## WEB-ONLY DATA SUPPLEMENT

## Oxygen uptake kinetics during continuous exercise

In addition to the continuous exercise (CE) and intermittent exercise (IE) bouts, seven subjects performed 1 or 2 additional constant-load transitions to $70 \%$ of peak power. The protocol for these additional exercise bouts involved 3 min of unloaded cycling followed by an abrupt application of the predetermined workload for 7 min . The breath-by-breath oxygen uptake $\left(\dot{\mathrm{V}}_{\mathrm{O}_{2}}\right)$ data from each trial (including the first 7 min of the CE bout) were inspected for aberrant breaths. Values exceeding 3 standard deviations from the local mean were removed. The $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ data from the repeated trials were then interpolated to 1 second values, time aligned and averaged, effectively smoothing the data and enhancing the underlying kinetic response. A fourcompartment model with three exponential terms was used to describe the time course of the $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ response (Equation 1):
$\left.\dot{\mathrm{V}}_{\mathrm{O}_{2}}(\mathrm{t})=\mathrm{A}_{\mathrm{B}}+\mathrm{A}_{\mathrm{C}}\left(1-\mathrm{e}^{-\left(\mathrm{t}-\mathrm{TD} \mathrm{D}_{\mathrm{C}}\right) / \tau_{\mathrm{C}}}\right)+\mathrm{A}_{\mathrm{P}}\left(1-\mathrm{e}^{-(\mathrm{t}-\mathrm{TD} \mathrm{D}) / \tau_{\mathrm{P}}}\right)+\mathrm{A}_{\mathrm{S}}\left(1-\mathrm{e}^{-(\mathrm{t}-\mathrm{TD}} \mathrm{S}\right) / \tau_{\mathrm{S}}\right)$,
where $\dot{\mathrm{V}}_{\mathrm{O}_{2}}(\mathrm{t})$ is the $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ at time t ; $\mathrm{A}_{\mathrm{B}}$ is the baseline $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ during unloaded cycling, while $A_{C}, A_{p}$ and $A_{S}$ represent the cardio-dynamic (phase I), primary (phase II) and slow component amplitudes, respectively; $\mathrm{TD}_{\mathrm{C}}, \mathrm{TD}_{\mathrm{P}}$ and $\mathrm{TD}_{\mathrm{S}}$, and $\tau_{\mathrm{C}}, \tau_{\mathrm{P}}$ and $\tau_{\mathrm{S}}$ are the time delays and time constants of phase I and the primary and slow components, respectively.

If the amplitude of the slow component $\left(\mathrm{A}_{\mathrm{s}}\right)$ was not significantly different from 0 , the model was reduced to three compartments with exponential terms describing phase I and phase II (Equation 2):

$$
\dot{\mathrm{V}}_{\mathrm{O}_{2}}(\mathrm{t})=\mathrm{A}_{\mathrm{B}}+\mathrm{A}_{\mathrm{C}}\left(1-\mathrm{e}^{-\left(\mathrm{t}-\mathrm{TD}_{\mathrm{C}}\right) / \tau_{\mathrm{C}}}\right)+\mathrm{A}_{\mathrm{S}}\left(1-\mathrm{e}^{-\left(\mathrm{t}-\mathrm{TD}_{\mathrm{S}}\right) / \tau_{\mathrm{S}}}\right)
$$

Figure 1 shows an example of the $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ response and model fit in a representative subject with chronic obstructive pulmonary disease (COPD) and Table 1 provides the individual and group mean values for selected model parameters.


Figure 1. $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ response to constant-load exercise in a representative subject. Data points represent mean second-by-second values for 2 transitions to $70 \%$ of peak power. The solid line indicates the model fit, with the residuals shown at the bottom. The predetermined workload was applied at 60 s . The $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ values from 0 to 60 s are for unloaded cycling.

Table 1: Oxygen uptake kinetics in the transition to constant-load exercise performed at $70 \%$ of peak power in patients with moderate COPD.

| Subjects | $\mathrm{A}_{\mathrm{B}}$ ( $\mathrm{ml} / \mathrm{min}$ ) | $\mathrm{A}_{\mathrm{C}}$ ( $\mathrm{ml} / \mathrm{min}$ ) | $A_{P}$ ( $\mathrm{ml} / \mathrm{min}$ ) | $\mathrm{A}_{\mathrm{s}}$ (ml/min) | $\tau_{\mathrm{P}}$ <br> (s) | $\mathrm{TD}_{\mathrm{P}}$ <br> (s) | $\mathrm{TD}_{\mathrm{s}}$ <br> (s) |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 1 | 555 | 24 | 310 | 59 | 86 | 9 | 217 |
| 2 | 580 | 53 | 324 | 55 | 64 | 25 | 224 |
| 3 | 477 | 19 | 146 | -- | 97 | 32 | -- |
| 4 | 646 | 115 | 361 | 24 | 85 | 27 | 141 |
| 5 | 551 | 47 | 352 | 26 | 74 | 26 | 229 |
| 6 | 714 | 127 | 581 | -- | 115 | 22 | -- |
| 7 | 747 | 158 | 663 | 80 | 55 | 8 | 174 |
| Mean (SE) | 610 (37) | 78 (21) | 391 (66) | 49 (11) | 82 (8) | 21 (3) | 197 (17) |

$A_{B}$ : baseline oxygen uptake amplitude; $A_{C}$ : phase I amplitude; $A_{P}$ : phase II amplitude; $A_{s}$ : slow component amplitude; $\tau_{\mathrm{P}}$ : phase II time constant; $\mathrm{TD}_{\mathrm{p}}$ : phase II onset time; $\mathrm{TD}_{\mathrm{s}}$ : slow component onset time.

## The effect of $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ kinetics on the intermittent exercise response

To determine if the lower $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ response observed during IE compared to CE was primarily the result of the exponential shape of the on-kinetic response, we used a similar methodology to that recently described by Morris and colleagues. ${ }^{1}$ Briefly, we calculated the $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ amplitude at 60 s of CE and compared this value with the measured IE response (Table 2). The $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ amplitude for IE was determined as the average $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ measured during the final 10 s of each 60-s exercise interval over the duration of the entire IE bout. The predicted $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ value was calculated using a single exponential term, omitting the Phase I response (Equation 3):

$$
\left.\dot{\mathrm{V}}_{\mathrm{O}_{2}}(\mathrm{t})=\mathrm{A}_{\mathrm{B}}+\mathrm{A}_{\mathrm{P}}\left(1-\mathrm{e}^{-(\mathrm{t}-\mathrm{T}} \mathrm{P}\right) / \tau_{\mathrm{P}}\right)
$$

The $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ amplitude at 60 s was also re-calculated using a time constant ( $\tau_{\mathrm{P}}$ ) value of 42 s . This value is the mean phase II $\tau$ for healthy older individuals performing constant-load cycling at $\sim 70 \%$ of peak power. ${ }^{2}$

Table 2. Measured and predicted oxygen uptake values for intermittent exercise.

| Subjects | $\begin{aligned} & \text { Measured } \dot{\mathrm{V}}_{\mathrm{O}_{2}} \\ & (1 / \mathrm{min}) \end{aligned}$ | $\begin{aligned} & \text { Predicted } \dot{\mathrm{V}}_{\mathrm{O}_{2}} \\ & (1 / \mathrm{min}) \end{aligned}$ | $\begin{aligned} & \text { Predicted } \dot{\mathrm{V}}_{\mathrm{O}_{2}} \\ & (1 / \mathrm{min}) \end{aligned}$ |
| :---: | :---: | :---: | :---: |
|  |  | $\tau_{\mathrm{P}}$ from Table 1 | $\tau_{\mathrm{P}}=42 \mathrm{~s}$ |
| 1 | 0.76 | 0.72 | 0.81 |
| 2 | 0.78 | 0.79 | 0.87 |
| 3 | 0.88 | 0.89 | 1.01 |
| 4 | 0.77 | 0.75 | 0.85 |
| 5 | 1.05 | 1.29 | 1.37 |
| 6 | 0.97 | 1.00 | 1.25 |
| 7 | 0.50 | 0.54 | 0.60 |
| Mean (SE) | 0.82 (0.07) | 0.85 (0.09) | 0.97 (0.10)* |

$\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ : oxygen uptake; $\tau_{\mathrm{p}}$ : phase II time constant. The measured $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ values represent the mean end-exercise interval values obtained over the entire intermittent exercise test duration. The predicted $\dot{\mathrm{V}}_{\mathrm{O}_{2}}$ values were calculated using a single-term (Equation 3) exponential model (see text for details). * $\mathrm{P}<0.05$, significantly different from measured $\mathrm{V}_{\mathrm{O}_{2}}$ value (repeated measures ANOVA).

## REFERENCES

1 Morris N, Gass G, Thompson M, et al. Physiological responses to intermittent and continuous exercise at the same relative intensity in older men. Eur J Appl Physiol 2003;90:620-625.

2 Sabapathy S, Schneider DA, Comadira G, et al. Oxygen uptake kinetics during severe exercise: a comparison between young and older men. Respir Physiol Neurobiol 2004;139:203-213.

