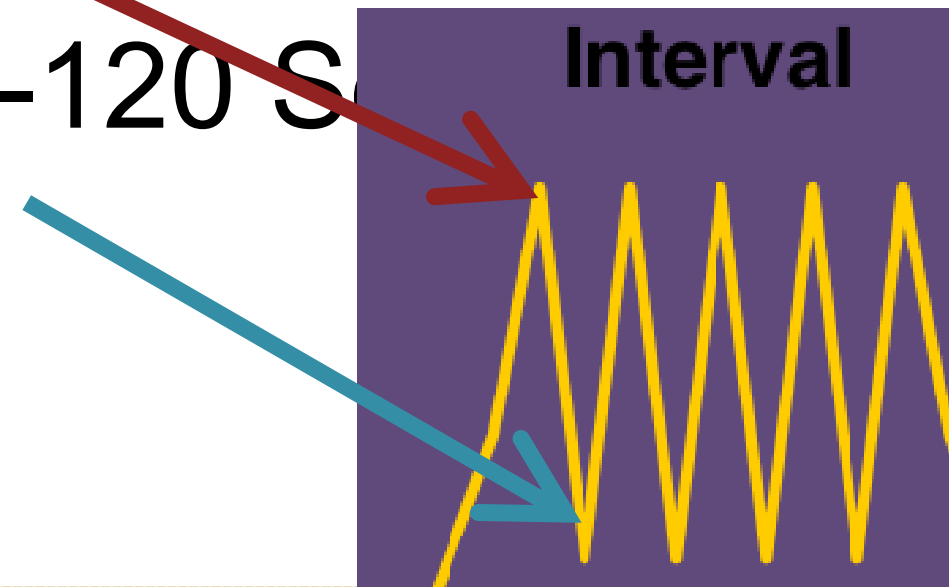


# Interval training

- Be able to apply more intense training
- Intense: 25-95%  $\text{VO}_{2\text{max}}$
- Interval time of Work Phase/Recovery Phase

Work Phase: 30-120 Sec

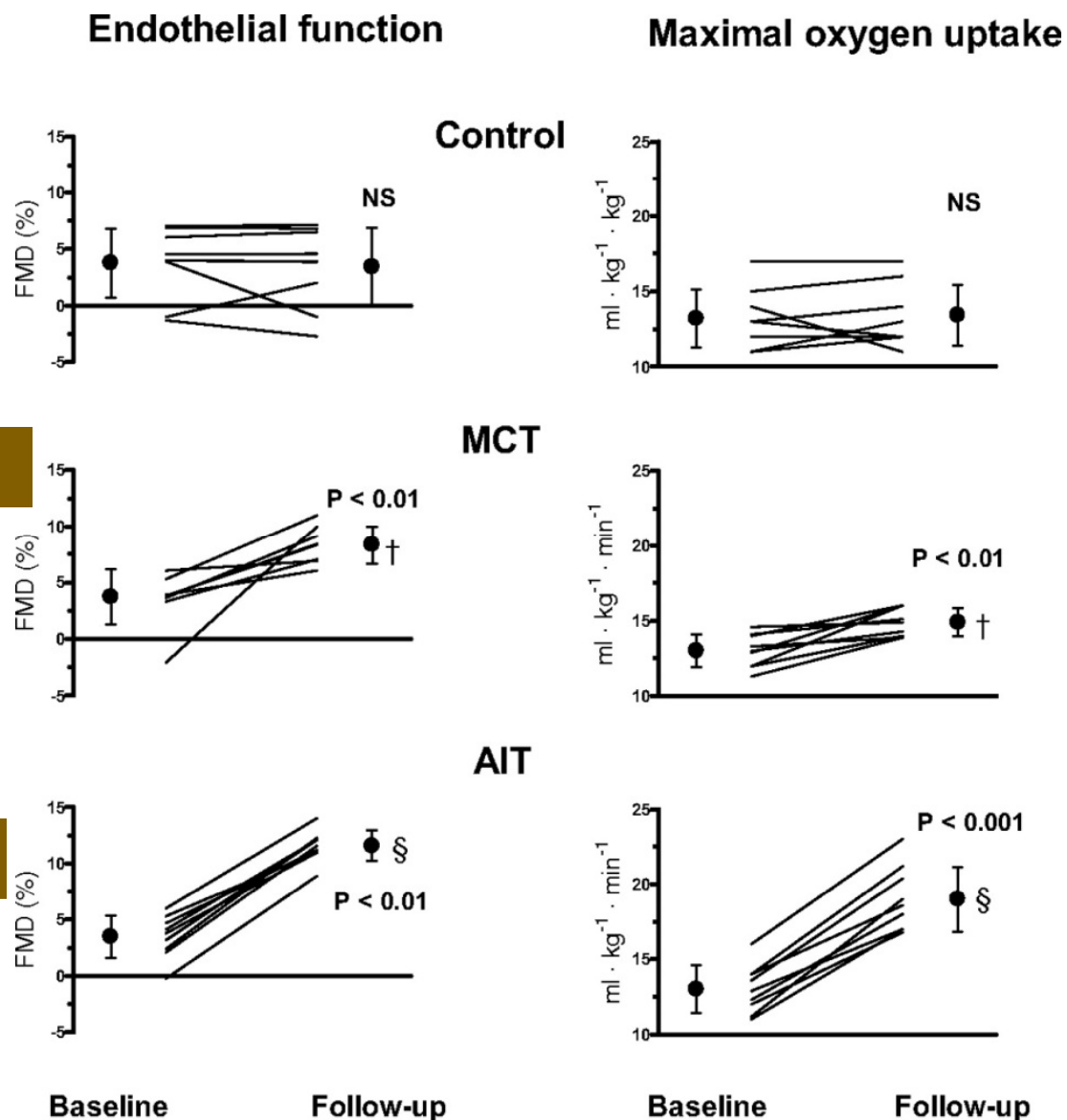
Recovery Phase: 60-120 S



# Moderate Continuous Training in HF patients: A Randomized Study

State

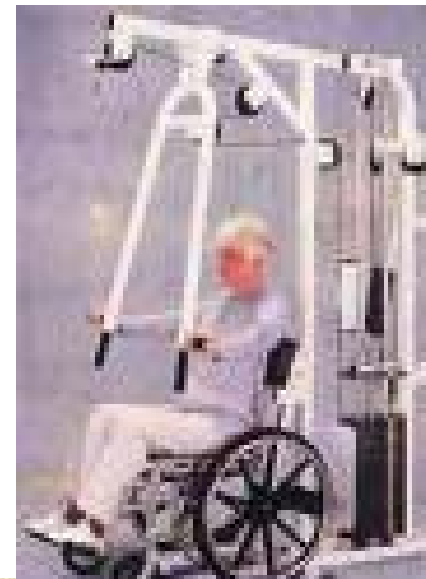
Interval



- 27 stable post-infarction HF with optimal medical treatment
- MCT: [70% peak HR]
- AIT: [95% peak HR]
- Control group
- 3 times/week for 12 weeks



# RESISTANCE TRAINING IN HF



# Efficient Modality Training in HF?

ci Sports Exer 2007

45 HF , FC II-III, LVEF < 35%

Resistance

10 rep /rest 2 minutes  
10 diff weight machines

Endurance

60%-75% of  $\text{VO}_2$  peak

Endurance-Resistance

Control

No significant difference between group:  
 $\text{VO}_{2\text{ peak}}$ , peak workload

# Cardiac Patients

Guidelines	Sets	Repetitions	# of Exercise	Frequency (day/week)
2000 AHA	1	10-15	8-10	2-3
2004 AACVPR	1	10-15	8-10	2-3

A: American Heart Association

VPR: American Association of Cardiovascular and Pulmonary Rehabilitation



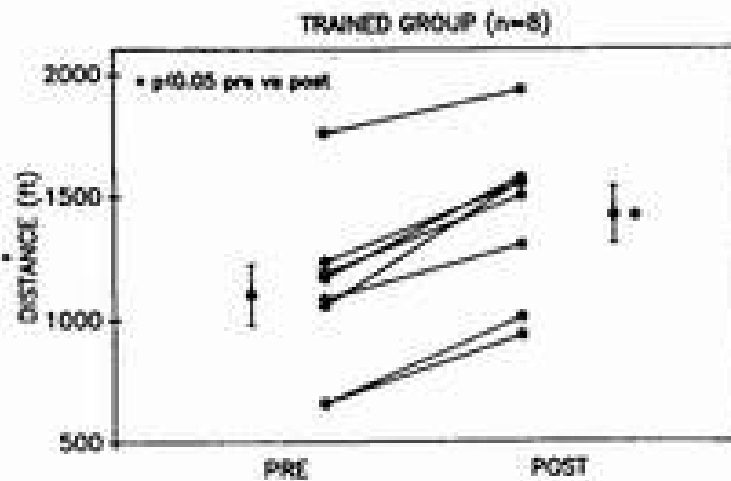
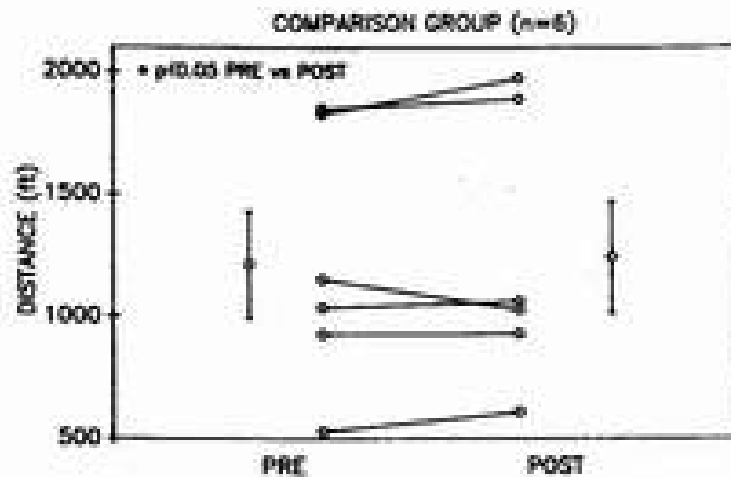


# RESPIRATORY MUSCLE TRAINING

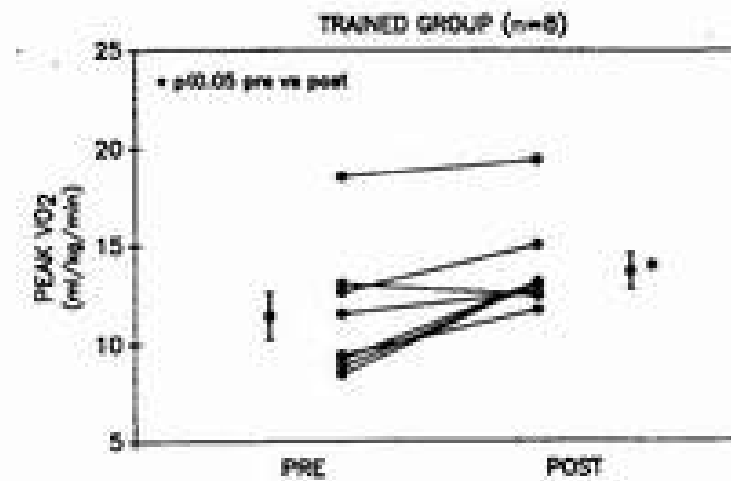
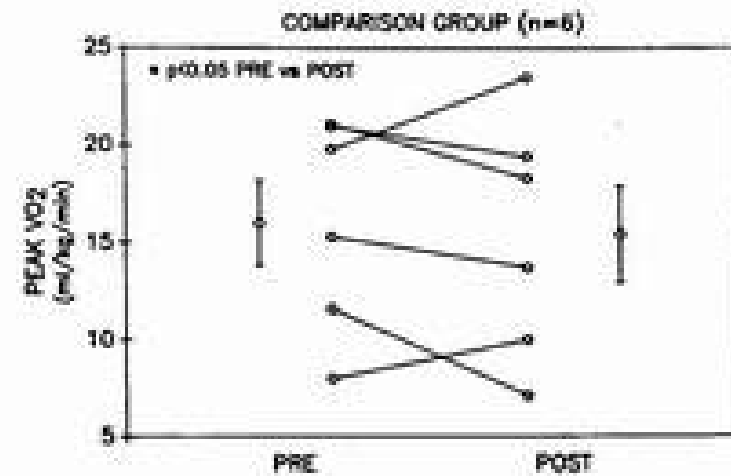


# Failure

## 6 minute walk test



## Peak VO<sub>2</sub>



# Failure

- **Inspiratory Muscle Training Improves Blood Flow to Resting and Exercising Limbs in Patients with CHF**

***JACC 2008.***

- **Effects of Inspiratory Muscle Training on Autonomic Activity,**
- **Endothelial Vasodilator Function, and NT pro-BNP levels in CHF**

***J Cardiopul Rehabil Prev 2008***

- **Inspiratory Muscle Training in Patients with HF**
- **and Inspiratory Muscle Weakness:**

***JACC 2006.***



# **ELECTRICAL STIMULATION: BENEFIT OF TRAINING IN HF**



**stimulation of the legs** and conventional **bicycle exercise training** for patients with chronic heart failure

	<b>Bike training group (n=24)</b>	<b>FES group (n=24)</b>
Exercise time(s)	544/654 < 0.001	501/568 0.02
Peak VO <sub>2</sub> (ml/kg/min)	19.0/19.8 0.276	18.6/18.6 0.932
Quadriceps strength (kg)	48.8-54.1 <0.001	42.3/47.6 0.009
Quadriceps fatigue	0.76/0.84 0.001	491/531 0.005
QOL score	0.105	0.094

# conventional bicycle exercise on endothelium and functional status indices in patients with heart failure

cohort study; compare the effect of muscle functional electrical stimulation (FES) on endothelial function to that of conventional bicycle training.

NYHC class II or III; LVEF  $\leq 35\%$

FES for 6 weeks, with a 6-week washout period then Bicycle training

Brachial artery flow-mediated dilation (FMD)

FES: significant improvement in FMD ( $5.9 \pm 0.5\%$  to  $7.7 \pm 0.5\%$ ,  $p < 0.001$ ).

Bicycle training: FMD ( $6.2 \pm 0.4\%$  to  $9.2 \pm 0.4\%$ ,  $p < 0.001$ )

effect of muscle FES in patients with heart failure

endothelial function, although not equivalent to that

conventional exercise, is substantial. Muscle FES protocols

useful in the treatment of patients with heart failure who

not or

⑩ Tai Chi: Reduced BP (Preventive Cardiol 2008)

Systolic: 3-32 mmHg; Diastolic 2-18 mmHg

⑩ Review 29 studies (9 RCT) (J Cardiopulm Reh  
Prev 2009)

Reduced BP, Improve Exercise capacity

⑩ Tai Chi in HF patients: (American J of Medicine  
2004)

Improve functional capacity and reduce BNP lev

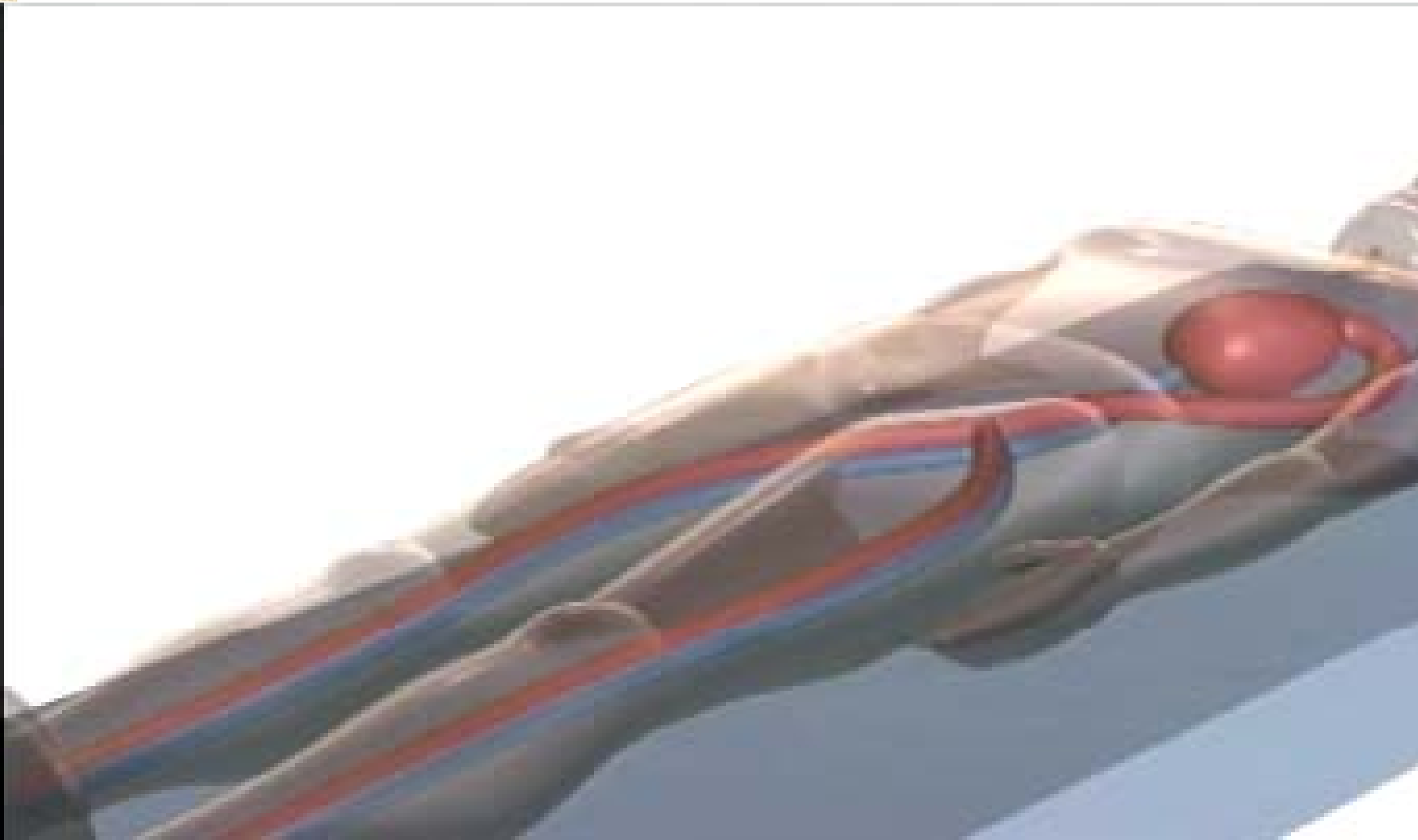


# program





# CounterPulsation: EECP



# US FDA approved indications

- **cardiogenic shock**
- **stable or unstable angina**
- **acute myocardial infarction**
- **congestive heart failure**

# PEECH trial

**P**rospective **E**valuation of **E**nhanced **E**xternal  
**C**outerpulsation in Congestive **H**eart Failure

**RCT:**            93 EECp + optimal medication  
                     94 optimal medication

**Mean LVEF 26% Ischemic: 2/3**

- ⑩ **Ex duration increase > 60 sec (35.4% vs 25.3%) at months (p = 0.016)**
- ⑩ **QOL (Minnesota Living with HF Questionnaire) at 1 wk, 3 mo significantly improve but not at 6 mo**

**Peak VO<sub>2</sub> increase > 1.25 ml/kg/min: not sig. diff**

# Mechanism For Sustained Clinical Benefit With EECp

## Physiologic Effects

- **Potential mechanisms include:**
  - **Recruitment of coronary collaterals.**
  - **Angiogenesis.**
  - **Improved endothelial function.**
  - **Training effects.**
  - **Other, yet unknown mechanism (s).**

# With EECp: Physiologic effects

## Training effects of Exercise

- Increase exercise capacity
- Reduce sympathetic activity
- Endothelial function
- Myocardial adaptation
- Peripheral adaptation
- Risk factors modification





**SAFETY**

## HF (FC II, III)

- Meta-analysis of 29 RCT, 1126 patients only one trial report complication (Cochrane Database 2004)
- RCT of 110 patients: no complication (Circulation 1999)
- Home Exercise programs with low intensity/electrical stimulation are safe.



# Summary

- Exercise training in HF is safe and has beneficial effects if performed properly.
- Benefits: reduce mortality, reduce morbidity, improve QOL, exercise capacity and etc.
- Every HF patient who had no contraindications to exercise should be encouraged to engage in physical activity and prescribed exercise program
- Aerobic Exercise Training: Interval Training may have more benefit than Steady State training
- Respiratory Muscle Training and Electrical stimulation are recommended.
- **Individualized exercise** program is very important for successful exercise prescription

# Goal of Management for HF Patient

## Cardiac Remodeling

