

Physiological Research Pre-Press Article

4 **Running Head: Central and Peripheral Response to Cycling in Older Men**

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6 **Central and Peripheral Response to Incremental Cycling Exercise in Older**

7 **Untrained Active Men: A Comparison of those In-Between**

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28 **Summary**

29 **Aim:** The aim of this study was to compare the central and peripheral components of
30 cardiorespiratory fitness during incremental to maximal exercise between older men who
31 were either recreational athletes (RA) or leisurely active (LA) men, i.e., those who fall
32 between trained and untrained. **Methods:** This was a cross-sectional study in which all
33 subjects completed an exercise test on a cycle ergometer. Maximal oxygen consumption
34 (VO_{2max}) and ventilatory threshold (VT) were assessed using gas analysis, and central
35 components of VO_{2max} were assessed using a non-invasive thoracic bio-impedance device.
36 **Results:** VO_{2max} (RA: 45.1 ± 4.8 ml/kg/min; LA: 32.2 ± 4.6 ml/kg/min, $p < 0.001$) and SV
37 at maximal exercise (RA: 133.5 ± 24.96 ml/beat; LA: 107.9 ± 17.6 ml/beat, $p = 0.005$) were
38 higher in the RA group compared to the LA group. A plateau in SV occurred between 30-
39 45% of maximal exercise capacity in the RA group. No differences in SV were observed
40 across workloads in the LA group. No differences in the calculated arterio-venous oxygen
41 difference ($(a-v)O_{2diff}$) were observed between groups. **Conclusion:** Training volume
42 appears to influence central components of cardiorespiratory fitness among a matched
43 sample of older men who are neither trained nor untrained. This builds a case for
44 increasing the volume of training to preserve cardiorespiratory fitness among older men.
45 **Key Words:** *stroke volume, aging, VO_{2max} , fitness*

46 **Introduction**

47 Endurance trained older adults have higher maximal cardiac output (Q), stroke
48 volume (SV), and arterio-venous oxygen difference ((a-v)O_{2diff}) compared to untrained
49 older adults (McLaren et al, 1997). While it is generally accepted that SV plateaus at 40-
50 50% of maximal exercise capacity (Astrand et al, 1964), there is some research to suggest
51 that highly trained individuals may have a continuous increase in SV to maximal exercise
52 (Rivera et al, 1989). Such research has typically compared highly trained older adults
53 (VO_{2max} > 50ml/kg/min) to untrained/sedentary older adults (VO_{2max} < 30ml/kg/min)
54 (Rivera et al, 1989; Dogra et al, 2012). The central and peripheral response of
55 recreationally active older adults that fall in between these extremes of trained and
56 untrained is not well described. Specifically, the cardiovascular exercise response profile
57 of older men who are recreational athletes (RA) or simply leisurely active (LA) is not
58 known.

59 Thus, the purpose of this study was to describe the cardiovascular response of
60 older active men, who are neither highly trained nor completely sedentary, to incremental
61 maximal cycling exercise. It was hypothesized that RA would have a similar response as
62 that previously observed in highly trained older men, and that the RA would exhibit
63 greater cardiovascular health than the LA.

64

65 **Materials and Methods**

66 *Study Design and Subjects*

67 Inclusion in this cross-sectional study was limited to males aged 60-80 years
68 without any chronic cardiovascular or respiratory conditions. Subjects were recruited from

69 local cycling groups. They completed a training log (7 day recall) prior to testing and were
70 divided into RA or LA based on this recall. Subjects were considered RA if they were
71 participating in moderate-vigorous cycling for a minimum of one hour 3-4 times per week
72 Subjects were considered LA if they were meeting the minimum recommendations of
73 150mins/week of moderate to vigorous physical activity (Paterson et al, 2010). A total of
74 36 older men were screened for participation; two subjects declined further participation
75 and four were considered ineligible due to age and cardiovascular health impairments.
76 Two subjects were excluded from analysis due to technological difficulties during testing,
77 resulting in unusable data. No subjects were taking any medications that would affect
78 cardiovascular response to exercise. All subjects were pre-screened to ensure they were at
79 minimal risk for participation in the exercise testing and all subjects provided written
80 informed consent prior to laboratory testing (Canadian Society of Exercise Physiology,
81 2013). All procedures were approved by the Research Ethics Boards of Acadia
82 University and the University of Ontario Institute of Technology.

83

84 *Methodology*

85 Subjects attended one laboratory session. Familiarization to the mode of exercise
86 was not necessary as all subjects were regularly cycling. Anthropometric measures of
87 height, weight, and waist circumference were assessed using a standard medical scale and
88 a tape measure to the nearest 0.5 cm, 0.1 kg and 0.1 cm respectively. Body mass index
89 (BMI) was calculated as $\text{weight (kg)}/\text{height}^2 \text{ (m)}$. Resting heart rate and blood pressure
90 were measured manually after resting in a seated position for 5 minutes. Blood pressure
91 was recorded manually a total of three times. Measurements were entered into the

92 software prior to and following calibration as per Physioflow instructions. Subjects were
93 then fitted with electrodes and connected to a non-invasive thoracic electric bio-impedance
94 device (Physioflow Enduro, Bristol, PA USA). Resting measures of Q and SV were
95 recorded for up to 2 minutes after sitting quietly for 3 minutes.

96 Subjects then completed a maximal exercise test on a cycle ergometer (LODE
97 Excalibur, Lode BV, Groningen, The Netherlands) using a ramp incremental protocol
98 (25Watts/min). Subjects maintained a self-selected pace between 70-100 revolutions per
99 minute. Maximal oxygen consumption (VO_{2max}) was determined by a plateau in VO_2 and
100 confirmed by a respiratory exchange ratio > than 1.1, achievement of age-predicted
101 maximum heart rate ($220-age$), a rating of perceived exertion > 19 (scale of 6-20), and/or
102 volitional exhaustion.

103

104 *Measures*

105 *Gas exchange measurements:* Expired CO_2 and O_2 were collected through a
106 pneumotachograph (Hans Rudolph 2700) and were analyzed using a gas collection system
107 (Parvo Medics OUSW 4.3, USA) at five-second intervals (to align with the impedance
108 cardiography output). VO_{2max} was recorded as an average of the highest 25 second period
109 (i.e. 5 data points). The first ventilatory threshold (VT) was visually determined
110 independently by two researchers as the point where ventilation increased non-linearly to
111 the increase in O_2 uptake and by identifying the point at which CO_2 production increased at
112 a faster rate than VO_2 .

113 *Central and Peripheral Components of VO_2 :* Heart Rate (HR), SV, end diastolic volume
114 (EDV), early diastolic filling ratio (EDFR) and Q were measured at 5 second intervals

115 non-invasively using thoracic electric bio-impedance signals. The Physioflow uses
116 changes in transthoracic impedance during cardiac ejection to calculate SV (Charloux et
117 al, 2000) via high-frequency and low-amperage alternating electric current from six
118 electrodes. The accuracy and reproducibility of the Physioflow has been assessed in
119 normal-weight and overweight participants during an incremental to maximal exercise test
120 against the direct Fick method (Richard et al, 2001). The mean difference between values
121 obtained by the Physioflow was 0.009 l min^{-1} , and the correlation coefficient between the
122 Physioflow and the direct Fick method was $r = 0.946$ (Richard et al, 2001). In the present
123 study, the Fick equation was used to calculate $(a-v)O_{2\text{diff}}$ as follows: $(a-v)O_{2\text{diff}} (\text{ml O}_2/100$
124 $\text{ml blood}) = [\text{VO}_2 (\text{l/min})/ Q (\text{l/min})] \times 100$. The Fick equation has been used in previous
125 studies to calculate $(a-v)O_{2\text{diff}}$ at rest and during exercise in normal-weight and obese
126 individuals (Vella et al, 2011) and has been deemed to be accurate as per Richard et al.,
127 2001. Measures of VO_2 and Q data were time aligned for analysis.

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129 *Statistical Analyses*

130 Data in tables are presented as means and standard deviation (SD). Independent
131 samples t-tests were used to compare subject characteristics and $\text{VO}_{2\text{max}}$ parameters
132 between RA and LA. Paired sample t-tests were used to determine differences among
133 parameters during exercise. A repeated measures analysis of variance was used to assess
134 for differences within groups during the incremental exercise test. Data in graphs are
135 presented as means and standard error. All statistics were conducted in SPSS v21 (SPSS
136 Inc., Armonk, NY). Statistical significance was declared at $p < 0.05$.

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138 **Results**

139 RA were cycling 186.9 (± 22.6) km per week and training 7.9 (± 1.5) hours per
140 week. LA were taking part in light to moderate intensity physical activity (e.g. walking,
141 gardening and cycling) 5.6 hours per week. One LA was a smoker. Additional sample
142 characteristics are presented in Table 1. Other than VO_{2max} and resting heart rate, there
143 were no differences between the RA and LA group i.e. they were well matched for age
144 and body composition.

145 VO_{2max} was significantly different between groups. VT occurred at 75% of VO_{2max}
146 (range: 51-88%) in the RA and 72% of VO_{2max} (52-88%) in the LA ($p=0.4$). However, the
147 absolute VO_2 at VT was significantly different between groups (RA: 2.6 l/min; LA: 1.8
148 l/min; $p<0.001$). Additional data at VT and maximal exercise for both groups are
149 available in Table 2. At VT and maximal exercise, VO_2 , Q and power (in Watts) were
150 significantly higher in RA compared to LA. SV at VT was approaching significance
151 ($p=0.055$) and was higher in the RA at maximal exercise ($p=0.01$). EDV was significantly
152 higher in the RA at VT and was approaching significance at maximal exercise ($p<0.01$).
153 No group differences were noted for HR or calculated (a-v) O_{2diff} at any workload. At
154 relative exercise intensities, between group differences in Q were observed across all
155 intensities, except for 45%. Differences observed in Q were largely attributed to
156 differences in SV at these intensities, as HR only differed between groups at 70, 90, and
157 100% of maximal exercise.

158 SV increased by 15% in RA compared to 6% in the LA from 100W to maximal
159 exercise. There were increases in Q, HR and (a-v) O_{2diff} during incremental to maximal
160 exercise within the LA and RA groups. SV in the RA group increased significantly during

161 incremental to maximal exercise ($p=0.02$); however, no differences in SV were observed
162 in the LA ($p=0.4$). The SV, Q, HR, and calculated $(a-v)O_{2diff}$ profiles are displayed in
163 Figure 1. SV was significantly higher in the RA compared to LA at maximal exercise and
164 across most submaximal exercise intensities. SV increased at 30% VO_{2max} compared to
165 25% VO_{2max} ($p=0.03$) and appeared to increase again at 45% VO_{2max} compared to 40%
166 VO_{2max} ($p=0.06$) in the RA.

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168 **Discussion**

169 The primary finding of this study is that RA appear to have a more dynamic SV
170 response during exercise than LA. Among the RA, SV plateaued at 30-45% of maximal
171 exercise, while the LA had no change in SV throughout exercise. The secondary finding is
172 that among a matched sample of older men, higher cardiorespiratory fitness in RA is
173 primarily due to a greater central response (i.e., $\uparrow SV$), as the peripheral response (i.e., a -
174 vO_{2diff}) between RA and LA appears to be the same. This novel investigation fills a gap in
175 current knowledge pertaining to the cardiovascular response to maximal exercise testing in
176 non-endurance trained and non-sedentary older men.

177 Our findings are the first, to our knowledge, to compare the cardiovascular
178 response to maximal cycling exercise in recreationally active older men. We observed a
179 SV plateau between 30-45% of maximal exercise in the RA, similar to what has been
180 previously reported in highly trained older adults. Specifically, in a study of older
181 endurance trained male runners and cyclists, (i.e. those who had trained consistently for at
182 least 3 years and who routinely ran a minimum of >30 km per week) participants
183 completed an exercise test on a cycle ergometer to volitional exhaustion. A plateau in SV

184 was observed at approximately 40% of VO_{2max} in the runners and 30% of VO_{2max} in the
185 cyclists (McLaren et al, 1997). Endurance trained men and master endurance trained
186 runners (age: 51-72 years) can progressively increase SV up to the point of 70% and 85%
187 of VO_{2max} , respectively (Proctor et al, 1998; Rivera et al, 1989). Thus, training level (i.e.
188 highly trained versus recreational athlete versus recreationally active) appears to have a
189 significant impact on the cardiovascular response to maximal exercise among older adults
190 as well.

191 Visually, a second increase in SV appeared after the initial plateau at 65% of
192 VO_{2max} in the RA ($p=0.15$ NS). This observation is of particular interest, as the bulk of the
193 research investigating such an increase in SV has primarily focused on young highly
194 trained adults. There is some evidence that such an increase may occur in moderately
195 active adults aged 18-30 years, but no such data are available in older men. From data on
196 middle-aged men with moderate fitness levels, it appears that the dominant response of SV
197 to maximal exercise is a plateau followed by a subsequent decrease in SV (Ferguson et al,
198 2011; Skof et al, 2012). A study by Skof and colleagues (2012) examined the SV response
199 to maximal exercise in a group of middle-aged men who were either highly trained
200 runners [VO_{2max} : 54.1 ± 3.8 (ml/kg/min)] or moderately trained runners (VO_{2max} : 36.8 ± 3.3
201 ml/kg/min). The highly trained group exhibited higher VO_{2max} , Q, and SV, similar to the
202 present study. Skof and colleagues (2012) also noted various responses in SV in both
203 groups: a plateau, a plateau with a subsequent drop, a progressive increase, and a plateau
204 with a significant decrease (in the highly trained group). Our data indicate that an increase
205 may be possible in older men who are RA as well.

206 The higher SV observed in the more trained groups among previous studies may
207 be related to a higher EDV. In the present study, EDV was greater in the RA at VT and
208 approaching significance at maximal exercise compared to the LA; however EDR was
209 not. A study comparing older trained and sedentary adults using Doppler
210 echocardiographic data found no differences in left ventricular diastolic filling
211 characteristics except for a trend for atrial filling fraction (Jungblut et al, 2000). Similarly,
212 Carrik-Ranson et al. (2012) did not find differences in left ventricle filling or lengthening
213 when comparing younger and older men. In other words, it seems that preload may not be
214 predicting differences in SV among older men, but rather factors such as blood volume or
215 ventricular contractility and autonomic response may be. Future research is needed to
216 determine which factors are leading to differences in SV among older men and to confirm
217 the possibility of an increase in SV after the initial plateau. Nevertheless, the present
218 finding offers interesting insight into cardiovascular aging and lifelong exercise
219 engagement.

220 The observation that differences in VO_{2max} were exclusively due to differences in
221 central components between groups was of significant interest. Research by Murias and
222 colleagues (2011) showed that older men who engaged in a 12 week aerobic training
223 program on a cycle ergometer were able to increase (a-v) O_{2diff} , citrate synthase and
224 capillarization (Murias et al, 2011). As such, it was expected that the RA group in the
225 present study would have a higher calculated (a-v) O_{2diff} than the LA. Evidence from
226 cross-sectional studies of men and women support the findings of the present study in that
227 they have also noted reliance on central components and no differences in peripheral
228 components between trained and untrained groups (Proctor et al, 1998; Dogra et al, 2012).

229 Specifically, in a study by Sagiv and colleagues (2007), 15 older aerobically trained
230 (VO_{2max} : 42.1 ± 2.1 ml/kg/min) and 15 older untrained (VO_{2max} 31.1 ± 2.4 ml/kg/min) men
231 underwent a maximal exercise test on a cycle ergometer. The trained group exhibited
232 greater reliance on central components of cardiorespiratory fitness; similar oxygen
233 extraction was noted in both trained and untrained groups (Sagiv et al, 2007). This
234 discrepancy in $(a-v)O_{2diff}$ between cross-sectional and laboratory based intervention
235 research requires further investigation as it may ‘shed light’ on important methodological
236 limitations or physiological mechanisms associated with the aging process.

237 The present study utilized technology that enabled continuous measurement of SV.
238 This may have allowed for more accurate identification of a peak in both our RA and LA
239 groups as well as the identification of a second increase in SV. Previous research has
240 primarily used methods such as acetylene wash in, which assesses Q and SV at
241 predetermined workloads only (Dogra et al, 2012). The continuous measurement of Q was
242 a significant strength of this study. Limitations of the present study were the lack of direct
243 measurement of $(a-v)O_{2diff}$, lack of a detailed history of exercise levels, and the cross-
244 sectional study design. Future research should make use of technology such as near infra-
245 red spectroscopy to better understand the physiological responses of muscle oxygen
246 extraction.

247

248 **Conclusions**

249 In conclusion, it appears that a higher cardiorespiratory fitness in older aerobically
250 active men (RA) is due primarily to a higher Q, specifically SV, when compared to LA
251 older men. Of note, the present findings suggest a secondary increase in SV may be

252 possible in RA whereas, the SV response in the LA remains unchanged during incremental
253 to maximal exercise. These findings provide novel insight into the cardiovascular
254 response to incremental cycling exercise among non-elite and non-sedentary older men.
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309 Table 1. Sample Characteristics (Mean \pm SD)

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Characteristic	Recreational Athletes (n=15)	Leisurely Active (n=13)
	Mean \pm SD	Mean \pm SD
Age (years)	65.4 \pm 3.5	67.6 \pm 4.1
Weight (kg)	79.6 \pm 7.3	78.7 \pm 11.5
Height (cm)	173.7 \pm 6.1	172.3 \pm 4.7
Waist Circumference (cm)	90.3 \pm 6.6	93.4 \pm 8.2
Body Mass Index (kg.m ⁻²)	26.3 \pm 2.1	26.5 \pm 3.5
Systolic Blood Pressure (mmHg)	117.0 \pm 8.1	116.2 \pm 8.9
Diastolic Blood Pressure (mmHg)	74.0 \pm 9.1	71.8 \pm 5.7
Resting Heart Rate (beats/min)	57.3 \pm 9.3	64.7 \pm 8.5*

*p<0.05

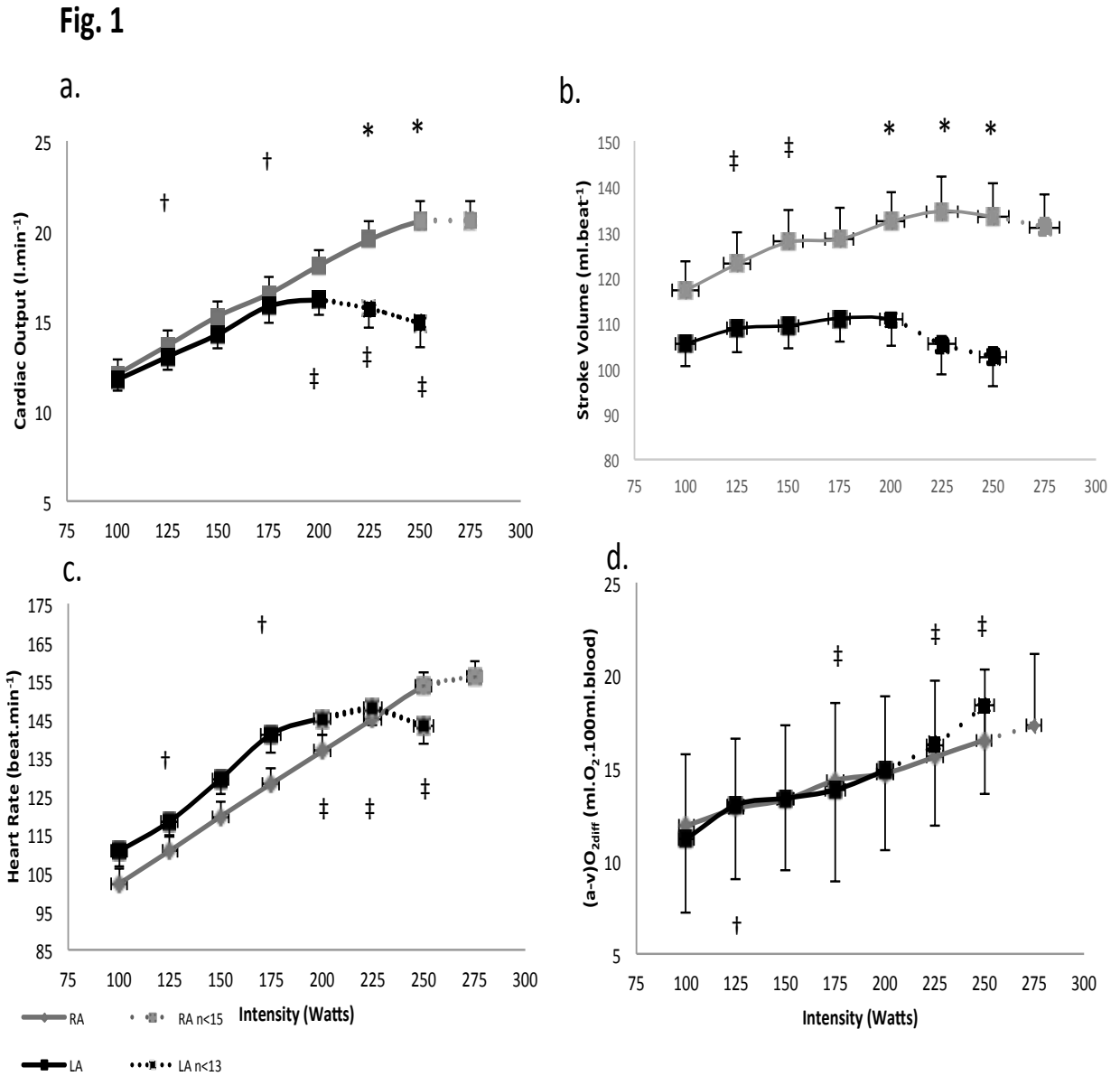
332 Table 2. Cardiovascular Parameters from Incremental to Maximal Cycling Exercise in
 333 Recreational Athletes (n= 15) and Leisurely Active (n=13) Older Men

Characteristic	At Ventilatory Threshold		At Maximal Exercise	
	Recreational Athletes	Leisurely Active	Recreational Athletes	Leisurely Active
	Mean \pm SD	Mean \pm SD	Mean \pm SD	Mean \pm SD
VO ₂ (l/min)	2.6 \pm 0.3	1.8 \pm 0.2*	3.5 \pm 0.4	2.5 \pm 0.4*
VO ₂ (ml/kg/min)	34.1 \pm 7.8	23.4 \pm 4.6*	45.1 \pm 4.8	32.2 \pm 4.6*
(a-v)O ₂ diff (mlO ₂ /100ml)	14.1 \pm 5.8	12.4 \pm 4.3	16.6 \pm 4.6	14.7 \pm 3.7
Heart Rate (beats/min)	138.3 \pm 21.6	130.2 \pm 12.0	156.0 \pm 15.6	150.5 \pm 15.0
Stroke Volume (ml/beat)	131.9 \pm 28.7	112.8 \pm 19.9 [^]	133.5 \pm 24.9	107.9 \pm 17.6*
Cardiac Output (l/min)	17.7 \pm 3.5	14.7 \pm 2.6*	20.9 \pm 4.0	16.3 \pm 3.0*
Power (Watts)	193.3 \pm 46.7	146.4 \pm 30.8*	280.0 \pm 31.6	209.6 \pm 29.8*
End Diastolic Volume (ml)	213.1 \pm 41.9	176.1 \pm 31.0*	214.9 \pm 33.4	188.1 \pm 38.1 [^]
Early Diastolic Filling Ratio	74.1 \pm 37.9	75.0 \pm 47.2	81.3 \pm 28.9	80.8 \pm 19.3

334 * p<0.05; [^]p 0.05 to <0.10

335 CI: Confidence Interval; VO₂: Oxygen Uptake; (a-v)O₂diff: arterio-venous Oxygen
 336 difference
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338 Figure 1. Cardiac Output a, Stroke Volume b, Heart Rate c, and (a-v)O_{2diff} d Response to
 339 Incremental to Maximal Cycling Exercise in Recreational Athletes and Leisurely Active Older
 340 Men.
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- * p<0.05 between groups
- ‡ p<0.05 within RA
- † p<0.05 within LA

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345 Figure Description



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351 Legend: Number of subjects who completed each workload by group:

Workload (Watts)	100	125	150	175	200	225	250	275
Recreational Athletes	15	15	15	15	15	15	15	13
Leisurely Active	13	13	13	12	11	7	5	NA

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