Supplemental Online Content

Scandalis L, Kitzman DW, Nicklas BJ, et al. Skeletal muscle mitochondrial respiration and exercise intolerance in patients with heart failure with preserved ejection fraction. *JAMA Cardiol*. Published online May 10, 2023. doi:10.1001/jamacardio.2023.0957

This supplementary material has been provided by the authors to give readers additional information about their work.

eTable 1. Correlation Between Skeletal Muscle Respiration and Physical Function

Physical Function Measurements	CXI, N_P		CXI+CXII, NS_P		MAX ETS, NS_E	
	r	*p*-value	*r*	*p*-value	*r*	*p*-value
Peak VO_2, mL/kg/min	0.70	<.001	0.69	<.001	0.69	<.001
Peak VO_2, mL/min	0.43	<.001	0.47	<.001	0.41	<.001
6-minute walk distance	0.69	<.001	0.69	<.001	0.69	<.001
SPPB total	0.44	<.001	0.44	<.001	0.46	<.001
SPPB gait speed	-0.51	<.001	-0.52	<.001	-0.53	<.001
SPPB chair time	-0.53	<.001	-0.52	<.001	-0.55	<.001
Leg Strength – right	0.24	.07	0.26	.05	0.21	.11
Leg Strength - left	0.42	.002	0.44	<.001	0.41	.002

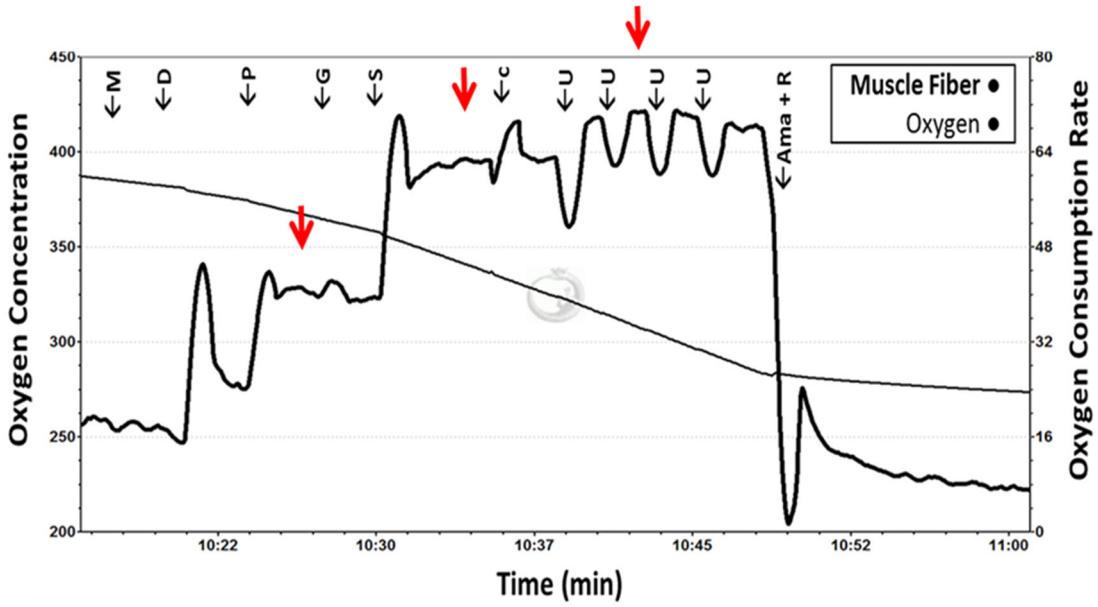
Pearson correlation coefficients and p-values for each association are shown.

CXI, Np, NADH-linked pathway through mitochondrial Complex I; CXI+CXII, NSp, NADH/Succinate combined pathway through Complexes I and II; MAX ETS, NS_E, maximum electron transport in the NADH/Succinate ET state; r, Pearson correlation coefficient; Peak VO_2, peak exercise oxygen consumption; SPPB, Short physical performance battery.

eTable 2. Correlation Between Skeletal Muscle Respiration and Peak VO_2 by HFpEF Status

	Peak VO_2, mL/kg/min			
	HFpEF (N=27)		**HC (N=45)**	
	R	p-value	r	p-value
CXI, N_P	0.45	.0194	0.26	.0787
CXI+CXII, NS_P	0.47	.0145	0.31	.0393
MAX ETS, NS_E	0.28	.1508	0.28	.0642

eFigure 1. Representative High-Resolution Respirometry Trace



Red arrows (from left to right) depict respiratory control states for OXPHOS capacity of the NADH pathway through Complex I (complex I, NP), the convergent NADH- and succinate pathway through Complex I and Complex II (complex I+complex II, NS_P), and the maximal electron transfer capacity of the systems-pathway (maximal capacity, NS_E).

eFigure 2. Association Between Skeletal Muscle Respiration and Peak VO_2 in HFpEF and Healthy Controls