Prefrontal Cortex Haemodynamics and Affective Responses during Exercise: A Multi-Channel Near Infrared Spectroscopy Study

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Abstract

The dose-response effects of the intensity of exercise upon the potential regulation (through top-down processes) of affective (pleasure-displeasure) responses in the prefrontal cortex during an incremental exercise protocol have not been explored. This study examined the functional capacity of the prefrontal cortex (reflected by haemodynamics using near infrared spectroscopy) and affective responses during exercise at different intensities. Participants completed an incremental cycling exercise test to exhaustion. Changes (Δ) in oxygenation (O2Hb), deoxygenation (HHb), blood volume (tHb) and haemoglobin difference (HbDiff) were measured from bilateral dorsal and ventral prefrontal areas. Affective responses were measured every minute during exercise. Data were extracted at intensities standardised to: below ventilatory threshold, at ventilatory threshold, respiratory compensation point and the end of exercise. During exercise at intensities from ventilatory threshold to respiratory compensation point, ΔO2Hb, ΔHbDiff and ΔtHb were greater in mostly ventral than dorsal regions. From the respiratory compensation point to the end of exercise, ΔO2Hb remained stable and ΔHbDiff declined in dorsal regions. As the intensity increased above the ventilatory threshold, inverse associations between affective responses and oxygenation in (a) all regions of the left hemisphere and (b) lateral (dorsal and ventral) regions followed by the midline (ventral) region in the right hemisphere were observed. Differential activation patterns occur within the prefrontal cortex and are associated with affective responses during cycling exercise.

Introduction

A dose-response relationship exists between the intensity of exercise and the pleasure-displeasure individuals feel [1], [2]. These affective responses, which underlie core affect [3], are linked to future exercise participation [4–6]. Recent guidelines from the American College of Sports Medicine (2013) recognise the importance of affective responses in exercise prescription and the detrimental impact of negative affect on future exercise participation [7]. This recognition provides the rationale to investigate a proposed neural basis of affective responses during exercise, particularly the role of the prefrontal cortex (PFC).

The dose-response relationship between the intensity of exercise and affective responses emerged following recommendations to standardise exercise intensities to individual gas exchange thresholds; the ventilatory threshold (VT) and the respiratory compensation point (RCP) [8]. The VT (approximately 60–70% peak oxygen uptake [VO2peak]) indicates the point of transition from aerobic to anaerobic metabolism and the RCP (approximately 80–90% VO2peak) indicates the point at which a physiological steady state can no longer be maintained [9]. At intensities of exercise substantially below VT, individuals report mostly positive (pleasant) affective responses [10], [11]. However at intensities above VT (proximal to RCP and exhaustion), individuals report uniformly negative (unpleasant) affective responses [12–15].

The dose-response relationship is explained by a theoretical framework, called the dual-mode model [16], [17]. According to the model, affective responses to exercise are regulated by the PFC and subcortical areas of the brain (which receive sensory input from the body). It is proposed that activation in the PFC serves to maintain positive (or actively avoid negative) affective responses during exercise. However, as the intensity increases (i.e. above VT) the functional capacity of the PFC becomes challenged by the intensified sensory (interoceptive) input from the body. Therefore, a deregulation in the PFC is proposed to occur, resulting in a negative-affective response, mainly driven by subcortical structures (i.e. the amygdala). Deregulation of the PFC would allow a shift of metabolic resources to areas of the brain directly associated with the activity [18], [19]. The basis of the dual-mode model is that the PFC provides top-down regulation (i.e. inhibition of subcortical structures) of affective responses through cognitive control processes. However at intensities of exercise above VT, deregulation of the PFC may occur and the functional capacity of the PFC becomes limited.

The PFC is involved in executive function and cognitive control processes [20]. Cognitive control describes an individual’s ability to facilitate or inhibit stimuli at a neural level, which may influence...
their behavioural (and affective) response [21]. Two main outcomes of affective (and emotional) regulation are (a) inhibition or suppression of stimuli through attentional processes, which serve to focus towards or distract away from stimuli and limit expressive action, and (b) cognitive reappraisal (defined by Lazarus [22]), which involves manipulating the meaning of stimuli in relation to current goals which neutralise negative experiences and therefore influence the on-going emotional response [21]. Evidence from neuroimaging studies has indicated differential activation patterns in areas of the PFC during cognitive control tasks in the presence of negative affective stimuli. For example, activation in the dorsolateral and ventromedial PFC is increased and is associated with reduced self-reported negative affect [23].

Activation in the right dorsolateral PFC is increased in attempts to inhibit [24] and suppress [20, 25], emotional responses. Cognitive control has been shown to induce greater activation in left-lateralised areas of the PFC in the presence of positive stimuli [26], [27], and greater activation in right-lateralised areas of the PFC in the presence of negative stimuli [27]. Further, neuroimaging studies have shown that activity in areas of the PFC induced by attempts of active cognitive coping with a negative affective stimulus, are associated with reduced activity in the amygdala [26–30].

Prefrontal lateralisation of affective (positive-negative) processing is related to the approach-withdrawal motivational system described by the valence asymmetry hypothesis [31], [32]. The approach system is associated with positive affect, which drives goal-orientation and behaviours towards something [22]. In opposition, the withdrawal system is associated with negative affect and aversion, which drives behaviours away from something. Studies have provided neurophysiological evidence of increased left anterior activation and approach-related positive affect and increased right anterior activation and withdrawal-related negative affect within the frontal cortex using electroencephalography [33–35], functional magnetic resonance imaging [36] and near infrared spectroscopy (NIRS) [37]. In exercise studies, an association between frontal cortex asymmetry and affective responses has previously been reported [38–40]. However, in line with the dual-mode model, the functional capacity of the PFC to regulate affective responses, presumably through cognitive control (i.e. appraisal, inhibition and/or suppression) and lateralised processes during exercise intensities relative to the VT and RCP has not been explored.

Advances in NIRS allow measures of cerebral haemodynamics to be taken during exercise. Optodes (light emitters and detectors) placed over the cortex indicate the change (A) in near-infrared light (due to absorption) as it passes through the underlying tissue and provides a measure of oxy-(O₂Hb) and deoxy-haemoglobin (HHb) concentration [41]. Cerebral ΔO₂Hb and ΔHHb reflects functional neuronal activation [42], [43]. Several studies have shown that cerebral oxygenation (ΔO₂Hb) in frontal areas is increased during sub-maximal exercise however it is reduced during exercise at near maximal exercise intensities [42], [44–47]. The interpretation of these findings is that a reduced supply of oxygen in frontal areas limits frontal cortex activation and contributes to the cessation of exercise. A meta-analysis by Rooks, Thom, McCully & Dishman [48] summarised the cerebral haemodynamic response during incremental exercise. The authors showed that at sub-maximal intensities (≥60% VO₂peak, around the VT) increases in cerebral oxygenation (ΔO₂Hb; blood flow), deoxygenation (ΔHHb) and blood volume (total haemoglobin [ΔHb]; the sum of ΔO₂Hb and ΔHHb) were observed. As exercise increased towards maximal intensities (VO₂peak) cerebral oxygenation (ΔO₂Hb) and blood volume (ΔHb) were lower, with a continued rise in deoxygenation (ΔHHb). These findings indicate that a reduced supply of oxygen (ΔO₂Hb) compared to relative demand (ΔHHb) occurs during exercise, therefore PFC activation is limited. Measures of cerebral haemodynamics using NIRS during incremental exercise are generally limited to single (as opposed to bilateral or multiple-) areas of the PFC [41], [49]. However, multi-channel NIRS has been used to examine cerebral haemodynamics from frontal and motor areas of the cortex, which have shown greater deoxygenation in frontal compared to motor areas at near maximal exercise intensities [49]. Yet no comparison of NIRS measures recorded from multiple areas (dorsal and ventral) of the PFC during incremental exercise is available. Ekkekakis [50] proposed that variations in the haemodynamic response of the PFC may represent individualised neuronal activity. Therefore measures of the haemodynamic response from multiple areas of the PFC could potentially indicate functional activation (i.e. top-down regulatory [cognitive control] processes) due to affective processing.

According to the dual-mode model, a reduction in oxygenation (ΔO₂Hb) in the presence of increased deoxygenation (ΔHHb) might indicate reduced neuronal activation and therefore a reduced functional capacity of the PFC, which may influence the onset of negative affective responses. Consistent with the dual-mode model, the transient ‘hypofrontality’ hypothesis [18], [19], states that the capacity of the PFC to maintain cognitive and affective processes is challenged during exercise due to a redistribution of metabolic resources from cortical to subcortical areas of the brain. During exercise, global cerebral blood volume remains relatively constant. However there is a shift in resources (oxygen delivery, metabolite removal) [44] or greater oxygen consumption [49] between areas required for cognitive, emotional, sensory and motor control. Regional activation in the brain comes at the expense of other regions [20]. Dietrich [18], [19], describes a five-stage hierarchy of consciousness of which the first to ‘deregulate’ are higher-order processes (in the dorsolateral PFC); the second, self-representation and emotional processes (in the ventromedial PFC); the third, memory, learning and basic emotions (the limbic system including the amygdala); the fourth, sensory processing (in the thalamus); and finally, arousal processes (in the brainstem). It is proposed that affective responses are largely influenced by the functional capacity of the PFC to maintain activation in competition with sensory (interoceptive) inputs from the body during exercise as the intensity becomes increasingly challenging.

Due to the limited depth resolution of NIRS only activation from surface regions of the cortex can be measured. No measures of activation in subcortical areas of the brain (i.e. amygdala and thalamus) can be recorded. To the authors’ current knowledge, haemodynamics in regions of the lateral PFC, such as dorsal and ventral areas, using multi-channel NIRS during incremental exercise has not been examined. In line with the dual-mode model and evidence of cognitive control and lateralised affective processes of the PFC, it is proposed that differential activation patterns would occur as the intensity of exercise increases. Differential activation of the PFC is theorised to coincide with reports of negative affective responses. It is further proposed that variability in the haemodynamic response highlighted by Rooks et al. [48], could be attributed to measures recorded from multiple regions of the PFC (as opposed to single uni- or bilateral measures not considered by Rooks et al.) during exercise standardised to metabolic markers.

Therefore, the present study investigates the functional capacity (activation in sub regions) of the PFC (reflected by haemodynamics using multi-channel NIRS) and affective responses during exercise...
at intensities standardised to below VT, VT, RCP and the end of exercise (recommended by Ekkekakis [50] and Rooks et al. [48]).

Three hypotheses were examined: 1) activation in the PFC is different in sub regions (dorsal and ventral area) at intensities of exercise above the VT; 2) affective responses are more positive at exercise intensities corresponding to below VT and more negative at exercise intensities corresponding to RCP and the end of exercise; and 3) activation in sub regions of the PFC is associated with affective responses during exercise.

Methods

Ethics Statement

This study was approved by The University of South Australia Ethics Committee. Participants read and signed an informed consent form prior to participation in the study.

Participants

Healthy participants (n = 25; male = 13; mean ± SD; age 25.6 ± 3.4 years; height 174.8 ± 6.8 cm; body mass 71.8 ± 8.3 kg; body mass index 23.3 ± 2.2; VO2peak 41.8 ± 5.2 ml·kg⁻¹·min⁻¹) were recruited. The volunteers completed the Adult Pre-exercise Screening System (APSS [51]) prior to the study to ensure suitability for participation. All participants were non-smokers and recreationally active. Participants reported partaking in intensities standardised to below VT, VT, RCP and the end of exercise; and 3) activation in sub regions of the PFC is associated with affective responses during exercise.

Statistical Analyses

The independent variables were Region (1–8; dorsal 1, 3, 5 and 7; ventral 2, 4, 6 and 8) and Time (corresponding to pre-exercise and intensities below VT, VT, RCP and End). The dependent variables were cerebral haemodynamics (ΔO2Hb, ΔHHb, ΔHb and ΔHbDiff) and affective responses. To examine hypothesis 1, a series of Region (3; 1–8) by Time (4; below VT, VT, RCP, End) analysis of variance (ANOVA) with repeated measures were conducted for each of the cerebral haemodynamic variables (ΔO2Hb, ΔHHb, ΔHb and ΔHbDiff). To examine hypothesis 2, a one-way ANOVA with repeated measures for Time (5; pre-exercise, and at time points corresponding to below VT, VT, RCP and End) was conducted for affective responses. Finally, to examine hypothesis 3, Pearson’s bivariate correlation analyses were conducted to examine the

Procedures

The study required individuals to visit the exercise physiology laboratory (temperature ~24°C, and relative humidity ~40%) on one occasion. Upon arrival, participants completed the informed consent form and exercise testing questionnaire and initial assessments were taken (age, height and weight). The procedures for the exercise tests were explained and a description of the Feeling Scale was provided. Participants were seated on a recumbent cycle ergometer (Lode Corival, Groningen, the Netherlands) and the NIRS optodes were carefully positioned. A facemask was fitted to measure metabolic data via online gas analysis (Cortex Metalizer 3B, Biophysik, Leipzig, Germany). The participants then completed an incremental [20 Watts/min; pedal cadence 70 rpm] cycling exercise test to exhaustion. The end of the test was determined by volitional cessation of exercise or failure to maintain a pedal cadence of 70 rpm despite strong verbal encouragement. Cerebral haemodynamic responses and expired gases were measured continuously and affective responses were recorded pre-, during (every minute) and at the end of exercise (VO2peak). The achievement of VO2peak was verified by a) a peak or plateau in oxygen consumption (changes <2 ml·kg⁻¹·min⁻¹) with increasing workload; and b) a respiratory exchange ratio of at least 1.10.

Data Analyses

The VO2peak was determined by the highest 30 second average of oxygen uptake (VO2 ml·kg⁻¹·min⁻¹). The VT was determined using the three-method procedure proposed by Gaskell et al. [54]. This involved agreement in the visual determination of VT using: a) ventilatory equivalent method; b) Excess CO2 method; and c) modified V-slope method. The RCP was determined accordingly to Beaver et al. [55]. This involved agreement in the visual determination of RCP between plots of a) minute ventilation over CO2 production; and b) ventilatory equivalent of CO2 over time. The determination of gas exchange thresholds; VT and RCP were agreed upon by two raters. No large differences between the raters were observed. Participants’ mean VT and RCP occurred at 68±3% and 90±2% VO2peak, respectively.

A 30 second baseline measure of cerebral haemodynamics (ΔO2Hb, ΔHHb, ΔHb and ΔHbDiff) was recorded prior to exercise. Data were exported every 10 seconds and normalised to express the magnitude of changes from the baseline period (arbitrarily defined as 0 μM) at the start of exercise. A 20 second average of cerebral haemodynamic variables was extracted at time points corresponding to intensities of exercise; 80% of VT (below VT), VT, RCP and end of exercise (End). Corresponding affective responses were selected at time points corresponding to each of the intensities of exercise.
relationships between cerebral oxygenation ($\Delta O_2$Hb) and affective responses during exercise. All statistical analysis was performed using SPSS v. 18.0 (IBM Corp., Armonk, NY, USA). Data were assessed for normality using Shapiro-Wilk tests. Greenhouse Geisser corrections were applied if the assumption of sphericity was not met and significant main and interaction effects ($p<.05$) were followed up using repeated measures contrasts. Effect sizes associated with $F$ statistics were expressed as eta squared ($\eta^2$) defined as small (.01), medium (.06) and large (.14) [56]. Values are mean ± SD unless otherwise stated.

Results

Participants’ physiological data (VO2) at each time point corresponding to each of the intensities of exercise were: below VT 22.8±2.9; at VT 28.5±3.7; RCP 37.7±4.6; and End 41.8±5.2 ml·kg⁻¹·min⁻¹. The mean duration of the exercise test was 641±126 seconds.

Cerebral Haemodynamics (Hypothesis 1)

**Cerebral $\Delta O_2$Hb.** Significant main effects of Region, $F(4, 105) = 3.67, p = .006, \eta^2 = .03$, and Time, $F(1, 27) = 16.21, p < .001, \eta^2 = .37$, and a Region by Time, $F(5, 125) = 4.54, p < .001, \eta^2 = .16$, were recorded for $\Delta O_2$Hb. Follow up tests indicated that $\Delta O_2$Hb was greater in Regions 2, 4, 6, 8 (ventral) than Regions 1, 3, 5, 7 (dorsal) (Table 1). Cerebral $\Delta O_2$Hb increased from below VT ($3.5±2.6$) to VT ($13.3±2.1$) and VT to RCP ($4.6±2.5$), but remained stable RCP to End ($4.9±5.7$). The two factor interaction indicated that from RCP to End, $\Delta O_2$Hb increased in Region 4 and declined in Region 1 (Figure 2A).

**Cerebral $\Delta$HHb.** A significant main effect of Time, $F(1, 27) = 77.54, p < .001, \eta^2 = .44$, was recorded for $\Delta$HHb. Follow up tests indicated that $\Delta$HHb remained stable from below VT ($7.0±2.7$) to VT ($8.7±2.6$), then increased from VT to RCP ($2.5±1.5$) and RCP to End ($4.3±2.0$).

**Cerebral $\Delta$HbDiff.** A significant main effect of Time, $F(1, 27) = 33.71, p < .001, \eta^2 = .42$, and a Region by Time, $F(5, 125) = 2.66, p = .024, \eta^2 = .01$, interaction were recorded for $\Delta$HbDiff. Follow up tests indicated that $\Delta$HbDiff increased from below VT ($1.0±1.7$), to VT ($2.1±2.1$), RCP ($7.1±5.7$) and End ($9.2±6.6$). The two factor interaction indicated that from RCP to End, $\Delta$HbDiff increased in Region 4 and remained stable in Region 1 (Figure 2B). Cerebral $\Delta$HbDiff was greater in Regions 2, 4, 6, 8 (ventral) than Regions 1, 3, 5, 7 (dorsal) (Table 1). The two factor interaction indicated that from VT to RCP and RCP to End, $\Delta$HbDiff increased and was greater in Regions 4 and 6, respectively, than Regions 1, 3, 5 and 7 (dorsal) (Figure 2C).

Affective Responses (Hypothesis 2)

A significant main effect of Time, $F(2, 39) = 68.63, p < .001, \eta^2 = .74$, was recorded for affective responses. Affective responses significantly declined from pre-exercise ($2.4±1.1$) to below VT ($1.7±1.0$), remained stable until VT ($1.5±1.0$) and significantly declined from VT to RCP ($−.8±1.8$) and RCP to End ($−2.0±1.8$).

Table 1. Oxygenation ($\Delta O_2$Hb) and haemoglobin difference ($\Delta$HbDiff; µM) by Region (1 to 8).

|                  | Right hemisphere | Left hemisphere |
|------------------|------------------|-----------------|
|                  | 1                | 3               | 5                | 7                |
| $\Delta O_2$Hb  | 1.51±3.86        | 2.35±4.45       | 2.22±3.38        | 2.72±3.09        |
| $\Delta$HbDiff  | −.46±4.31        | −.09±4.83       | −.09±3.37        | 0.53±3.18        |
|                  |                  | 2               | 4               | 6               | 8               |
| $\Delta O_2$Hb* | 1.21±3.79        | 3.43±3.65       | 3.46±3.33        | 3.45±3.68        |
| $\Delta$HbDiff* | 0.99±4.31        | 1.52±3.93       | 1.82±3.43        | 1.29±3.86        |

Mean ± SD. *indicates significantly greater than other regions, $p < .05$.

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Associations between Cerebral Haemodynamics and Affective Responses (Hypothesis 3)

Below VT and at VT, no significant relationships were shown between affective responses and $\Delta O_2Hb$ in any region of the PFC. However, at RCP, significant inverse relationships were shown between affective responses and $\Delta O_2Hb$ in Regions 1 and 2 in the right hemisphere; and in all Regions (3-8) in the left hemisphere. At End, significant inverse relationships were shown between affective responses and $\Delta O_2Hb$ in Region 4 in the right hemisphere; and in all Regions (3-8) in the left hemisphere (Table 2).

Discussion

The present study investigated the functional capacity (activation in 8 sub regions in the left and right hemispheres) of the PFC using multi-channel NIRS during incremental exercise to volitional exhaustion. The pattern of the cerebral haemodynamic and affective responses were examined at intensities of exercise standardised to individual gas exchange thresholds [48], [50], and data were extracted at time points corresponding to below VT, VT, RCP and the end of exercise. In line with the dual-mode model [16], [17], and evidence of cognitive control and lateralized processes, a reduced functional capacity or deregulation may occur in different regions of the PFC during exercise at intensities of exercise above VT, which may explain the regulation of affective responses during exercise.

Hypothesis 1: Cerebral Haemodynamic Responses

The results support hypothesis 1 with differential haemodynamic responses dependent on the level of exercise intensity. At intensities of exercise below VT, deoxygenation ($\Delta HHb$) remained stable whereas oxygenation ($\Delta O_2Hb$) increased; resulting in an increase in blood volume ($\Delta tHb$) indicating activation (increased $\Delta HHb$). However, the dorsal regions showed increased deoxygenation ($\Delta HHb$) increased in all regions, resulting in an increase in blood volume ($\Delta tHb$) indicating activation (increased $\Delta HHb$). However, greater oxygenation ($\Delta O_2Hb$) and activation ($\Delta HHb$) were shown in ventral regions (2, 4, 6, 8) than all dorsal regions (1, 3, 5, 7).

At intensities of exercise from RCP to the end of exercise, oxygenation ($\Delta O_2Hb$) continued to increase in ventral regions (greatest in region 4 followed by regions 6 and 8), but remained stable or declined in dorsal regions (1, 3, 5, 7). Deoxygenation ($\Delta HHb$) continued to increase in all regions and blood volume ($\Delta tHb$) increased in most regions except the most lateral area in the right dorsolateral region of the PFC (region 1). The increase in blood volume (sum of $\Delta O_2Hb$ and $\Delta HHb$) indicates activation in ventral regions and reduced activation in the dorsal regions at these intensities. This is because the ventral regions showed an increase in oxygenation ($\Delta O_2Hb$) coupled with an increase in deoxygenation ($\Delta HHb$); therefore adequate oxygen supply for demand; increased $\Delta HHb$). However, the dorsal regions showed a decline or plateau in oxygenation ($\Delta O_2Hb$) coupled with an increased deoxygenation ($\Delta HHb$); therefore lack of oxygen supply to demand; decreased $\Delta HHb$). Activation in bilateral ventral regions located closest to the midline (regions 4 and 6; the anterior PFC, approximately BA 10) indicated the greatest activation at the end of exercise. The overall pattern of the cerebral haemodynamic response during incremental exercise is consistent with previous studies [48]; however the different responses between the multiple regions of the PFC measured is a novel finding.

The dual-mode model explains that exercising at intensities below VT would not induce sufficient sensory input from the body to challenge the individual, and therefore the capacity of the PFC. However, as the intensity of exercise increases, sensory input from the body would challenge the capacity of the PFC to maintain activation. According the transient hypofrontality hypothesis [18], [19], the first stage of deregulation would occur in the dorsolateral areas of the PFC and the second stage of deregulation would occur in the ventromedial areas of the PFC. In this study during exercise above the VT, reduced activation was observed in the left and right dorsolateral frontal areas (corresponding to regions 1, 3, 5, 7) and greater activation in ventral regions. As the intensity increased, increased deoxygenation ($\Delta HHb$) increased in all regions, resulting in an increase in blood volume ($\Delta tHb$) indicating activation (increased $\Delta HHb$). However, greater oxygenation ($\Delta O_2Hb$) and activation ($\Delta HHb$) were shown in ventral regions (2, 4, 6, 8) than all dorsal regions (1, 3, 5, 7).

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Table 2. Associations ($r$) between cerebral oxygenation ($\Delta O_2Hb$) and affective responses.

| Region | Time (corresponding to intensities of exercise) | below VT | VT | RCP | End |
|--------|-----------------------------------------------|---------|----|-----|-----|
| 1      |                                               | -.17    | -.09| -.55**| -.38 |
| 2      |                                               | -.12    | -.08| -.53**| -.33 |
| 3      |                                               | -.06    | -.08| -.29 | -.30 |
| 4      |                                               | -.00    | -.11| -.36 | -.46* |
| 5      |                                               | -.30    | -.17| -.50*| -.52*|
| 6      |                                               | -.07    | -.09| -.59**| -.56*|
| 7      |                                               | -.09    | -.08| -.62**| -.55*|
| 8      |                                               | -.04    | -.11| -.53**| -.46*|

*p<.05, **p<.01.

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increased, greater activation was observed in ventral regions (4 and 6) indicating the potential shift in metabolic resources from dorsal regions (regions 1, 3, 5 and 7; stage one) towards ventromedial areas (located beneath regions 4 and 6; stage two). Therefore, during incremental exercise the cerebral haemodynamic response appeared to follow a top-down (dorsal-ventral) direction from VT to RCP, and outside-in (lateral to midline) direction from RCP to the end of exercise.

**Hypothesis 2: Affective Responses**

The results support hypothesis 2 with different affective responses dependent on the level of exercise intensity. Despite a decline in affective responses from pre-exercise to intensities below VT, affective responses remained positive until intensities above VT. However, at RCP and the end of exercise affective responses were uniformly negative. This pattern of affective responses is consistent with the dose-response relationship of the intensity of exercise [1]. According to the dual-mode model, sensory input from the body is intensifies (above VT) and the functional capacity of the PFC is challenged, presumably due to increased activity in subcortical areas of the brain. Maintained activation in the PFC would serve to maintain a more positive affective response through cognitive control processes. Prior work has shown that maintained PFC activation through the use of a cognitive strategy results in a more positive affective response during exercise at intensities above the VT, but not the RCP [57].

**Hypothesis 3: Associations between Cerebral Haemodynamic and Affective Responses**

The results support hypothesis 3 with associations between haemodynamic and affective responses dependent on the level of exercise intensity. No associations between cerebral oxygenation and affective responses were shown at intensities below VT and at VT. From pre-exercise to VT, the PFC appeared to be adequately saturated (sufficient oxygen supply [ΔO2Hb] vs. demand [ΔHHb]) therefore greater ΔHbDiff (and a positive affective response was reported). It could be expected that at these intensities, PFC activation would be associated with positive affective responses if cognitive control processes influenced the regulation of affective responses. However, the correlations between cerebral oxygenation and affective responses are very weak (see Table 2). It is proposed that at these intensities sensory input from the body is not sufficient enough to induce negative affect and threaten the capacity of the PFC; therefore cognitive control processes may not be actively engaged.

At intensities above VT (i.e. RCP and End), cerebral oxygenation (ΔO2Hb) in both the left and right hemispheres was inversely associated with affective responses. In the left hemisphere at intensities of exercise at the RCP and the end of exercise, greater cerebral oxygenation (ΔO2Hb) in all regions was associated with a more negative affective response. Greater activation in the left hemisphere in the presence of a negative affective response may reflect increased cognitive control processes towards a less negative/more positive affective response [21], [23].

In the right hemisphere at intensities of exercise at RCP, a more negative affective response was associated with greater cerebral oxygenation in the most lateral (ventral-dorsal) regions (1 and 2), but not those closest to the midline (3 and 4). However, at the end of exercise, only the ventral region closest to the midline (region 4) showed an inverse relationship between cerebral oxygenation (ΔO2Hb) and affective responses. This region indicated a significant increase in oxygenation (ΔO2Hb) between RCP and the end of exercise. Although some caution should be taken in the interpretation of these moderate linear correlations, the association between oxygenation (ΔO2Hb) in both hemispheres, but particularly in the right hemisphere at intensities above VT is novel. Lateralised areas of the PFC have been shown to be involved in the cognitive control of negative affect [24], [27]. Furthermore, relative right hemisphere activation is associated with negative affect [31], [32], which may indicate processes related to active avoidance of negative affect. These findings provide preliminary indications that activation in sub regions of the PFC may be involved in the regulation of negative affective responses during exercise at intensities above VT. It is proposed that increased oxygenation (i.e. blood flow) in sub regions of the PFC is induced by attempts of active cognitive coping with a negative affective stimulus, which would serve to reduce activity in the amygdala [23], [26], [28–30].

**Limitations and Future Directions**

The interpretation of these results is predominantly speculative for two reasons. Firstly, cognitive control processes proposed to regulate affective responses were not measured during exercise. Secondly, no measures of subcortical areas of the brain (i.e. the amygdala) were recorded. However, this study did measure functional capacity of the PFC (reflected through haemodynamics) to host such cognitive control processes to provide top-down regulation of subcortical areas. An extension of this study would be to employ a cognitive control paradigm to determine the effects upon PFC haemodynamics and affective responses in a control and exercise condition. An understanding of the role of the PFC during exercise will provide some confirmation of a neural basis of the dose-response relationship of affective responses to the intensity of exercise.

The different patterns of the cerebral haemodynamic response in sub regions of the PFC as the intensity of exercise increased may explain some of the variability previously reported. Rooks et al. [48] indicated large variability in cerebral oxygenation (ΔO2Hb) at sub maximal (>60% VO2peak) and near maximal (VO2peak) intensities. The authors attributed some of this variability to the individual’s training status (VO2peak), reported by Rooks et al. [48] to have an average VO2peak of 40.3±10.4 ml·kg⁻¹·min⁻¹ similar to that reported in the present study (41.8±5.2 ml·kg⁻¹·min⁻¹). The participants in the current study were all physically active however no measure of training status was assessed. As recommended by the authors, the intensities of exercise in this study were standardized to individual metabolic markers. Using multi-channel NIRS, altered patterns of the cerebral haemodynamic response were observed in the different sub regions measured (i.e. dorsal regions indicate less activation [reduced ΔO2Hb] than ventral regions) and between hemispheres (i.e. greater ΔO2Hb in region 4) at intensities above VT (68% VO2peak). These findings indicate that variability in the cerebral haemodynamic response may be due to the regions (i.e. the placement of optodes) from which the haemodynamic response is measured. This was not considered by Rooks et al. [48]. Although, the authors did examine the distance between optodes (< or >4 cm), no effect upon the cerebral haemodynamic response was indicated. Prior studies examining the influence of cerebral oxygenation during incremental exercise have measured cerebral and muscle oxygenation under conditions such as hypoxia [49] and hyperoxia [58]. Measures of cognitive and affective processes in these types of studies are warranted to encompass additional factors associated with limited cerebral oxygenation during exercise, aside from those related to motor control. Future research should consider methodological limitations associated with using NIRS during exercise such as approximate optode placement (within-between subjects and sessions) and sampling procedures as highlighted.
previously [46,49] and population characteristics (i.e. fitness and
training status). Although, the spatial resolution of NIRS is poor
compared to other neuroimaging techniques (such as magnetic
resonance imaging), NIRS is less influenced by movement artifact
making it suitable for use during exercise.

Conclusion

The findings of the current study indicate the potentially limited
functional capacity of the PFC during exercise at intensities above
the VT. Some support for the dual-mode model as a potential
neural basis of the dose-response effects of the intensity of exercise
and affective responses has been provided. The study has indicated
that prefrontal haemodynamics (which reflect functional activa-
tion) during exercise shifts in a top-down manner from dorsal to
ventral regions, with greatest activation in anterior regions (closest
to the midline) near exhaustion. In the left hemisphere, consistent
inverse associations between affective responses and oxygenation
in all (dorsal and ventral) regions were observed at intensities
above the VT. In the right hemisphere, inverse associations
between affective responses and oxygenation in the lateral (dorsal
and ventral) and the midline (ventral) regions were observed at
intensities at the RCP and the end of exercise, respectively.

Author Contributions

Conceived and designed the experiments: GT RE GP. Performed the
experiments: GT. Analyzed the data: GT RE GP. Wrote the paper: GT
RE GP.

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