# Comparison between esophageal and intestinal temperature responses to upper-limb exercise in individuals with spinal cord injury

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29 ABSTRACT

30 Study Design: Experimental study.

31 **Objective**: Individuals with spinal cord injuries (SCI) may present with impaired sympathetic 32 control over thermoregulatory responses to environmental and exercise stressors, which can 33 impact regional core temperature ( $T_{core}$ ) measurement. The purpose of this study was to 34 investigate whether regional differences in  $T_{core}$  responses exist during exercise in individuals 35 with SCI.

**Setting**: Rehabilitation centre in Wakayama, Japan.

**Methods**: We recruited 12 men with motor-complete SCI (7 tetraplegia, 5 paraplegia) and 5 38 able-bodied controls to complete a 30-minute bout of arm-cycling exercise at 50%  $\dot{V}O_2$ peak.  $T_{core}$  was estimated using telemetric pills (intestinal temperature;  $T_{int}$ ) and esophageal probes  $(T_{eso})$ . Heat storage was calculated from baseline to 15 and 30 minutes of exercise.

41 **Results**: At 15 minutes of exercise, elevations in  $T_{eso}$  ( $\Delta 0.39 \pm 0.22^{\circ}$ C; P<0.05), but not  $T_{int}$ 42  $(\Delta 0.04 \pm 0.18^{\circ}C; P=0.09)$ , were observed in able-bodied men. At 30 minutes of exercise, men 43 with paraplegia and able-bodied men both exhibited increases in  $T_{eso}$  (paraplegia:  $\Delta 0.56 \pm 0.30^{\circ}$ C, P<0.05; able-bodied men:  $\Delta 0.60\pm0.31^{\circ}$ C, P<0.05) and T<sub>int</sub> (paraplegia:  $\Delta 0.38\pm0.33^{\circ}$ C, P<0.05; 44 able-bodied men:  $\Delta 0.30\pm0.30^{\circ}$ C, P<0.05). T<sub>eso</sub> began rising 7.2 min earlier than T<sub>int</sub> (pooled, 45 46 P < 0.01). Heat storage estimated by T<sub>eso</sub> was greater than heat storage estimated by T<sub>int</sub> at 15 minutes (P=0.02) and 30 minutes (P=0.03) in men with paraplegia. No elevations in T<sub>eso</sub>, T<sub>int</sub>, or 47 48 heat storage were observed in men with tetraplegia.

49 Conclusions: While not interchangeable, both T<sub>eso</sub> and T<sub>int</sub> are sensitive to elevations in T<sub>core</sub>
 50 during arm-cycling exercise in men with paraplegia, although T<sub>eso</sub> may have superior sensitivity
 51 to capture temperature information earlier during exercise.

## 53 INTRODUCTION

Exercise-induced elevations in core body temperature ( $T_{core}$ ) are regulated within an acceptable range by the sweating and skin blood flow responses, mediated in part by the sympathetic nervous system in a negative feedback loop [1, 2]. While this is an adaptive process in ablebodied individuals, individuals with spinal cord injuries (SCI) may present with varying degrees of autonomic dysfunction as a result of damage to the spinal cord. Altered autonomic function has been shown to impair sympathetically-mediated regulatory processes below the lesion level, such as the thermoregulatory response to heat and exercise [3, 4].

61 Autonomic function in SCI may differentially affect the validity and sensitivity of different temperature measurement tools through alterations in regional thermoregulatory 62 control. Rectal and esophageal temperatures ( $T_{eso}$ ) provide close estimates of  $T_{core}$  during 63 64 exercise, but are restricted to 'lab-based' assessments that are not feasible to measure in the field [5, 6]. Ingestible telemetric pills can provide wireless measurement of intestinal temperature 65 (T<sub>int</sub>), an estimate of T<sub>core</sub>, and have been shown to be sensitive to heat- and exercise-related 66 thermoregulatory challenges in both able-bodied men and individuals with SCI [7–9]. However, 67 given the known differences in both gut motility [10] and abdominal skin temperature with 68 69 higher level SCI [11, 12], it is unknown whether the telemetric pill provides equally sensitive 70 assessment of T<sub>core</sub> in all individuals with SCI. Furthermore, much of the SCI thermoregulation research has been conducted in athletes with SCI [7, 11, 13–16], with relatively little knowledge 71 72 on the generalizability of methods to untrained adults with SCI who may have altered 73 thermoregulatory sensitivity to exercise [17] and lower levels of heat production related to reduced aerobic capacity [18]. The ability to rely on non-invasive methods to estimate exercise-74 75 or environment-induced T<sub>core</sub> elevations is particularly important to untrained adults with SCI

who are not habitually exposed to temperature stressors, and therefore may be at elevated risk forheat stress exposure.

The purpose of this study was to investigate whether T<sub>int</sub> and T<sub>eso</sub> provide comparable 78 79 information about the thermoregulatory response to upper-limb exercise in untrained men with paraplegia and tetraplegia. To simulate realistic exercise conditions in this population,  $T_{core}$ 80 sensitivity was tested during arm cycling at 50% of VO<sub>2</sub>peak reserve, corresponding to 81 moderate-to-vigorous intensity exercise recommended by the latest physical activity guidelines 82 for adults with SCI [19]. Based on previous work in able-bodied individuals [9], we 83 84 hypothesized that T<sub>int</sub> and T<sub>eso</sub> would both reflect exercise-related elevations in T<sub>core</sub> in individuals with paraplegia and tetraplegia. 85

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#### 87 METHODS

#### 88 Experimental Design

Seventeen men were recruited for this study: seven individuals with tetraplegia, five individuals 89 90 with paraplegia, and five able-bodied controls (Table 1 for participant characteristics). Participants were asked to come to the laboratory for two separate visits for a baseline testing 91 session, as well as an acute exercise trial. During the first visit, degree of autonomic dysfunction 92 93 was assessed by sympathetic skin responses (SSR) and the sit-up test to identify orthostatic 94 hypotension (OH), followed by a graded arm exercise test to exhaustion. At the second visit, participants performed 30 minutes of arm cycling exercise at 50% VO<sub>2</sub>peak, during which T<sub>core</sub> 95 and skin temperature  $(T_{sk})$  were recorded throughout. The testing environment was maintained at 96 97 25°C at a relative humidity of 50%.

Sympathetic Skin Responses: SSR testing was completed as previously described [20, 21], with 98 99 modification. In the supine position, electrodes were placed on the dorsal and ventral sides of the 100 left hand and foot, and were attached to a commercial data acquisition unit (PowerLab 26T 101 (LTS), AD Instruments, Colorado Springs, CO, USA). In 30-40 second intervals, electrical stimulation was applied to the left median nerve, the left posterior tibial nerve, and the left supra-102 103 orbital nerve, in series. Five stimuli were applied at each nerve, with the pulse amplitude set to a 104 threshold where a motor response could be elicited (12-20 mA, 0.2 ms, single pulse). Presence or 105 absence of a response was noted for each stimulation, which were summated at each site for SSR 106 scoring.

107 Sit-up Test: Participants were instrumented with an oscillometric blood pressure cuff (STBP-780, 108 Colin, Komaki, Japan), and electrocardiogram for impedance estimation (PhysioFlow®, 109 NeuMeDx, Bristol, PA, USA). Participants were instructed to maintain a breathing rate of 15 110 breaths/min, timed to a metronome, to account for respiratory sinus arrhythmia. After five 111 minutes of supine measurement, participants were passively sat up in a reclinable chair to a 112 seated position for 15 minutes. Automated oscillometric blood pressure was recorded every minute, or until participant demonstrated signs of syncope. OH was defined as a drop in systolic 113 blood pressure (SBP) of  $\geq$  20 mmHg, or a drop in diastolic blood pressure (DBP) of  $\geq$  10 mmHg 114 115 [22].

116  $\dot{VO}_2 peak Testing:$  Peak oxygen consumption ( $\dot{VO}_2 peak$ ) was assessed with a graded arm cycling 117 test to volitional exhaustion. After 10 minutes of rest, participants were instructed to cycle at 50 118 rpm on a manually-braked arm ergometer (Monark Rehab Trainer 881E; Monark Exercise AB, 119 Varberg, Sweden) at 0W, 5W, and 10W for three-minute intervals, thereafter increasing the 120 power output by 5W/min for individuals with tetraplegia or 10 W/min for both individuals with paraplegia and able-bodied controls until test termination (cycling rate < 40 rpm). Ventilatory</li>
data were collected with a portable metabolic unit (MetaMax 3B, Cortex, Leipzig, Germany).

123 Skin temperatures: Skin temperatures were measured at 10 different sites with iButton sensors 124 (DS1922T, Maxim Integrated Products, Inc., Sunnyvale, CA, USA): on the forehead and on the left side of the body at the chest, upper back, upper arm, forearm, dorsal hand, lower abdomen, 125 anterior thigh, lateral calf, and dorsal foot. Data were sampled every 10 seconds, which were 126 then averaged every minute for analysis. T<sub>sk</sub> was averaged between the arm, forearm, and hand 127 for upper-limb temperature, and averaged between the thigh, leg, and foot for lower-limb 128 129 temperature. In the absence of SCI-specific T<sub>sk</sub> standards, mean T<sub>sk</sub> was calculated as per 130 Ramanathan 1964 by the following equation [23]:

 $Mean Skin Temperature = (0.3 \times T_{chest}) + (0.3 \times T_{arm}) + (0.2 \times T_{thigh}) + (0.2 \times T_{calf})$ 

Core temperature: Core temperature was measured using both an ingestible telemetric capsule 131 132  $(T_{int})$ , as well as an esophageal probe  $(T_{eso})$ . The telemetric capsule was factory-calibrated and swallowed 8-10 hours prior to arrival for the acute exercise trial (e-Celsius<sup>™</sup>; BodyCap, Paris, 133 France). Data were sampled every 30 seconds by a portable receiver (e-Performance; BodyCap), 134 which were then averaged every minute for analysis. Teso was assessed by thermocouples in a 135 136 polyethylene tube (PE90; Nippon Becton Dickinson Company, Tokyo, Japan), which was 137 inserted through the right nostril and extended to the esophagus at the level of the right atria 138 (0.25\*supine length), the placement of which was confirmed via a cold-water swallow. Immediately after each exercise trial, the esophageal probe was immersed in both room 139 140 temperature and 40°C water for calibration. The delay in temperature elevation was identified between initiation of the test and the first sustained elevation in both  $T_{\text{eso}}$  and  $T_{\text{int}}.$ 141

142 *Heat Storage*: Heat storage during exercise was calculated as per [24] by the following equation:

Heat storage = 
$$((0.8 \times [\Delta T_{eso} \text{ or } \Delta T_{int}]) + (0.2 \times \Delta T_{sk})) \times c_b$$

where  $c_b$  is the specific heat capacity of body tissue (3.49 J·g<sup>-1.o</sup>C<sup>-1</sup>) and  $T_{sk}$  is the mean skin temperature as calculated from above. Heat storage was calculated from rest to each bout of exercise (ending at 15 min and 30 min) for both  $T_{eso}$  and  $T_{int}$ .

*Exercise Trial*: Exercise was performed in a temperature and humidity-controlled room at 25°C and 50% humidity. Exercise intensity was chosen to reflect the most recent physical activity guidelines for individuals with SCI [19]. After instrumentation, participants were instructed to sit quietly for 10 minutes of rest, followed by two 15-minute bouts of exercise at 50%  $\dot{V}O_2$  reserve [25], separated by two minutes of rest. Throughout exercise, heart rate was monitored by singlelead ECG.

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153 Statistics: Statistical analyses were performed using IBM SPSS Statistics (version 20.0.0; IBM 154 Corp., Armonk, NY, USA). Data were visually assessed for normality using histograms and statistically using the Kolgomorov-Smirnov test. Resting characteristics and cardiopulmonary 155 156 responses to exercise were compared between groups with one-way independent ANOVAs. Oneway repeated measures ANOVAs were used to assess the change in T<sub>core</sub> and T<sub>sk</sub> at discrete 157 158 intervals over the exercise bout within each group (i.e., timepoints: rest, 15 min, recovery, 30 159 min), where significant effects were followed by Tukey's HSD post-hoc test. Paired t-tests were 160 used to examine the difference in heat storage between measurement types. Bland-Altman analysis was used to examine the difference between T<sub>int</sub> and T<sub>eso</sub> sensitivity during exercise, 161 162 reporting the mean bias and 95% limits of agreement. Values are reported as mean±SD unless otherwise noted. In all analyses, the level of significance was set at  $\alpha = 0.05$ . 163

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## 166 RESULTS

Participant characteristics and baseline autonomic function are described in Table 1. Autonomic testing revealed that men with tetraplegia had autonomic dysfunction as they presented with an absence of SSR and presence of OH, while men with paraplegia had generally preserved upperlimb SSR responses and no OH. As a methodological control, autonomic testing was also performed on able-bodied men, indicating intact SSR responses and only one case of diastolic OH.

Peak and sub-maximal exercise responses are described in Table 2. There was a graded effect of lesion level on fitness, where men with SCI had lower  $\dot{V}O2$ peak than able-bodied men (P < 0.05). During the acute exercise bout, men with SCI worked at lower power output (P < 0.05) compared to able bodied men, while men with tetraplegia had lower heart rate responses at both 15 and 30 minutes of exercise (both P < 0.01), with no differences between men with paraplegia and able-bodied controls (both P > 0.90).

179 At rest, there were no differences between tetraplegia, paraplegia, or able-bodied men in core temperature as measured by either  $T_{int}$  (P = 0.14) or  $T_{eso}$  (P = 0.35), although  $T_{eso}$  was lower 180 than T<sub>int</sub> by 0.63 °C for all individuals (36.14±0.36 °C vs. 36.76±0.40 °C, P < 0.01). T<sub>int</sub> and T<sub>eso</sub> 181 182 demonstrated similar response patterns to exercise in men with paraplegia and able-bodied men, with both methods increasing by the end of the exercise bout (P < 0.05; Figure 1). T<sub>eso</sub> in able-183 bodied men demonstrated additional sensitivity to temperature changes at the 15-minute and 184 resting timepoints, without concurrent changes in Tint. There were no elevations in either Teso 185 186 (P=0.24) or T<sub>int</sub> (P=0.71), in men with tetraplegia. When all groups were pooled, we observed a delay in temperature elevations, where  $T_{eso}$  began rising 7.2 min earlier than  $T_{int}$  (5.2±1.9 vs. 187

188 12.4 $\pm$ 6.6 min; *P*<0.01). To assess the overall difference between temperature assessment 189 sensitivity during exercise, a Bland-Altman analysis was conducted using the pooled data 190 between the 15- and 30-minute timepoints, separated by group (Figure 2). An average bias of a 191 0.17°C greater T<sub>eso</sub> response was observed, with the 95% limits of agreement ranging from -0.25 192 to 0.59°C.

Average upper limb skin temperature did not significantly change from rest in any group (*P*>0.05; Figure 3), although forearm temperature exhibited a time x group interaction (P<0.01), where temperature decreased in men with tetraplegia throughout the entire exercise bout ( $\Delta$  -1.80°C from rest to 30min; all *P*<0.05). In contrast, lower limb skin temperature demonstrated a main effect for time (*P*<0.01) where temperatures were lower during the exercise bout than during baseline rest for all participants ( $\Delta$  -0.49°C from rest to 30min; all *P* < 0.05; Figure 3).

Heat storage was increased when estimated from  $T_{eso}$  compared to  $T_{int}$  after the first 15 minutes of exercise in both men with paraplegia and able-bodied controls (P < 0.05; Figure 4). After the second bout of exercise, heat storage estimated by  $T_{eso}$  was only greater in men with paraplegia (P = 0.03). Heat storage increased from the first bout to the second bout only in men with paraplegia ( $T_{eso}$  estimate P = 0.03;  $T_{int}$  estimate P < 0.01). There were no differences between heat storage method estimates or heat storage over time in men with tetraplegia (P > 0.05).

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# 207 DISCUSSION

We investigated whether two regional indicators of  $T_{core}$  demonstrate similar responses during normothermic exercise in individuals with different levels of SCI. The primary findings were: 1) amongst able-bodied men and men with paraplegia, both  $T_{eso}$  and  $T_{int}$  were sensitive to  $T_{core}$  elevations during exercise in normothermia, with further  $T_{eso}$  sensitivity in able-bodied men earlier during exercise; and 2) individuals with tetraplegia demonstrated signs of autonomic dysfunction at rest, and had no elevations in  $T_{eso}$  or  $T_{int}$  during 30 minutes of cycling at an intensity reflecting recommended workloads for healthy habitual physical activity. Our results indicate that  $T_{int}$  and  $T_{eso}$  can provide similar estimates of  $T_{core}$  elevation during moderateintensity arm-cycling exercise in men with paraplegia, although our able-bodied data indicates  $T_{eso}$  likely has superior sensitivity to capture temperature information earlier during exercise.

The use of telemetric pills provides a non-invasive estimate of T<sub>core</sub> which can be used for 218 219 field assessments of thermoregulation during exercise in able-bodied individuals [9, 26, 27]. 220 Ambulatory temperature monitoring is particularly important for individuals with SCI who may 221 have temperature dysregulation secondary to autonomic dysfunction and are therefore at elevated risk for heat-related illness. Here, we report similar sensitivity in simultaneously measured Teso 222 and T<sub>int</sub> during upper limb exercise in individuals with paraplegia and able-bodied men. T<sub>eso</sub> 223 224 demonstrated an earlier rise during exercise, demonstrating rapid sensitivity to brief elevations in 225 T<sub>core</sub>. These findings agree with previous reports in able-bodied men that longer exercise 226 durations are required to achieve a 0.1°C change in T<sub>int</sub> compared to T<sub>eso</sub> [28]. Circulating blood 227 from the active limbs likely contributes to the robust temperature sensitivity, as warmed venous 228 return from the working limbs directly impacts temperature conduction through the pulmonary artery to the level of the esophageal probe. Upper-limb exercise also results in greater heat 229 230 production due to poor work efficiency (compared to the lower limbs), as well as less afferent 231 input to the central nervous system resulting in greater heat storage [18]. It has been previously suggested that T<sub>int</sub> might be impacted by the location of the pill in the intestinal tract, and as 232 233 such, there have been recommendations to standardize pill ingestion 6 hours prior to

234 measurement time for optimal intestinal transit time [9]. It is, however, unknown whether these 235 assumptions hold true for individuals with SCI who have slower colon transit time (previously 236 reported as SCI: 0.63±0.33 cm/hr, controls: 2.58±1.20 cm/hr) [29] and autonomically-mediated 237 peristaltic dysfunction [30]. Previous studies in able-bodied men and athletes with SCI have demonstrated minimal differences between T<sub>core</sub> estimations using esophageal, intestinal, or 238 rectal probes during high-intensity exercise [27, 31]. Our findings are likely specific to 239 240 individuals with SCI with poor aerobic fitness performing moderate-intensity exercise, which 241 represents the fitness and activity of the majority of untrained adults with SCI [32]. It is 242 unknown whether exercise in heat or longer duration exercise would diminish observed 243 differences between methods in men with paraplegia, although an exercise duration of 30 244 minutes is a realistic and relevant physiological challenge for most individuals with SCI.

245 While the thermoregulatory responses to prolonged exercise have previously been 246 documented in athletes with SCI [7, 8, 11, 13–16, 31, 33, 34], relatively little is known about the 247 exercise response in untrained adults with SCI, in particular, individuals with tetraplegia. Using a 248 moderate-intensity exercise protocol recommended by national SCI physical activity guidelines [19], we observed increases in T<sub>core</sub> in men with paraplegia and able-bodied controls, but not in 249 individuals with tetraplegia, likely due to lower exercise power output amongst men with 250 251 tetraplegia. The exercise intensity levels in individuals with tetraplegia were likely below a 252 threshold required to generate sufficient heat stress as evidenced by a lack of elevations in upperlimb T<sub>sk</sub> or heat storage during exercise in a normothermic environment. Low cardiorespiratory 253 254 loads at a similar relative exercise intensity were evident by both the lower exercise heart rate as well as moderate RPE scores during the final minute of exercise, likely as a result of autonomic 255 256 dysfunction (i.e., lack of SSR, symptoms of OH) and low cardiorespiratory fitness. Previously,

Webborn *et al.* (2005) demonstrated robust increases in  $T_{int}$  in athletes with tetraplegia in response to an intermittent sprint exercise protocol on an arm ergometer, albeit at four times the power output of our untrained participants. Although our data do not indicate heat stress in men with tetraplegia and poor cardiorespiratory fitness, it remains unknown whether environmental heat stress would impact the effect of exercise on regional elevations in  $T_{core}$  in this population.

262 Telemetric measurement of T<sub>core</sub> in individuals with SCI is important for the dynamic 263 assessment of thermoregulation, a key homeostatic process that may be impaired by autonomic 264 dysfunction following SCI. In our study, we observed similar increases in Teso and Tint during 30 265 minutes of upper-limb exercise in men with paraplegia as well as able-bodied controls. The 266 absence of exercise-induced heat stress in men with tetraplegia, as reflected by a lack of 267 increases in T<sub>core</sub> or T<sub>sk</sub>, was likely due to limitations to aerobic capacity and low power outputs during the acute exercise. Our results indicate that  $T_{int}$  and  $T_{eso}$  can provide similar estimates of 268 T<sub>core</sub> elevation during moderate-intensity arm-cycling exercise in men with paraplegia. 269

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### 271 DATA ARCHIVING

The datasets generated during the current study are available from the corresponding author onreasonable request.

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279 STATEMENT OF ETHICS

All methods and procedures were approved by the Medical Ethical Committee of Wakayama Medical University (#2076), adhering to the Declaration of Helsinki except for registration in a database. All testing was completed in the Wakayama Medical University Genki Development Institute (Wakayama, Japan). All participants gave verbal and written consent prior to participation in this study. We certify that all applicable institutional and governmental regulations concerning the ethical use of human volunteers were following during the course of the research.

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# 288 CONFLICTS OF INTEREST

289 The authors declare that they have no conflicts of interest.

290

# 291 AUTHOR CONTRIBUTIONS

JA was responsible for designing the protocol, conducting the experiments, extracting andanalyzing the data, interpreting the results, as well as drafting the final manuscript.

294 YK was responsible for designing the protocol, conducting the experiments, extracting the data,

interpreting the results, as well as providing feedback on the final manuscript.

VG was responsible for designing the protocol, interpreting the study findings, as well asproviding feedback on the final manuscript.

CL was responsible for designing the protocol, interpreting the study findings, as well asproviding feedback on the final manuscript.

300 MM was responsible for designing the protocol, interpreting the study findings, as well as301 drafting the final manuscript.

302 YM was responsible for conducting the experiments, extracting the data, as well as providing303 feedback on the final manuscript.

FT was responsible for designing the protocol, interpreting the study findings, as well asproviding feedback on the final manuscript.

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**396** FIGURES

- FIGURE 1. Changes in core temperature ( $T_{core}$ ) during the exercise bout measured by esophageal (open circles;  $T_{eso}$ ) and intestinal (closed circles;  $T_{int}$ ) probes. Shaded areas indicate rest periods. Plotted points are mean+SEM. \**P*<0.05 different from resting temperature.
- FIGURE 2. Bland-Altman analysis indicating systematically larger changes in  $T_{eso}$  compared to T<sub>int</sub> during exercise in men with tetraplegia, men with paraplegia, and able-bodied men. The difference between changes in temperature ( $T_{eso} - T_{int}$ ) is plotted against the range of average temperature changes, with the mean bias (0.17 °C) and 95% limits of agreement (-0.25 to 0.59
- 404 °C) indicated by dashed lines.

FIGURE 3. Changes in upper (top) and lower (bottom) skin temperature ( $T_{sk}$ ) during the exercise bout in individuals with tetraplegia (open triangles), paraplegia (open circles), and able-bodied men (closed circles). Shaded areas indicate rest periods. Plotted points are mean+SEM. †*P*<0.05 all groups different from resting temperature.

FIGURE 4. Differences in heat storage estimated from esophogeal ( $T_{eso}$ ) and intestinal ( $T_{int}$ ) temperatures after the first and second bouts of exercise in A) men with tetraplegia, B) men with paraplegia, and C) able-bodied men. Mean±SE. \**P*<0.05 different *vs*.  $T_{eso}$ ; +*P*<0.05 different *vs*. 15min.

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414 TABLES

415 Table 1. Participant characteristics.

416 Table 2. Cardiorespiratory outcomes during graded exercise test and exercise bout (mean±SD).







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Time (min)

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0

-10

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Table 1. Participant characteristics.
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Outcome	Tetraplegia	Paraplegia	Able-bodied
	(n=7)	(n=5)	(n=5)
Age (yrs)	38±12	53±7	37±13
Height (m)	$1.72 \pm 0.07$	$1.67 \pm 0.04$	$1.69 \pm 0.05$
Body mass (kg)	57±14	55±7	64±14
Injury level range	C4 - C7	T4 <b>-</b> T11	
AIS			
А	4	5	
В	3	0	
SSR			
Radial Stimulation	1/0	4/0	5/5
(Hand/Foot)	1/0	4/0	5/5
Tibial Stimulation	0/0	0/0	5/5
(Hand/Foot)	0/0	0/0	5/5
Supra-orbital Stimulation	0/0	3/0	5/5
(Hand/Foot)	0/0	5/0	5/5
Cases of OH from SUT	6/7	0/5	1/5
(/total)	0/7	0/0	115

 $\overline{AIS} = American Spinal Injury Association Impairment Scale; = OH = orthostatic hypotension; RER=respiratory exchange ratio; RPE = ratings of perceived exertion; SSR = sympathetic skin response; SUT = sit-up test; V<sub>E</sub> = ventilatory rate. SSR are group averages, out of a maximum of five stimulations per limb. Values are mean±SD unless otherwise noted.$ 

Outcome	Tetraplegia	Paraplegia	Able-bodied
Graded Exercise Test			
Resting $\dot{V}O_2(L \cdot min^{-1})$	0.19±0.03+	$0.21 \pm 0.02$	$0.25 \pm 0.04$
Peak power output (W)	17±11*†	56±20	77±11
$\dot{V}O_2$ peak (L·min <sup>-1</sup> )	0.56±0.10*†	1.11±0.31+	$1.64 \pm 0.28$
Peak heart rate (bpm)	97±11*†	161±19	$168 \pm 22$
Peak $V_E(L \cdot min^{-1})$	26.8±5.2*+	49.4±13.0†	$80.0 \pm 18.8$
Peak RER	$1.05 \pm 0.05 \pm$	$1.22 \pm 0.20$	$1.39 \pm 0.22$
Exercise Responses at 50%	, D		
VO <sub>2</sub> peak			
Power output (W)	7±2*†	25±11+	43±7
Power output (%W <sub>peak</sub> )	$47 \pm 17$	$43 \pm 7$	55±3
@15 min Heart rate (bpm)	75±12*†	$131 \pm 14$	$127 \pm 18$
@15 min RPE	$13 \pm 2$	15±1	15±1
$\overline{a}$ 30 min Heart rate (bpm)	77±6*†	139±19	$138 \pm 21$
@30 min RPE	$14 \pm 2$	16±1	16±1

Table 2. Cardiorespiratory outcomes during graded exercise test and exercise bout (mean±SD).

 $\overrightarrow{RPE}$  = ratings of perceived exertion;  $V_E$  = ventilatory rate; RER=respiratory exchange ratio; \**P*<0.05 vs paraplegia; †*P*<0.05 vs able-bodied.