



Ultimate team collaboration:  
Xtreme Everest 10 years on

**Kay Mitchell**

University Hospital Southampton



# Introduction

- Introduction to Xtreme Everest
- Some views on teamwork in science and research
- Caudwell Xtreme Everest 2007 – a case study of critical care research teamwork
- What can we learn from Xtreme Everest ten years on

# Introduction to Xtreme Everest

Xtreme Everest Oxygen Research Consortium – a collaboration between (founding partner) University College London, University of Southampton and Duke University, North Carolina, USA

# CASE Medicine

- UCL Centre for Altitude, Space and Extreme Environment Medicine
- Founded at the end of the nineties
- Clinicians and scientists with specialist interests and training in medicine and the physiology of extreme environments.
- Aimed to use the study of human systems exposed to these environments to increase our understanding of critically ill patients.





# One approach to critical care research

(Image shown with permission)





# Use models



Laboratory



Computer



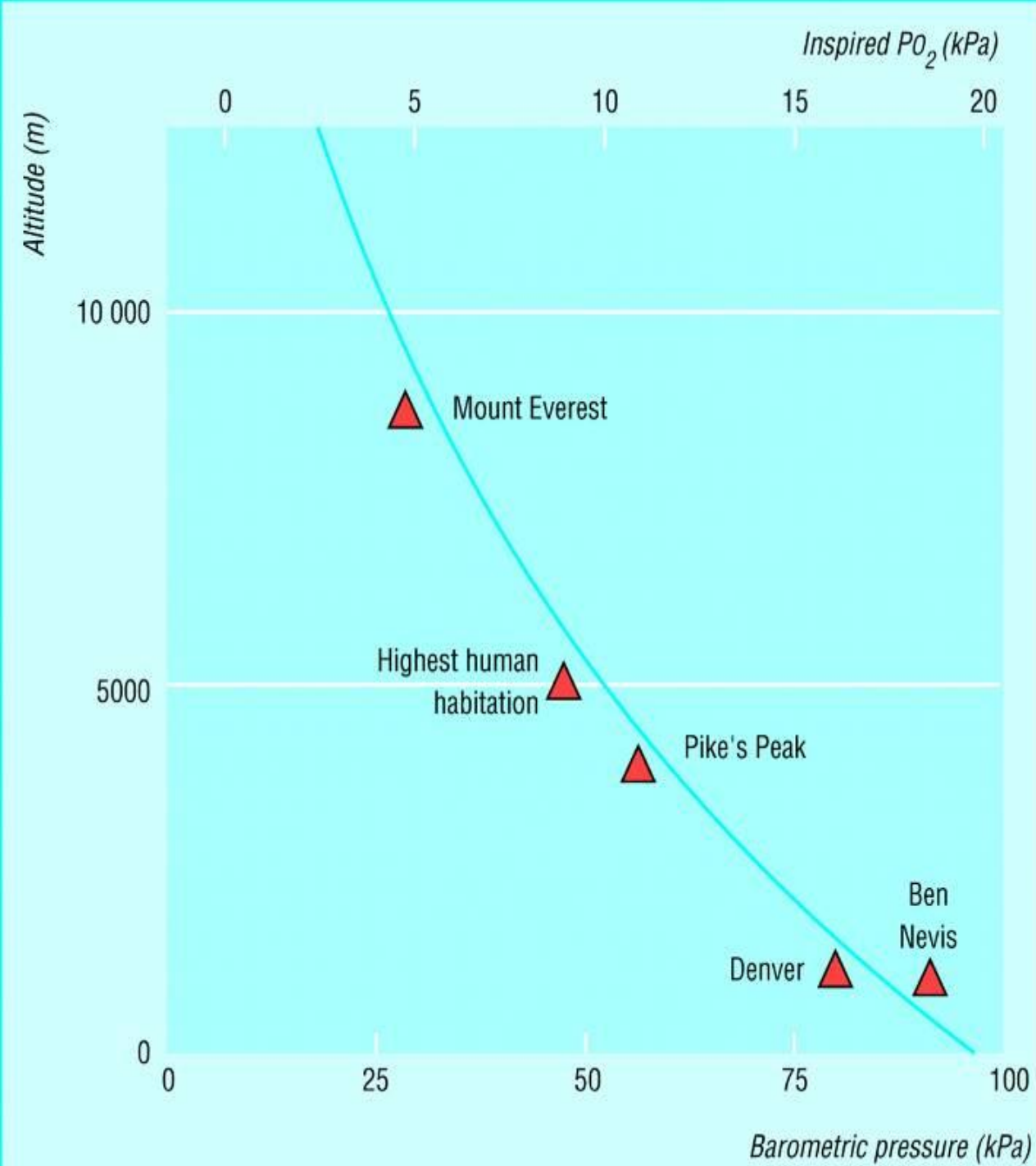
Healthy volunteers



# Why research at high altitude?

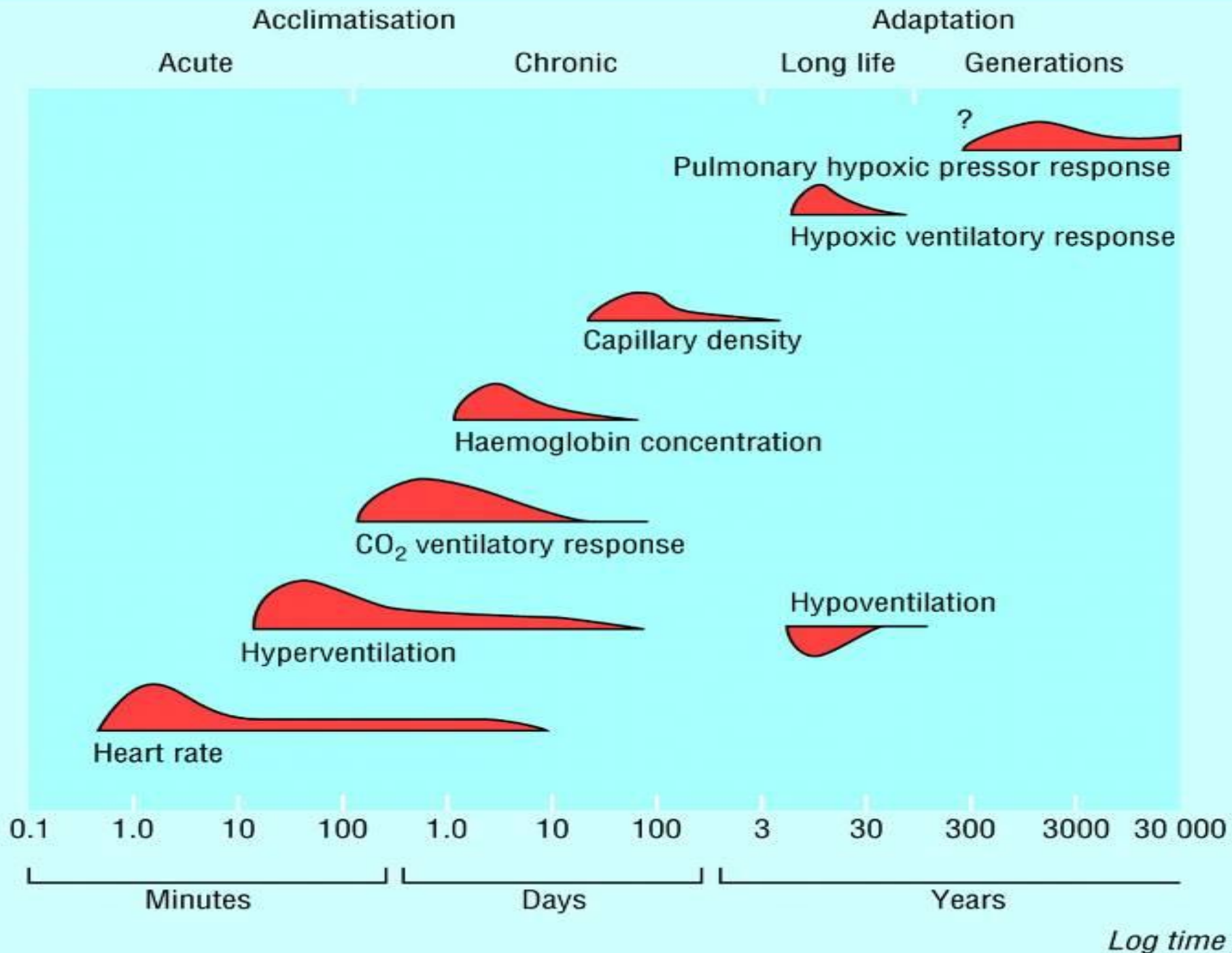
- Model for understanding critical illness
- Exercising volunteers at high altitude are experiencing similar physiological challenges to critically ill patients





# Acute hypoxia







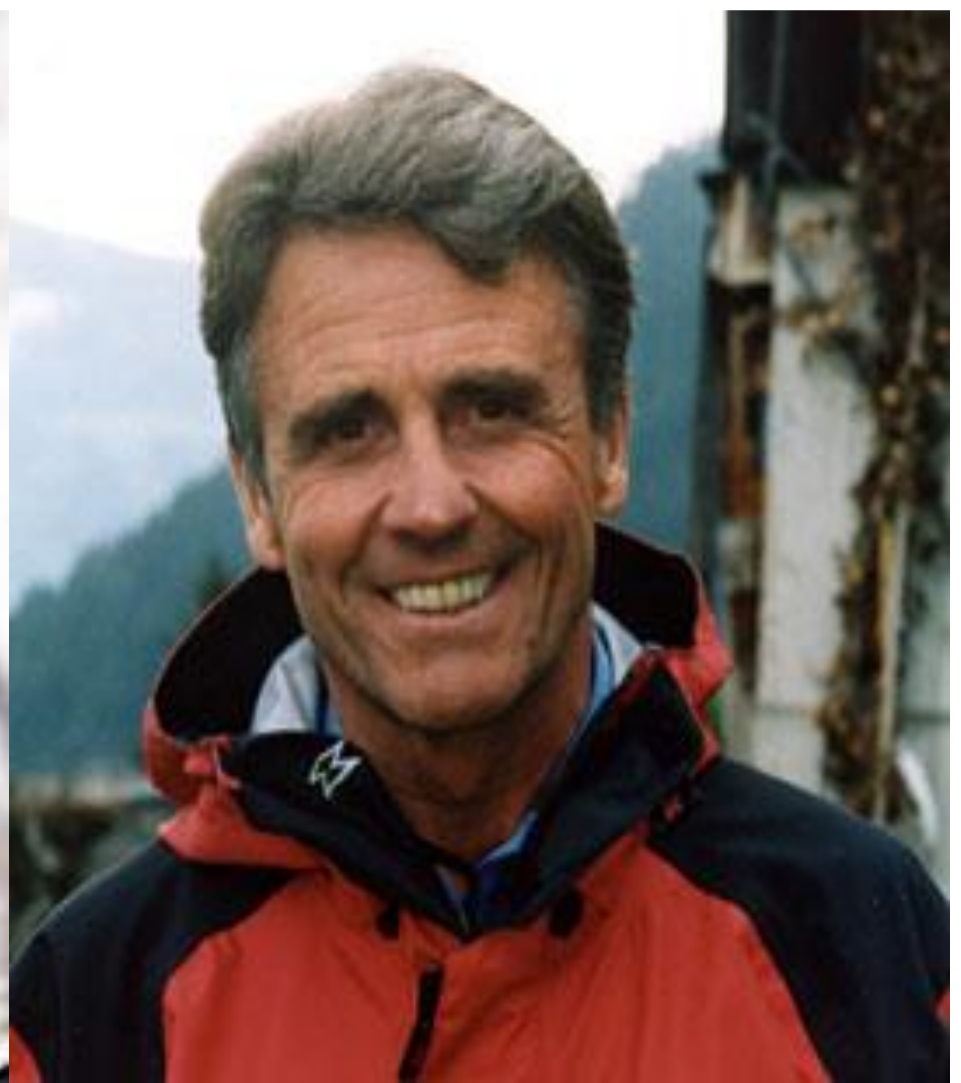
# Why Research on Everest?

- Traditional explanation of adaptation to altitude fails to explain differences in performance between individuals



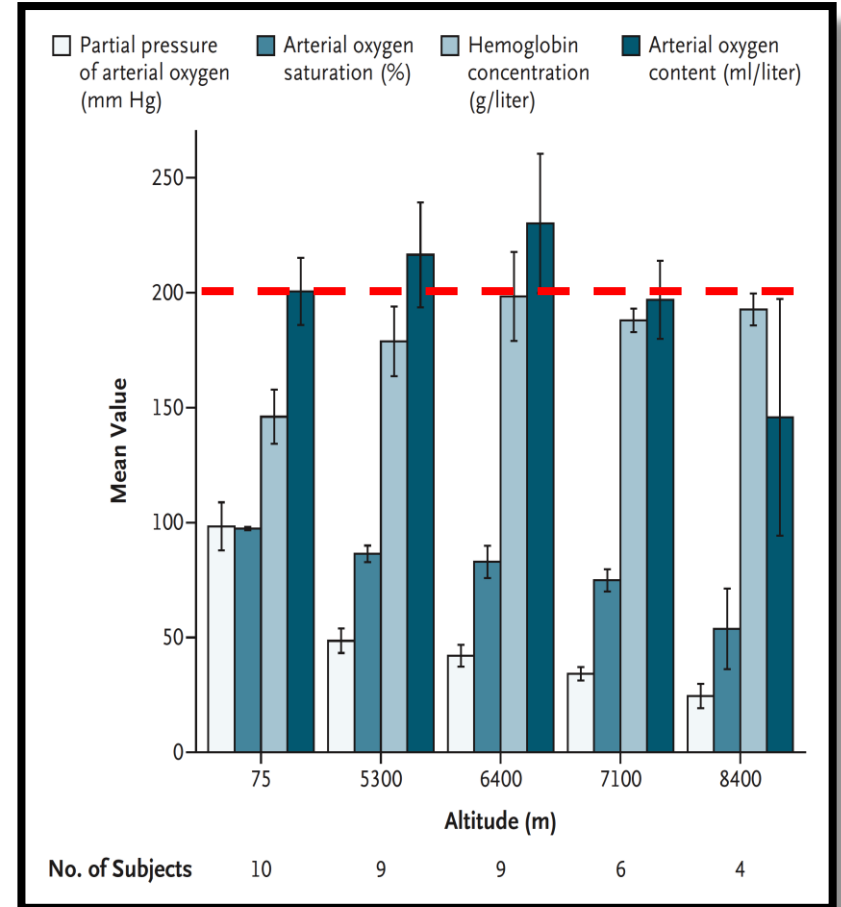
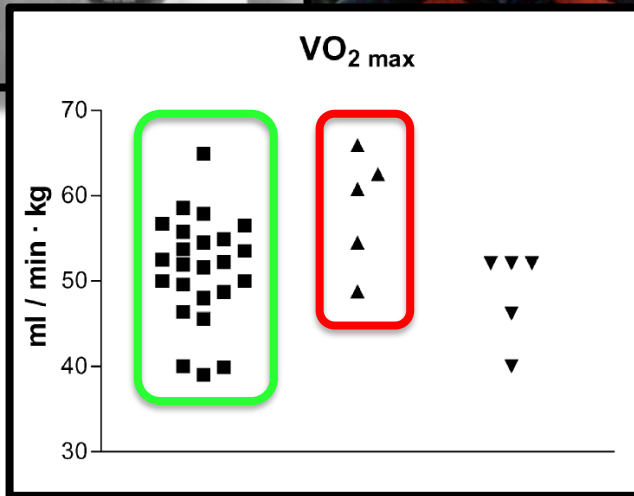


**Edmund Hillary and  
Tenzing Norgay**  
*29<sup>th</sup> May 1953*

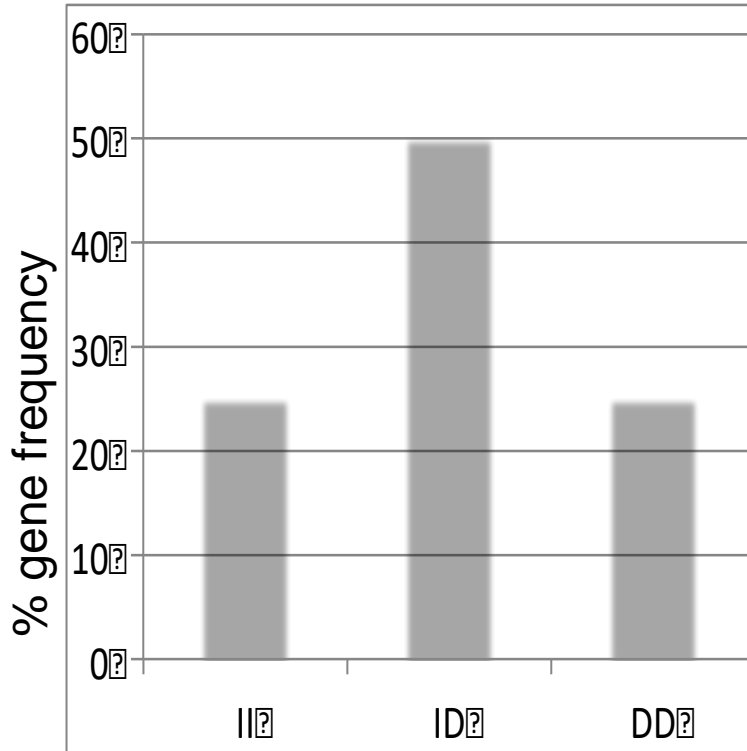


**Reinhold Messner and  
Peter Habeler**  
*8<sup>th</sup> May 1978*

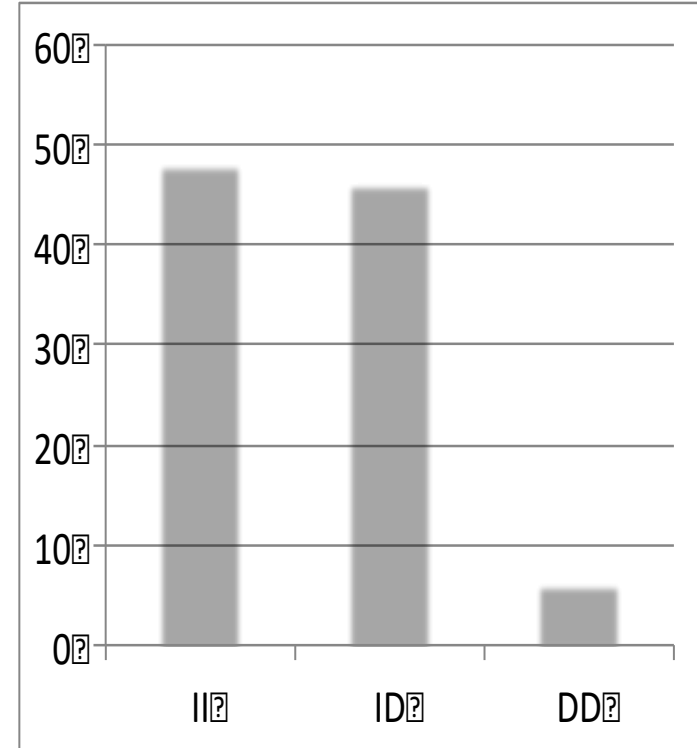
# Performance $\neq$ O<sub>2</sub> delivery



# ACE gene



Normal distribution

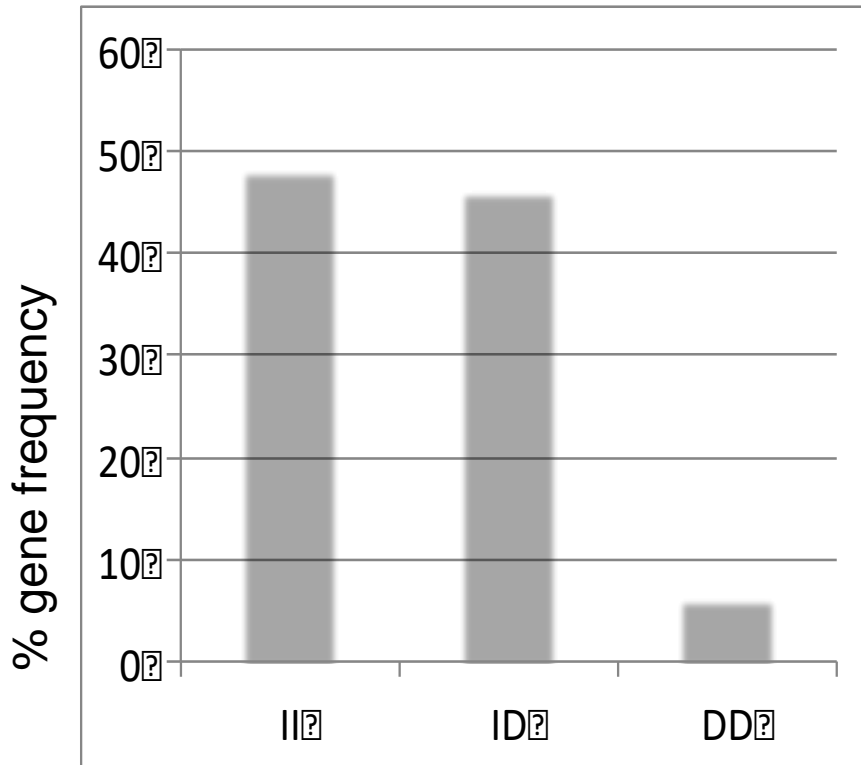


Elite High Altitude Climbers

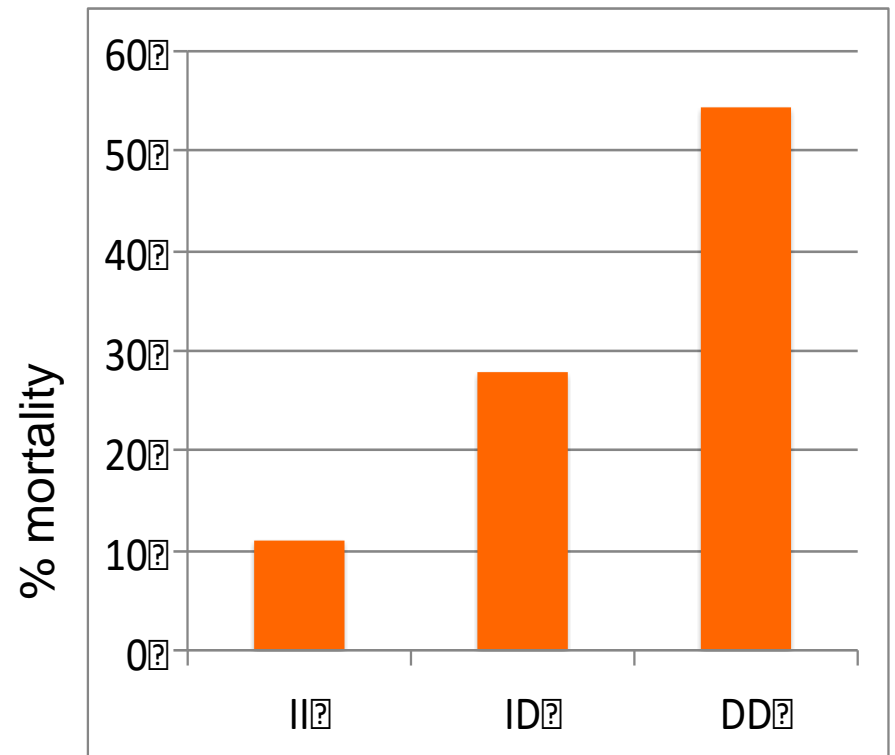
Montgomery *Nature* 1998

Marshall *AJRCCM* 2002

# ACE gene



Elite High Altitude Climbers



Critically Ill Patients (ARDS)

Montgomery *Nature* 1998  
Marshall *AJRCCM* 2002



# Areas for investigation

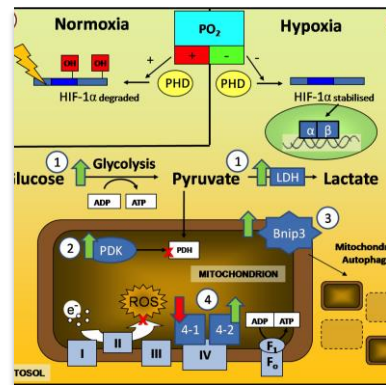
Metabolic function



Microcirculatory function



Mitochondrial function



Genetic differences





# Some views on teamwork in science and research

# Teamwork in healthcare

A systematic concept analysis in 2008 concluded teamwork to be "a dynamic process involving two or more healthcare professionals with complementary backgrounds and skills, sharing common health goals and exercising concerted physical and mental effort in assessing, planning, or evaluating patient care"

Xiryichis et al 2008

# “The Scientist”, Culotta 1993

- No single person is able to contribute all the necessary expertise to solve increasingly complex problems
- Research teams need a shared mission, a good organisational structure, and plenty of interpersonal interactions, says John Kutzbach
- Collaborations with an element of fun are most productive

# Science of team science (SciTS)

- “Mapping a research agenda for the science of team science”, Research Evaluation, 2011
- “An increase in cross-disciplinary, collaborative team science initiatives over the last few decades has spurred interest by multiple stakeholder groups in empirical research on scientific teams, giving rise to an emergent field referred to as the science of team science (SciTS).”
- NIHR and NIH supported

# Improving teamwork in anaesthesia and critical care - BJA, Brindley 2014

“To create a high-performing team we must understand the reality of the multidisciplinary team, how human factors impact on their performance, and the nature of how the different disciplines work, interact, and train.”

Five teamwork principles

- Leadership
- Coordination
- Mutual support
- Situation monitoring
- Communication

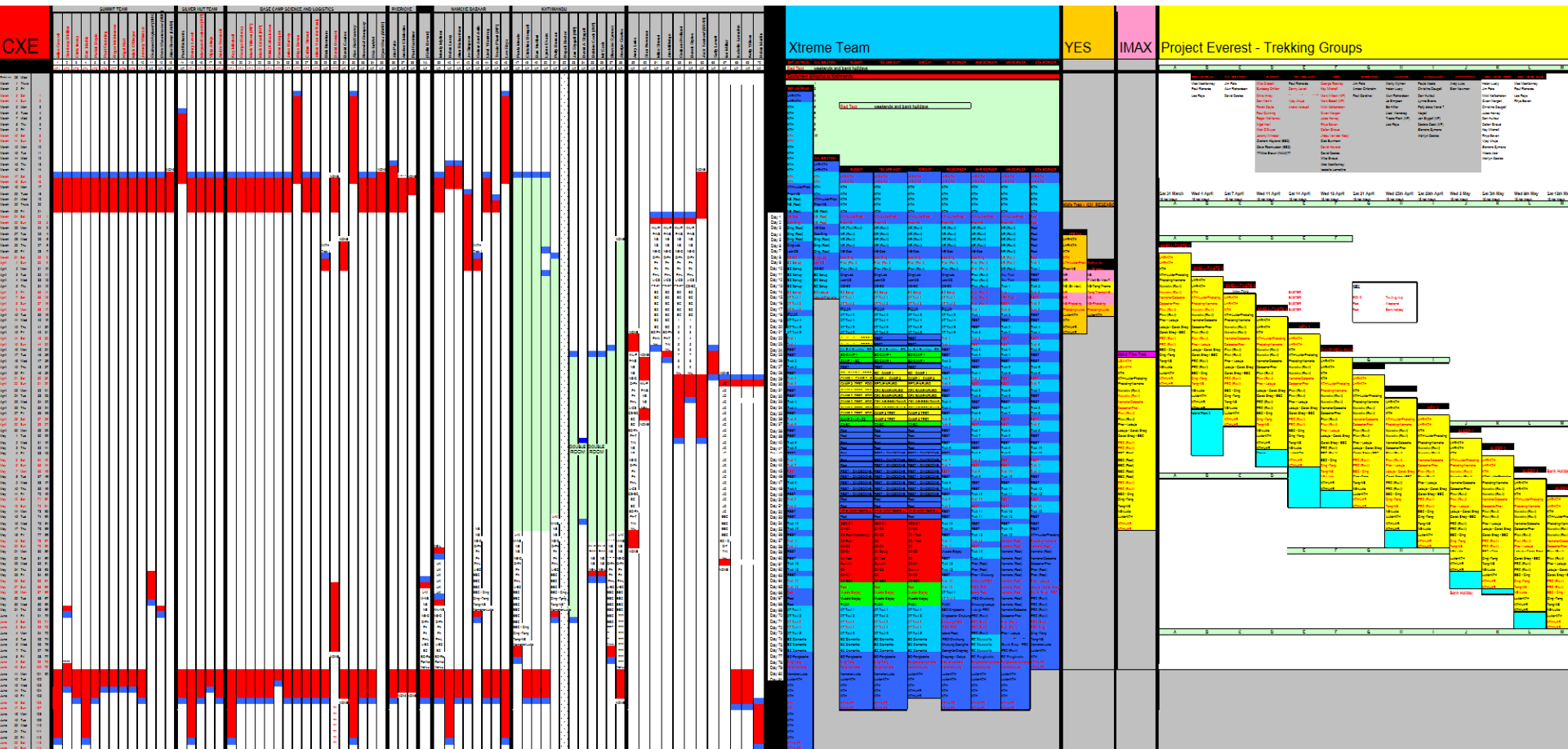
# Caudwell Xtreme Everest 2007 – a case study of critical care research teamwork

Using Brindley's five teamwork principles

# Leadership



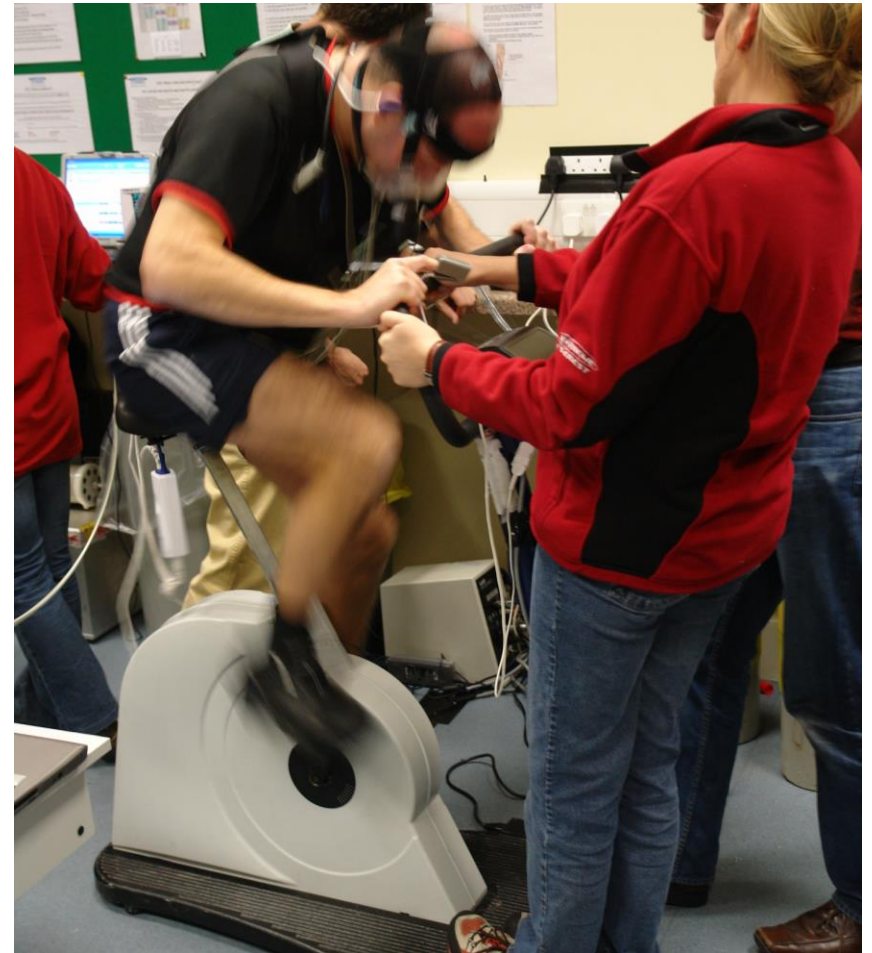
# Coordination (Project Management)





# Investigators – 45

- Medics, anaesthetists, surgeons and GPs
- Nurses, physiotherapists and dieticians
- Physiologists
- Research scientists
- Medical students
- Gap year students



# Subjects – 222

- 24 investigators
- 198 trekkers included:
  - ✓ Record producer
  - ✓ Dry stone waller
  - ✓ Retired fighter pilot



# Sherpa colleagues

**Climbing**



**Laboratory and trekking**





# Logistics team



# Equipment



- >26 metric tonnes
- >1000 containers
- >1m items

Including

Devices

Disposables

Medical kit

Computers

Generators and inverters

Food treats



















# Power!









# Mutual support



# Situation Monitoring

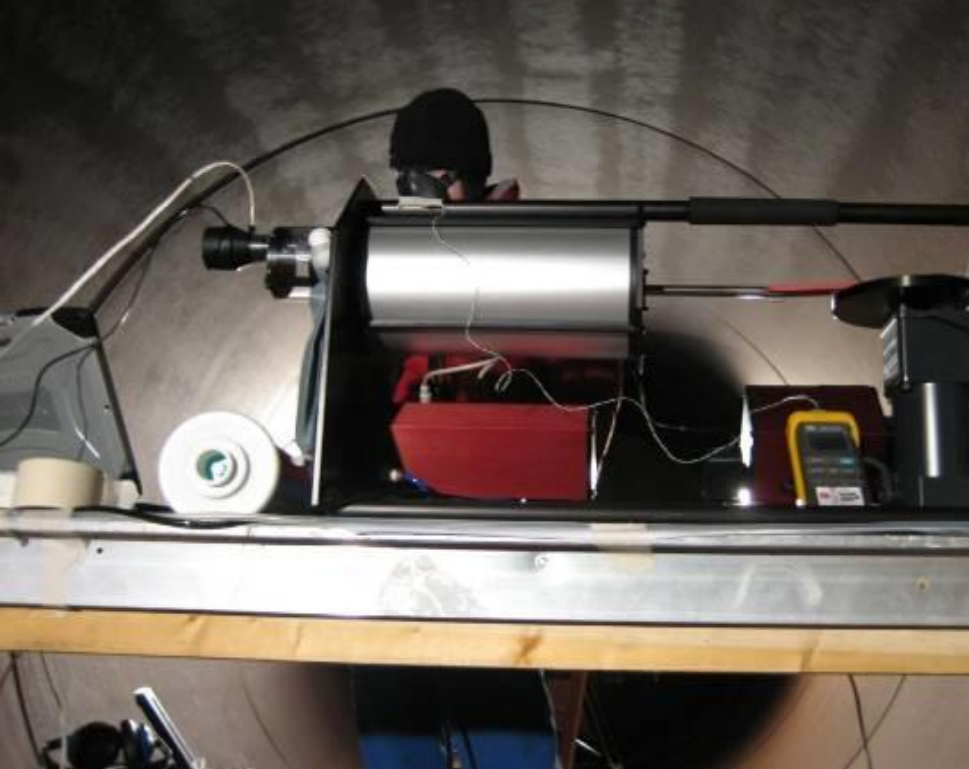
- Safety
- Science
- (Summit)
- Expedition Management Group
- +/- Climbing Management Group
- Laboratory Management
- Medical management
- Publicity
- Use of photographs
- Scientific publications

# Delivering robust data



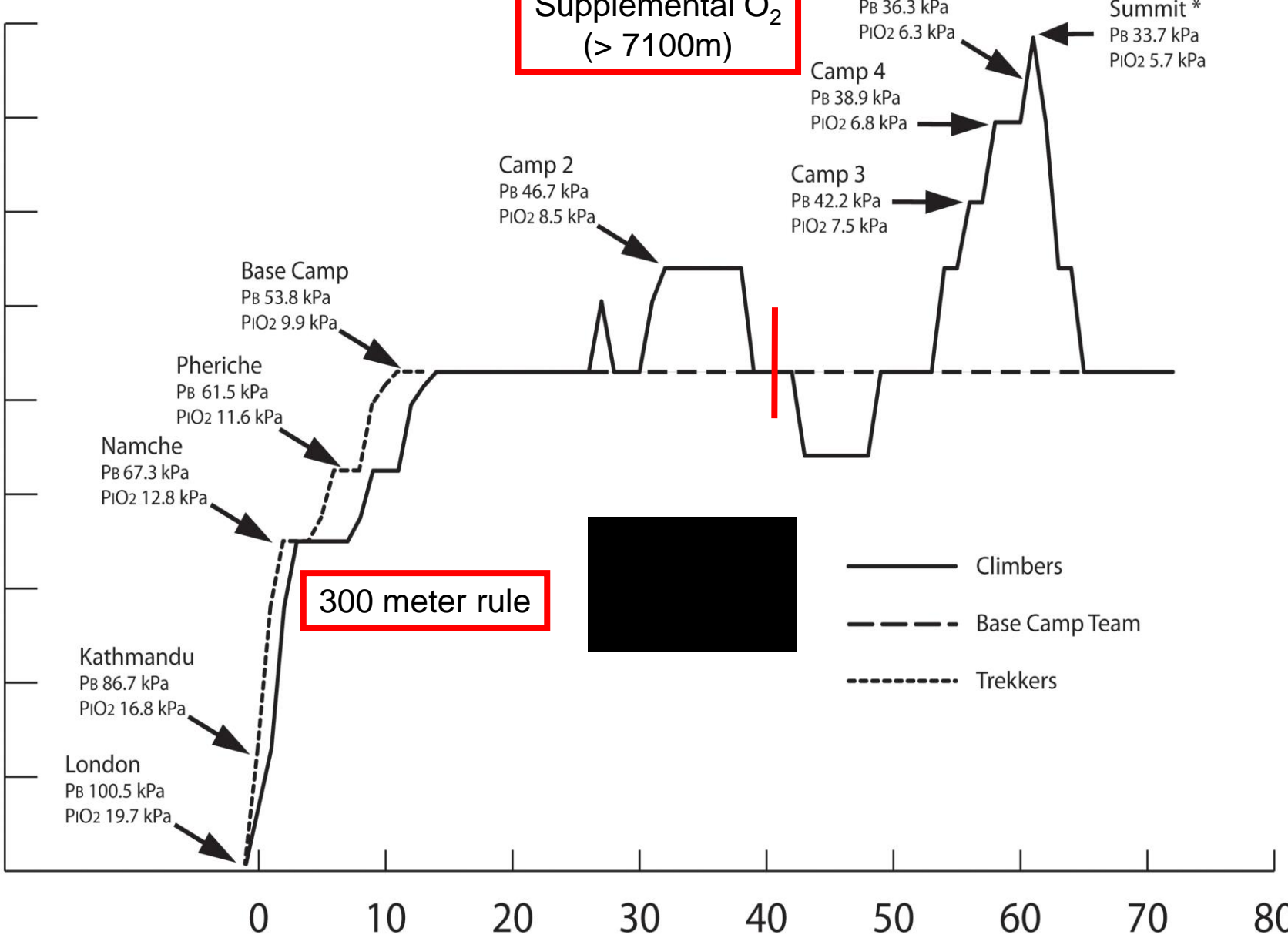
- Feasibility
- Reliability
- Validation
  - Pilot data
- Sea-level studies
  - Physiology laboratory
  - Environmental chamber
  - Hypobaric Chamber
- Field studies
  - Alps 2005 & 2006
  - Cho Oyu 2005
  - Cho Oyu 2006





Altitude (metres)

9000  
8000  
7000  
6000  
5000  
4000  
3000  
2000  
1000



Intermittent Supplemental O<sub>2</sub> (> 7100m)

300 meter rule

- Climbers
- - - Base Camp Team
- · · Trekkers

Expedition Day

0 10 20 30 40 50 60 70 80



# Communication

## Internal

- Set up and project development
- Inter-laboratory communications –using satellite phones and two way radios
- Overcoming isolation from family and friends

## External

- Website (using BGAN)
- Press and public relations
- Journal articles

# Study groups

**CXE**

## 'Trekking'

n = 198

Healthy volunteers  
13 treks of 16 subjects  
Study length 14 days

London 50m - EBC 5300m

## 'Investigators'

n = 24

Volunteer Investigators  
Study length 75 days

## EBC Staff

n = 10

London - EBC  
+ EBC (week 9)

## Climbers

n = 14

London - EBC  
+ 5300-8400m  
+ EBC (week 9)

# Phenotypic Variables

**CXE**

## *Trekkers (n = 198)*

VO<sub>2</sub> max and AT  
Oxygen efficiency  
Hb and Hct  
Inflammatory markers  
Muscle NIRS  
Brain NIRS  
Spirometry  
Neurocognitive testing  
Weight change  
Sleep studies  
Smell and taste  
Retinal Photography

[Pulmonary vascular response]

## *Investigators (n =24)*

*In addition...*

Arterial blood gases  
Skeletal muscle biopsies  
Sublingual microcirculation  
Gastric tonometry / DO<sub>2</sub>  
Cerebral Doppler  
Body composition change  
Resting metabolic rate  
Nutritional studies  
Structural brain MRI  
MR spectroscopy  
Eye saccades  
Laryngoscopy  
Thromboelastography

# Research expedition model

- Repeated measures of a core data set
- as environmental oxygen decreased
- on ascent to Everest Base Camp,
- and in sub group, to Summit



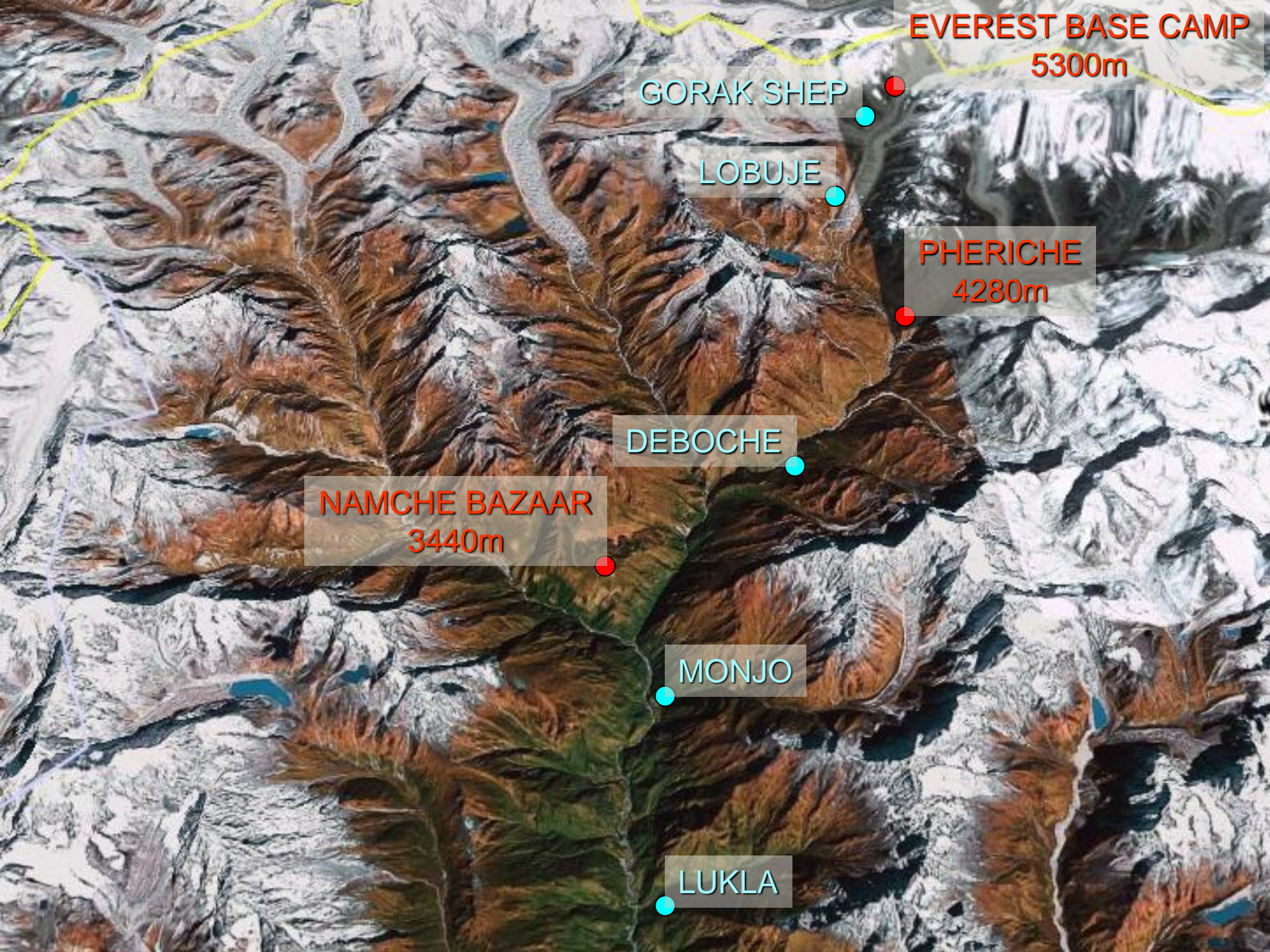












EVEREST BASE CAMP  
5300m

GORAK SHEP

LOBUJE

PHERICHE  
4280m

DEBOCHE

NAMCHE BAZAAR  
3440m

MONJO

LUKLA

















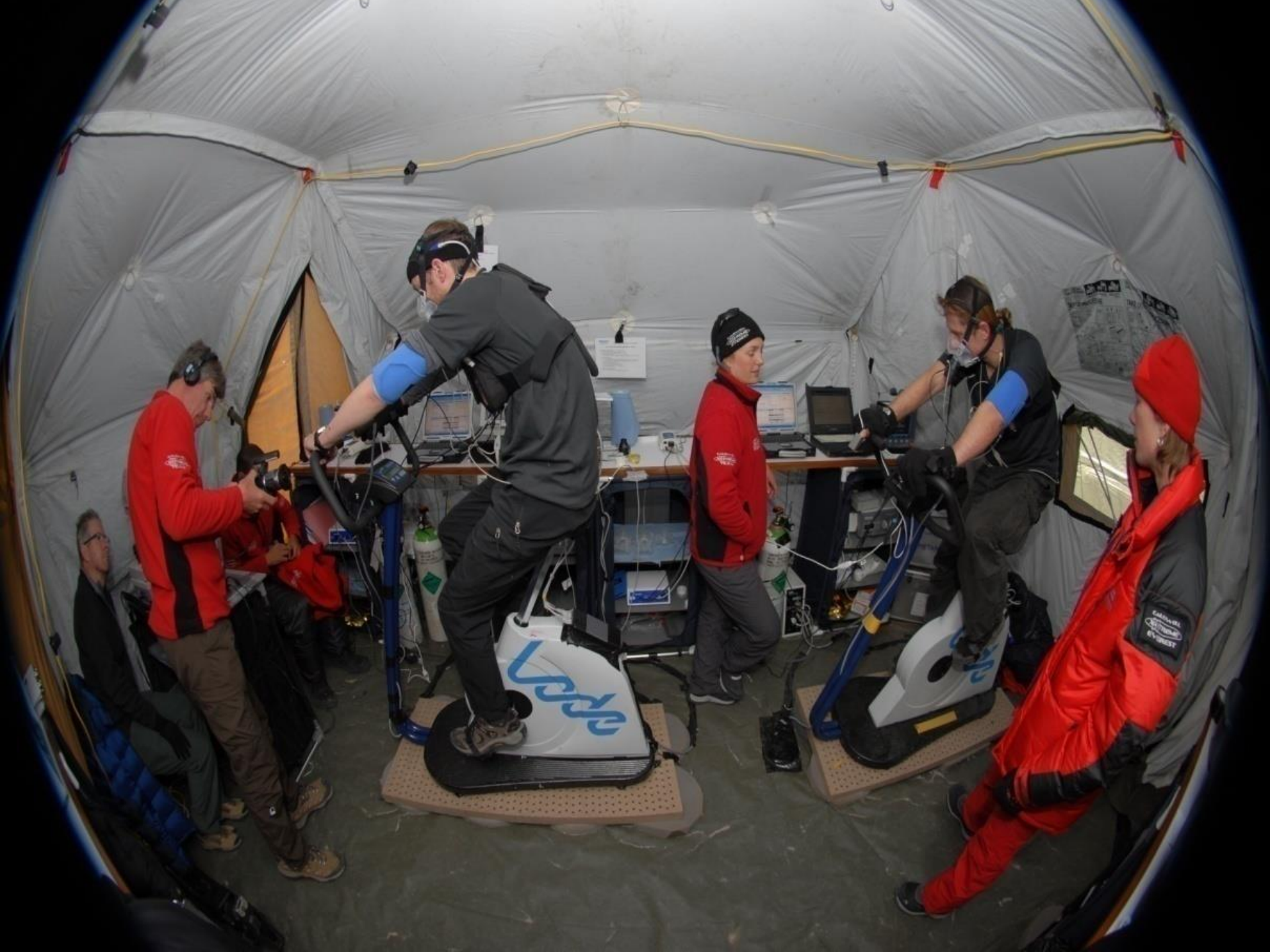




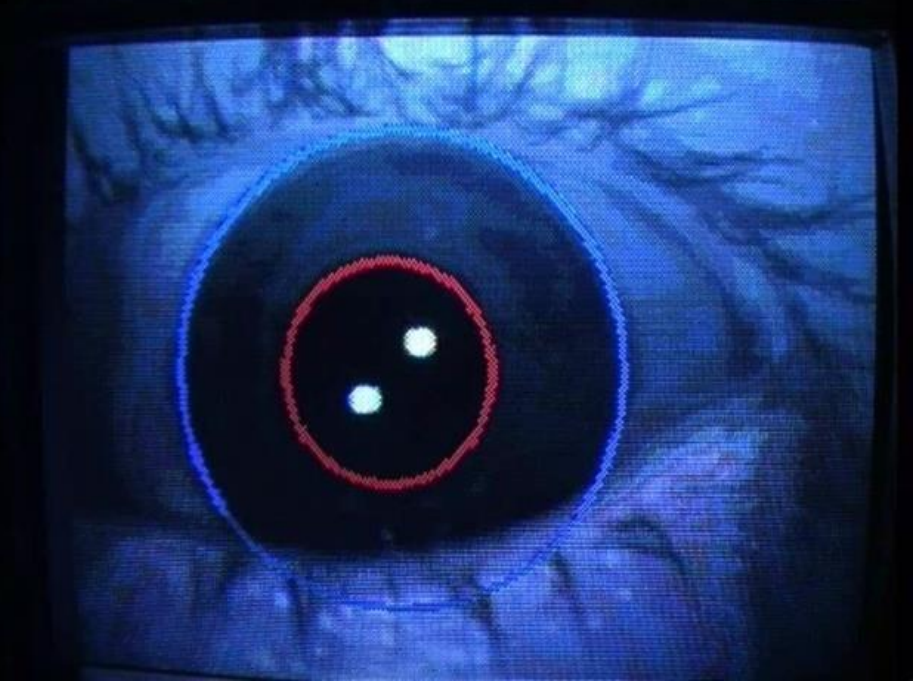
























SUMMIT  
8848m

Balcony (8400m)

South Col (7950m)

CAMP 4

CAMP 3

Western Cwm (6400m)

CAMP 2

CAMP 1

EBC

Image © 2007 DigitalGlobe  
Image © 2007 TerraMetrics  
© 2007 Europa Technologies

© 2007

Go















































CAUDWELL  
XTREME  
EVEREST



# Success?

- **Safety**
  - All significant illness evacuated safely with good outcome
  - 8 turned back, 4 evacuations
- **Science**
  - >93% of planned testing completed
  - Reliable and valid measurements
  - Largest controlled human physiology experiment at altitude
  - Highest measurements of several variables
- **Climbing success**
  - 190/198 trekkers to basecamp
  - 25 climbers to summit (8 doctors, 2 cameramen, 15 sherpas)

What can we learn from Xtreme Everest ten years on?



# Xtreme Everest Oxygen Research Consortium

## 12 research studies

- Cho Oyu 2005
- Cho Oyu 2006
- **Caudwell Xtreme Everest 2007**
- Young Everest Study 2007
- **CXE 2009**
- Xtreme Alps 2010
- **Bedford School 2012**
- **Simulated Altitude Study 2012**
- **EMC2 2012/3**
- **XE2 2013**
- YES 2 2013
- XEX 2017



**High-altitude physiology and pathophysiology: implications and relevance for intensive care medicine**

Michael Grocott, Hugh Montgomery and Andre Vercueil

Centre for Altitude, Space and Extreme Environment Medicine (CASE Medicine), UCL Institute of Human Health and Performance, Ground Floor, Charterhouse Building, UCL Archway Campus, Highgate Hill, London, N19 5LW, UK  
Exp Physiol 95.3 pp 463–470

Corresponding author: Michael Grocott, mike.grocott@ucl.ac.uk

Published: 1 February 2007  
This article is online at <http://ccforum.com/content/11/1/203>  
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Critical Care 2007, 11:203 (doi:10.1186/cc5142)

THE NEW ENGLAND JOURNAL OF MEDICINE

ORIGINAL ARTICLE

**Arterial Blood Gases and Oxygen Content in Climbers on Mount Everest**

Michael P.W. Grocott, M.B., B.S., Daniel S. Martin, M.B., Ch.B., Denny Z.H. Levett, B.M., B.Ch., Roger McMorrow, M.B., B.Ch., Jeremy Windsor, M.B., Ch.B., and Hugh E. Montgomery, M.B., B.S., M.D., for the Caudwell Xtreme Everest Research Group<sup>§</sup>

**Caudwell Xtreme Everest: a field study of human adaptation to hypoxia**

Daniel S. Martin<sup>1</sup>, Denny Z. H. Levett<sup>1</sup>, Mike P. W. Grocott<sup>1</sup> and Hugh E. Montgomery<sup>1</sup>  
Mike Grocott<sup>1</sup>, Alan Richardson<sup>2</sup>, Hugh Montgomery<sup>1</sup> and Monty Mythen<sup>1</sup>

<sup>1</sup>Centre for Altitude, Space and Extreme Environment Medicine (CASE Medicine), UCL Institute of Human Health and Performance, Ground Floor, Charterhouse Building, UCL Archway Campus, Highgate Hill, London, N19 5LW, UK  
<sup>2</sup>Chelsea School, University of Brighton, Hillbrow, Denton Road, Eastbourne BN20 7SR, UK

Corresponding author: Monty Mythen, montymythen@btinternet.com

Published: 1 August 2007  
This article is online at <http://ccforum.com/content/11/4/151>  
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Critical Care 2007, 11:151 (doi:10.1186/cc5921)

Original article

**The Young Everest Study: effects of hypoxia at high altitude on cardiorespiratory function and general well-being in healthy children**

E Scrase,<sup>1</sup> A Lavery,<sup>1</sup> J C D Gavlak,<sup>1</sup> S Sonnappa,<sup>2</sup> D Z H Levett,<sup>3</sup> D Martin,<sup>3</sup> M P W Grocott,<sup>3</sup> J Stocks<sup>2</sup>

The FASEB Journal • Research Communication

**Cardiac response to hypobaric hypoxia: persistent changes in cardiac mass, function, and energy metabolism after a trek to Mt. Everest Base Camp**

Oliver J. Rider,<sup>1</sup> Denny Z. H. Levett,<sup>2</sup> Damian J. Tyler,<sup>3\*</sup> Jane M. Francis,<sup>1</sup> Stefan Neubauer,<sup>1</sup> Michael P. W. Grocott,<sup>2</sup> and Kieran Clarke,<sup>3\*</sup> for the Caudwell Xtreme Everest Research Group

<sup>1</sup>Department of Physiology, Anatomy, and Genetics and <sup>2</sup>The University of Oxford Centre for Clinical Magnetic Resonance Research, University of Oxford, Oxford, UK; and <sup>3</sup>University College

Research

**Changes in skeletal muscle oxygenation during exercise measured by near-infrared spectroscopy on ascent to altitude**

Daniel S Martin<sup>1</sup>, Denny ZH Levett<sup>1</sup>, Michael Mythen<sup>1,2</sup> and Mike PW Grocott<sup>1</sup>, for the Caudwell Xtreme Everest Research Group

**Experimental Physiology – Review Article**

**Variation in human performance in the hypoxic mountain environment**



**The role of nitrogen oxides in human adaptation to hypoxia**

SUBJECT AREAS:  
CHEMICAL BIOLOGY  
MEDICAL RESEARCH  
PHYSIOLOGY

Denny Z. Levett<sup>1\*</sup>, Bernadette O. Fernandez<sup>2\*</sup>, Heather L. Riley<sup>3</sup>, Daniel S. Martin<sup>1</sup>, Kay Mitchell<sup>4</sup>, Carl A. Leckstrom<sup>5</sup>, Can Ince<sup>6</sup>, Brian J. Whipp<sup>6</sup>, Monty G. Mythen<sup>1</sup>, Hugh E. Montgomery<sup>1</sup>, Mike P. Grocott<sup>1\*</sup> & Martin Feilisch<sup>7\*</sup>, for the Caudwell Xtreme Everest Research Group



**Anaesthesia**  
Journal of the Association of Anaesthetists of Great Britain and Ireland

Anaesthesia, 2011, 66, pages 348–353

doi:10.1111/j.1365-2044.2011.

ORIGINAL ARTICLE

**A novel ambulatory closed circuit breathing system for during exercise**

R. C. N. McMorrow,<sup>1,2</sup> J. S. Windsor,<sup>3,4</sup> M. G. Mythen,<sup>5,6</sup> M. P. W. Grocott<sup>7,8</sup> and the Caudwell Xtreme Everest Research Group\*

**Changes in sublingual microcirculatory flow index and vessel density on ascent to altitude**

Daniel S. Martin<sup>1</sup>, Peter Goedhart<sup>2</sup>, Andre Vercueil<sup>1,3</sup>, Can Ince<sup>2</sup>, Denny Z. H. Levett<sup>1</sup> and Mike P. W. Grocott<sup>1</sup> for the Caudwell Xtreme Everest Research Group

<sup>1</sup>UCL Centre for Altitude, Space and Extreme Environment Medicine (CASE Medicine), Portex Unit, Institute of Child Health, 30 Gullford Street, London WC1N 1EH, UK

<sup>2</sup>Department of Translational Physiology, Academic Medical Center, University of Amsterdam, Meibergdreef 9, 1105 AZ Amsterdam, The Netherlands

<sup>3</sup>King's College Hospital, Denmark Hill, London SE5 8RX, UK  
[OPEN ACCESS Freely available online](https://doi.org/10.1186/1745-2975-13-151)



**The Effect of High-Altitude on Human Skeletal Muscle Energetics: <sup>31</sup>P-MRS Results from the Caudwell Xtreme Everest Expedition**

Available online <http://ccforum.com/content/13/S5/S7>

Lindsay M. Edwards<sup>1,2,\*</sup>, Andrew J. Murray<sup>1,2</sup>, Damian J. Tyler<sup>1,2</sup>, Graham J. Kemp<sup>3</sup>, Cameron J. Holloway<sup>2</sup>, Peter A. Robbins<sup>1</sup>, Stefan Neubauer<sup>2</sup>, Denny Levett<sup>4</sup>, Hugh E. Montgomery<sup>4,5</sup>, Mike P. Grocott<sup>4</sup>, Kieran Clarke<sup>1</sup>, Caudwell Xtreme Everest Research Group

<sup>1</sup>Department of Physiology, Anatomy, and Genetics, University of Oxford, Oxford, UK; <sup>2</sup>The Oxford Centre for Clinical Magnetic Resonance Research, University of Oxford, Oxford, UK; <sup>3</sup>School of Clinical Sciences, University of Liverpool, Liverpool, Merseyside, United Kingdom; <sup>4</sup>UCL Centre for Altitude, Space and Extreme Environment Medicine (CASE Medicine), UCL Institute of Human Health and Performance, Charterhouse Building, UCL Archway Campus, Highgate Hill, London, N19 5LW, UK; and <sup>5</sup>Institute for Human Health and Performance, UCL Archway Campus, Highgate Hill, London, N19 5LW, UK



# Pubmed search – 22<sup>nd</sup> May 2017

- 34 listed articles
  - Cho Oyu 2006 - 1
  - CXE - 26
  - CXE 2009 – 1
  - Xtreme Alps – 2
  - XE 2 – 3
  - Miscellaneous – 1



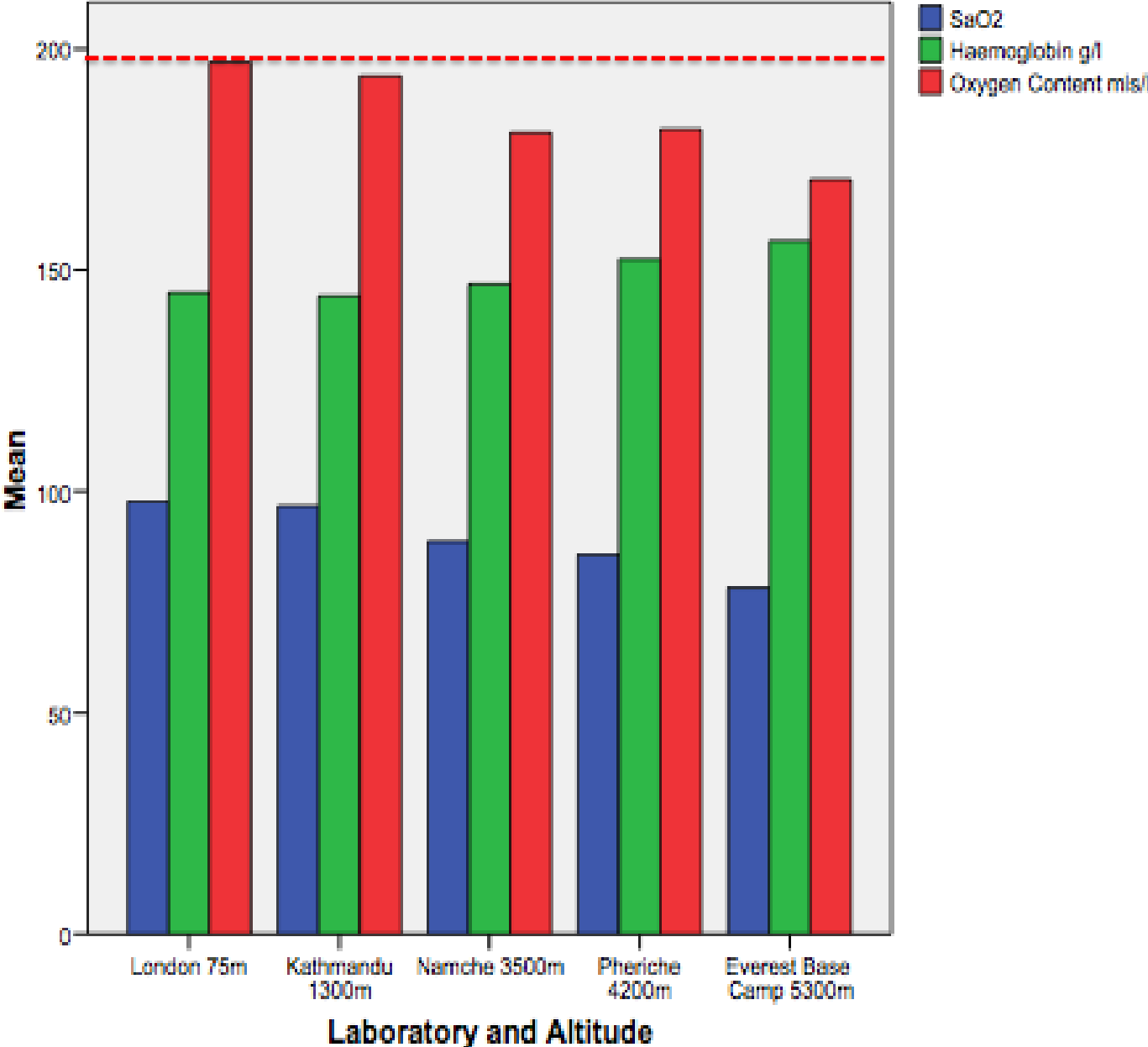
**HUMAN PHYSIOLOGY AT EXTREME ALTITUDE**

Does not include  
numerous invited reviews

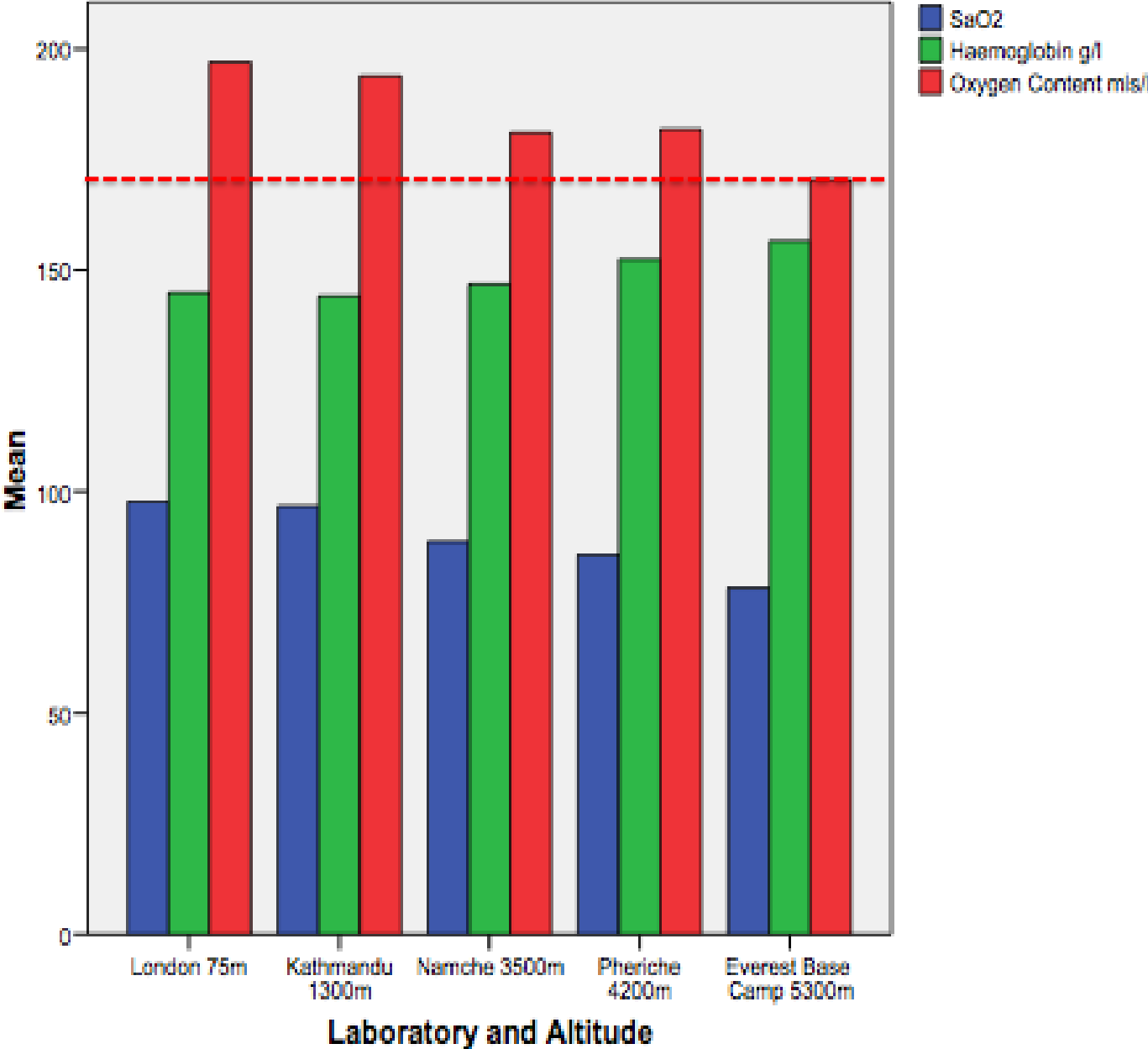




# Oxygen Saturations, Haemoglobin and Oxygen Content (Trekking)

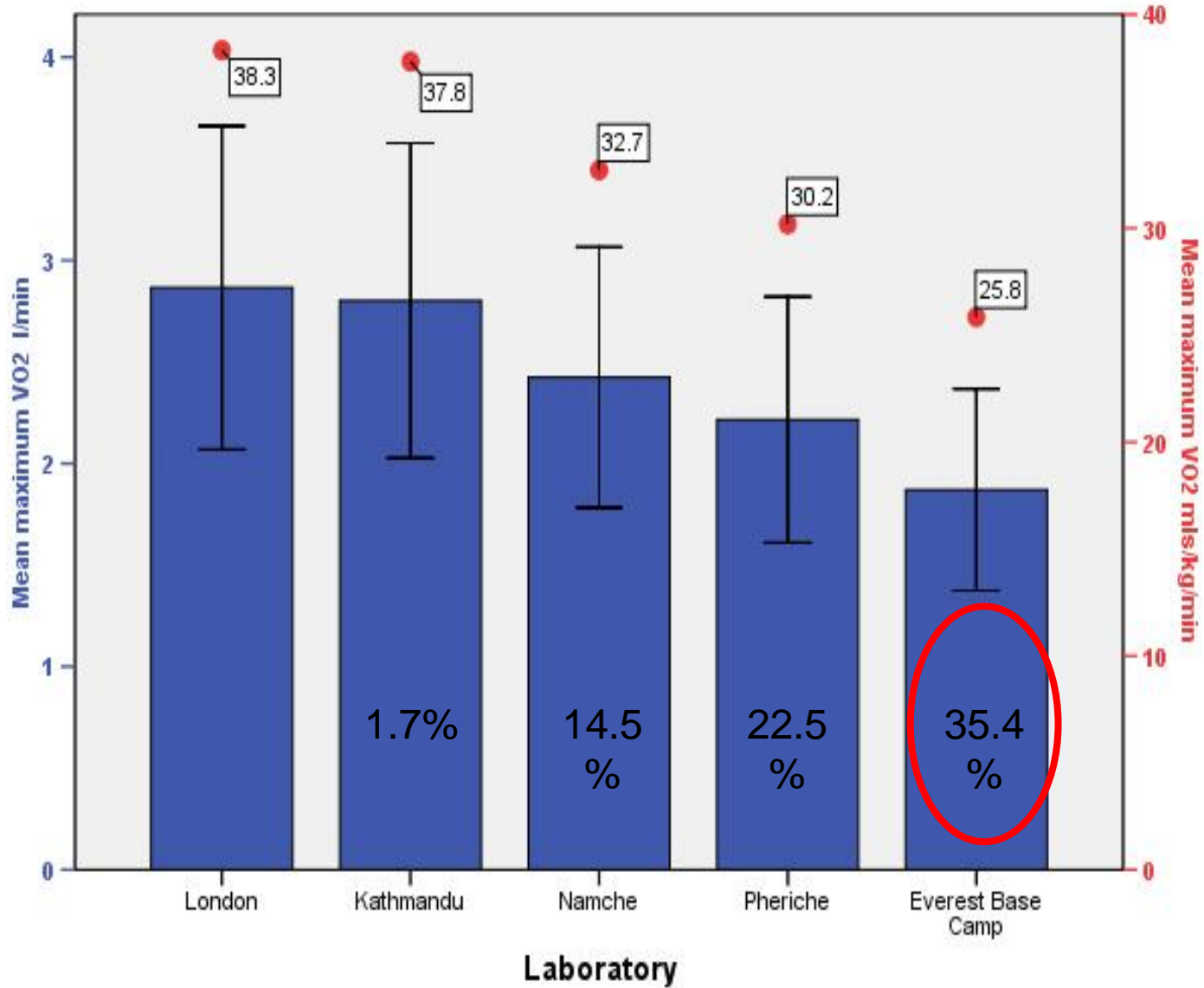


# Oxygen Saturations, Haemoglobin and Oxygen Content (Trekking)





# Maximum Oxygen Consumption at Altitude, Trekkers n = 190



Error Bars: +/- 1 SD

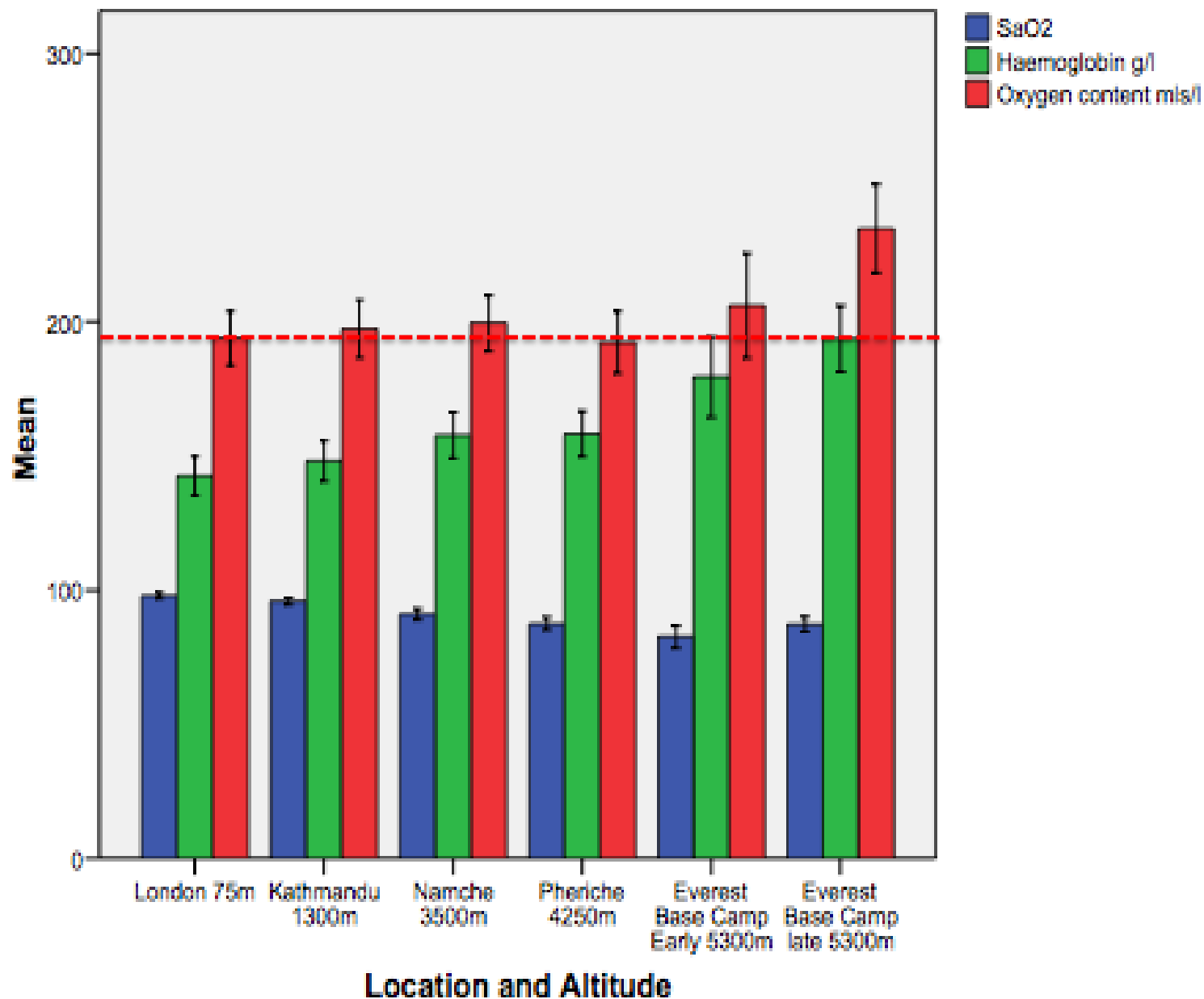




# VO<sub>2</sub> max at 7950m

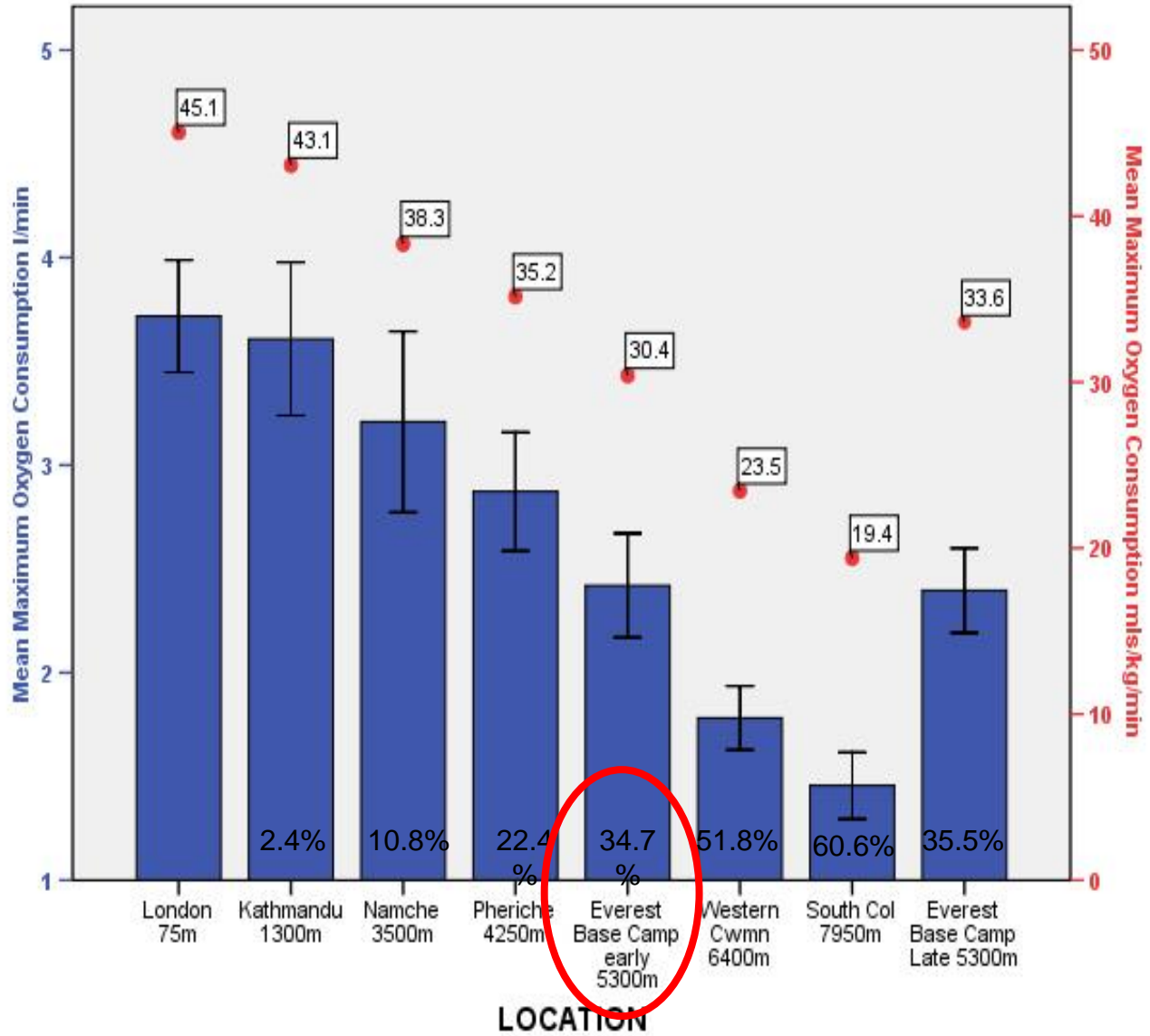


# Haemoglobin, Arterial Oxygen Saturations and Oxygen Content (Climbers n=14)



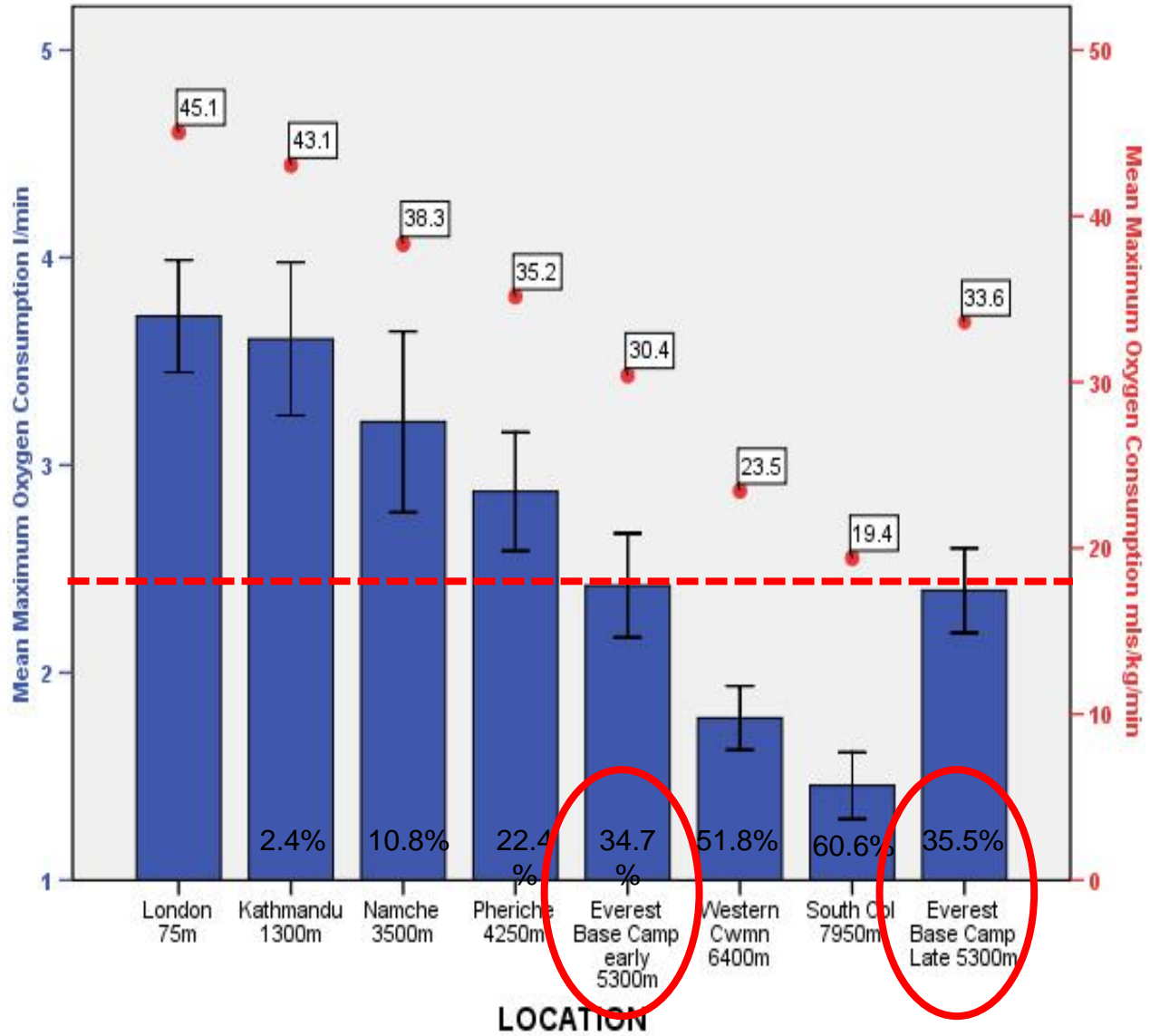


# Maximum Oxygen Consumption at Altitude, South Col Team, [REDACTED]



Error Bars: +/- 1 SD

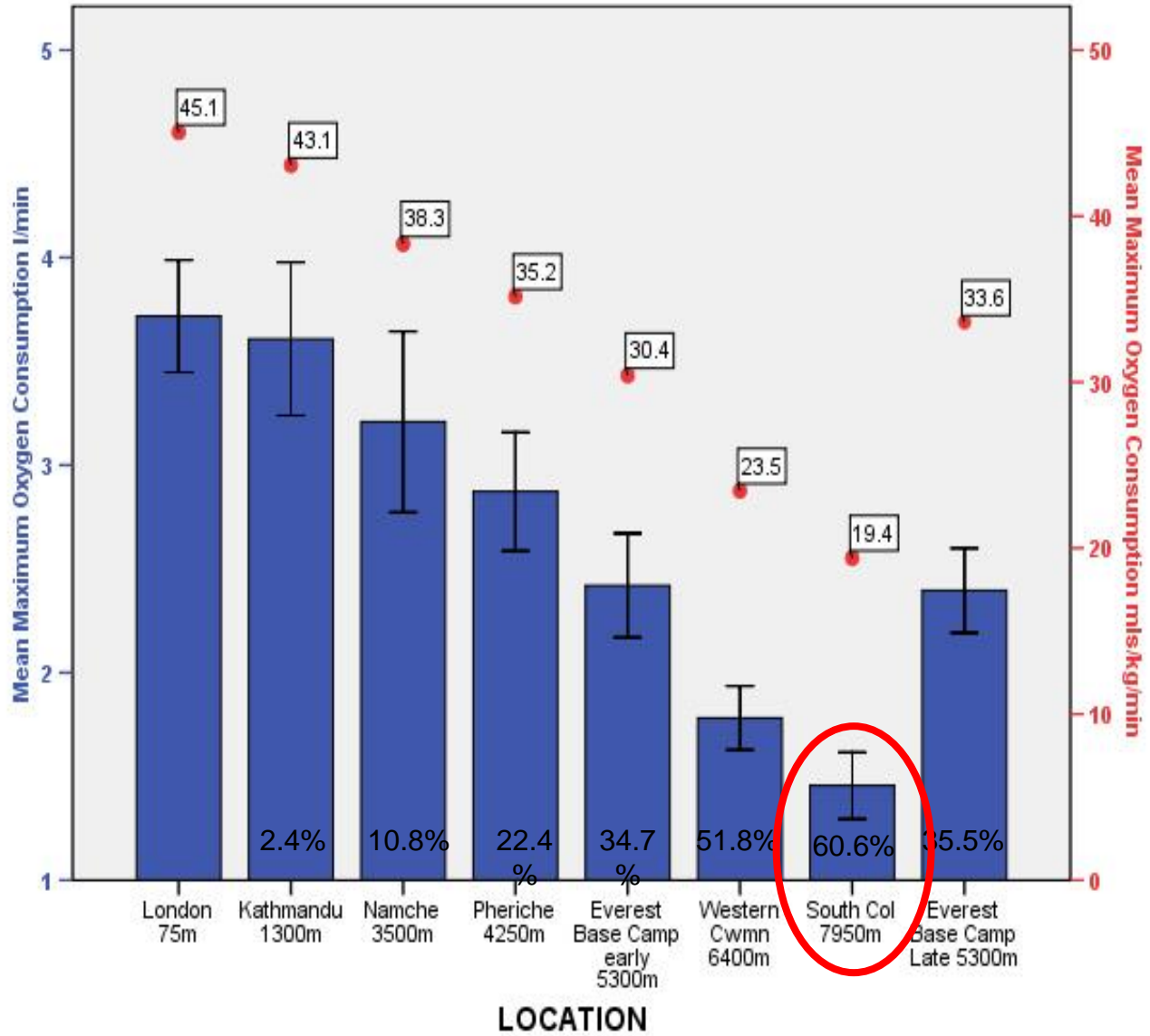
# Maximum Oxygen Consumption at Altitude, South Col Team, [REDACTED]



Error Bars: +/- 1 SD

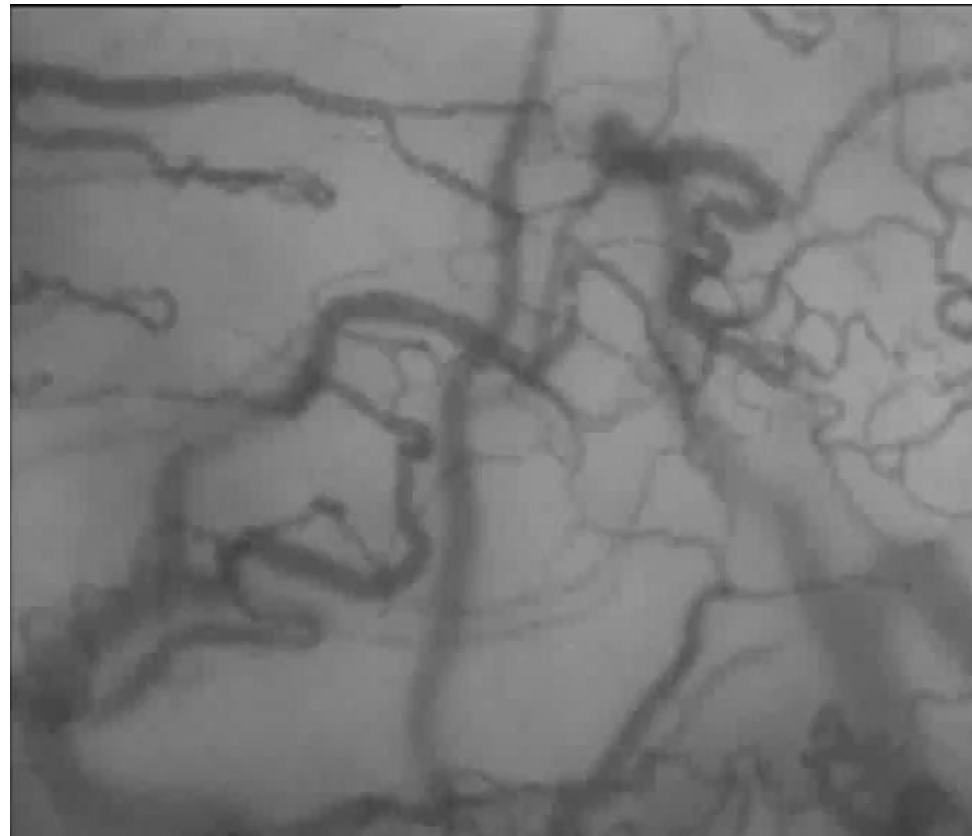
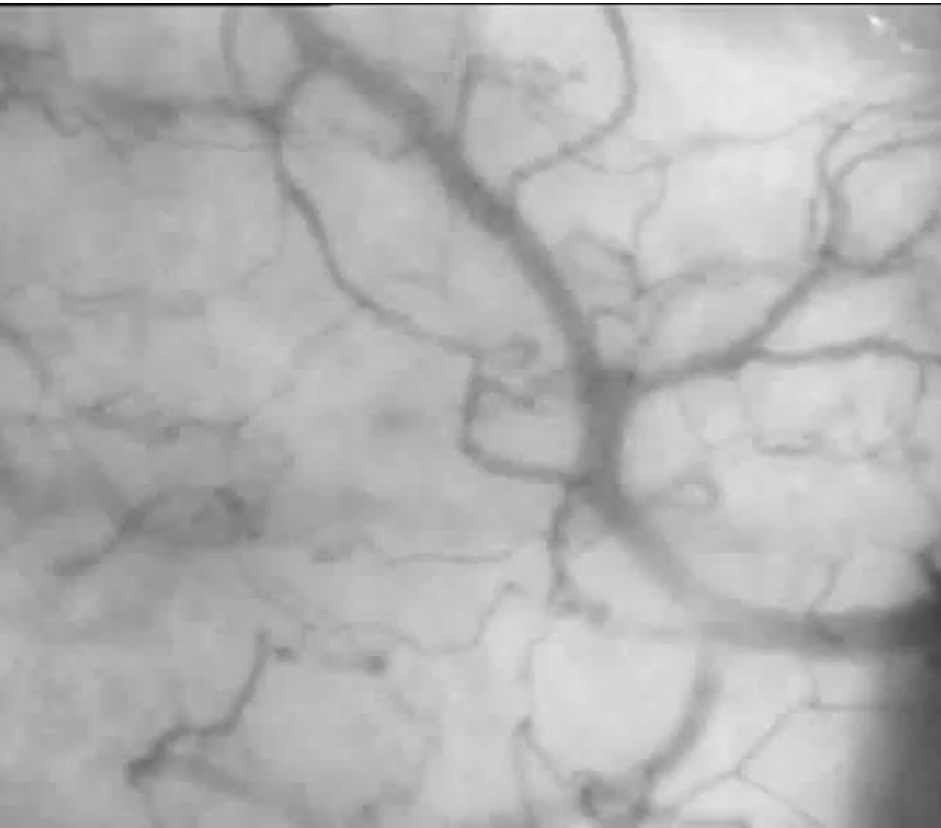


# Maximum Oxygen Consumption at Altitude, South Col Team, [REDACTED]



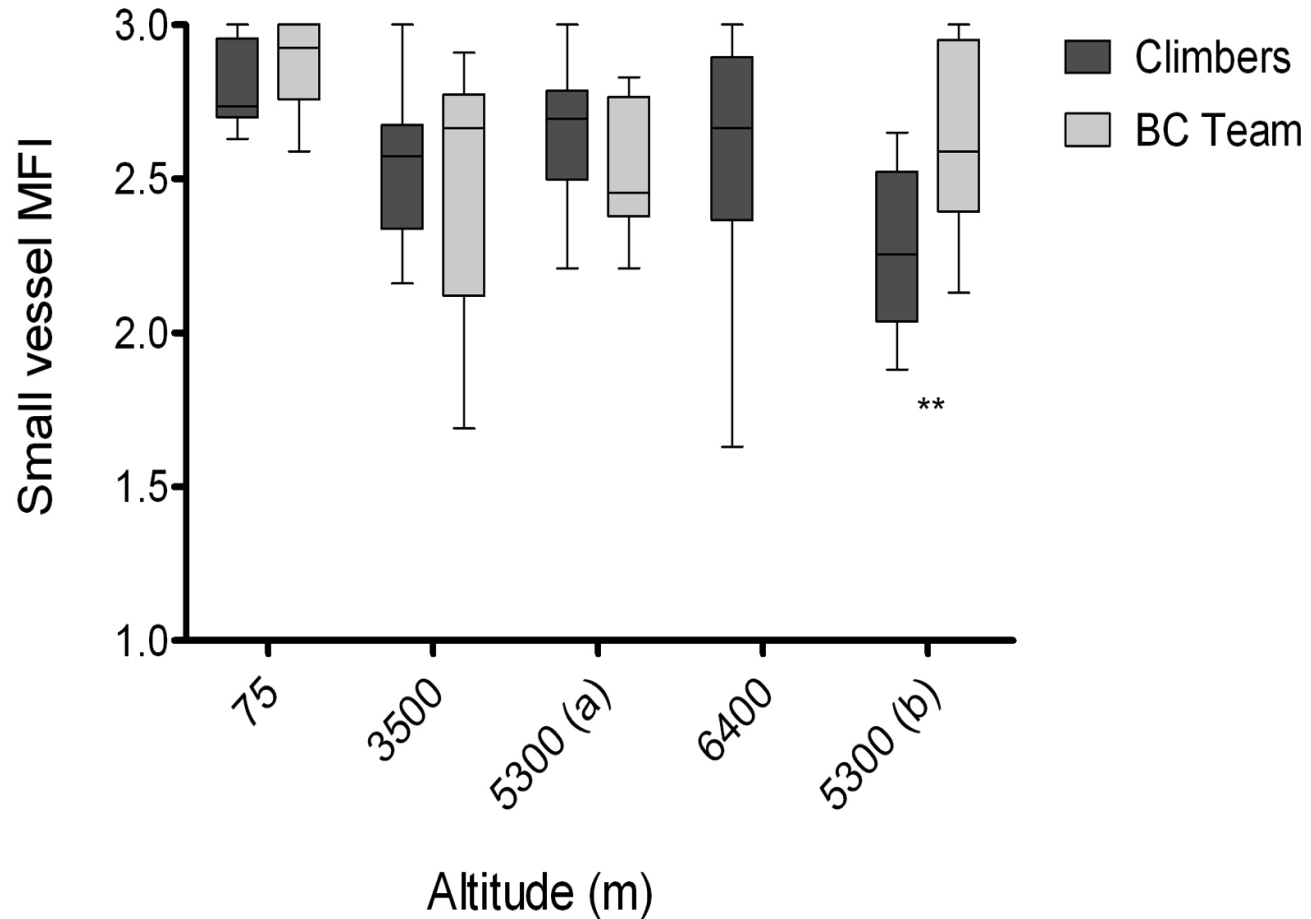
Error Bars: +/- 1 SD

# Microcirculatory dysfunction





# Microcirculation



# Metabolic basis to Sherpa altitude adaptation – Horscroft et al, 2017

- Himalayan Sherpas highly adapted to life in a low oxygen environment
- Does enhanced tissue oxygenation and delivery play a role?
- In Sherpa/Lowlander comparison, Sherpas showed lower capacity for fatty acid oxidation, enhanced efficiency of O<sub>2</sub> utilisation, improved muscular energetics, and protection against oxidative stress





# **CXE: A prospective study of the effects of environmental hypoxia on cognitive functioning – Griva et al, 2017**

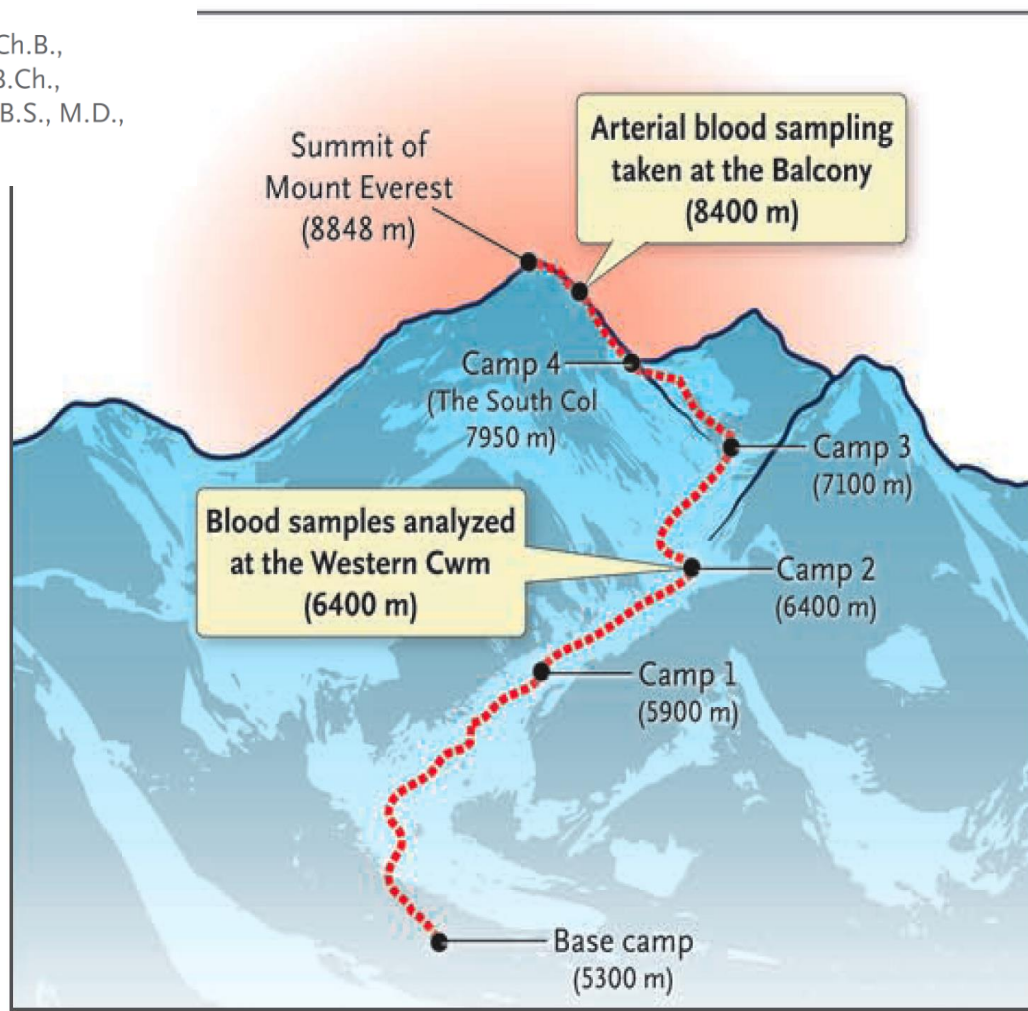
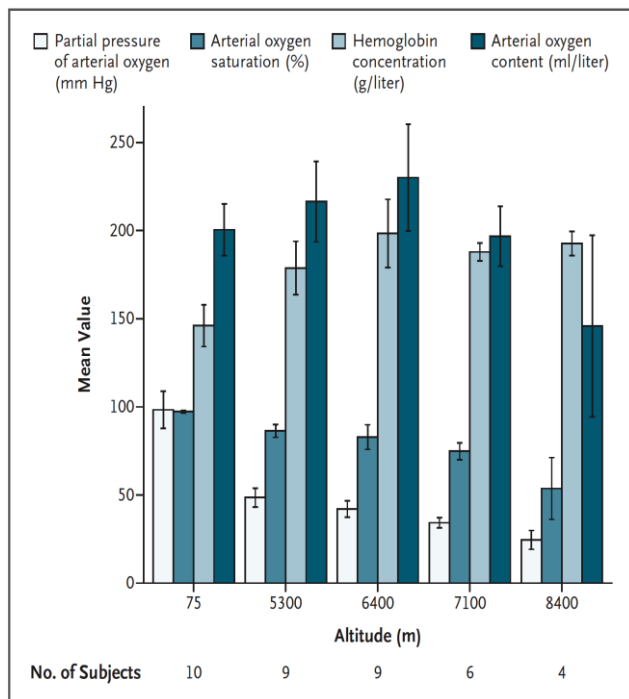
- Overall, attention, verbal ability and executive function declined in those exposed to hypoxia compared with controls
- Memory and psychomotor function showed decline at highest ascent only
- Considerable inter-individual variability in the response to sustained hypoxia
- Cognitive decline was greater amongst older people
- Not associated with mood, socio-demographic, or physiological variables



ORIGINAL ARTICLE

# Arterial Blood Gases and Oxygen Content in Climbers on Mount Everest

Michael P.W. Grocott, M.B., B.S., Daniel S. Martin, M.B., Ch.B.,  
 Denny Z.H. Levett, B.M., B.Ch., Roger McMorrow, M.B., B.Ch.,  
 Jeremy Windsor, M.B., Ch.B., and Hugh E. Montgomery, M.B., B.S., M.D.,  
 for the Caudwell Xtreme Everest Research Group\*

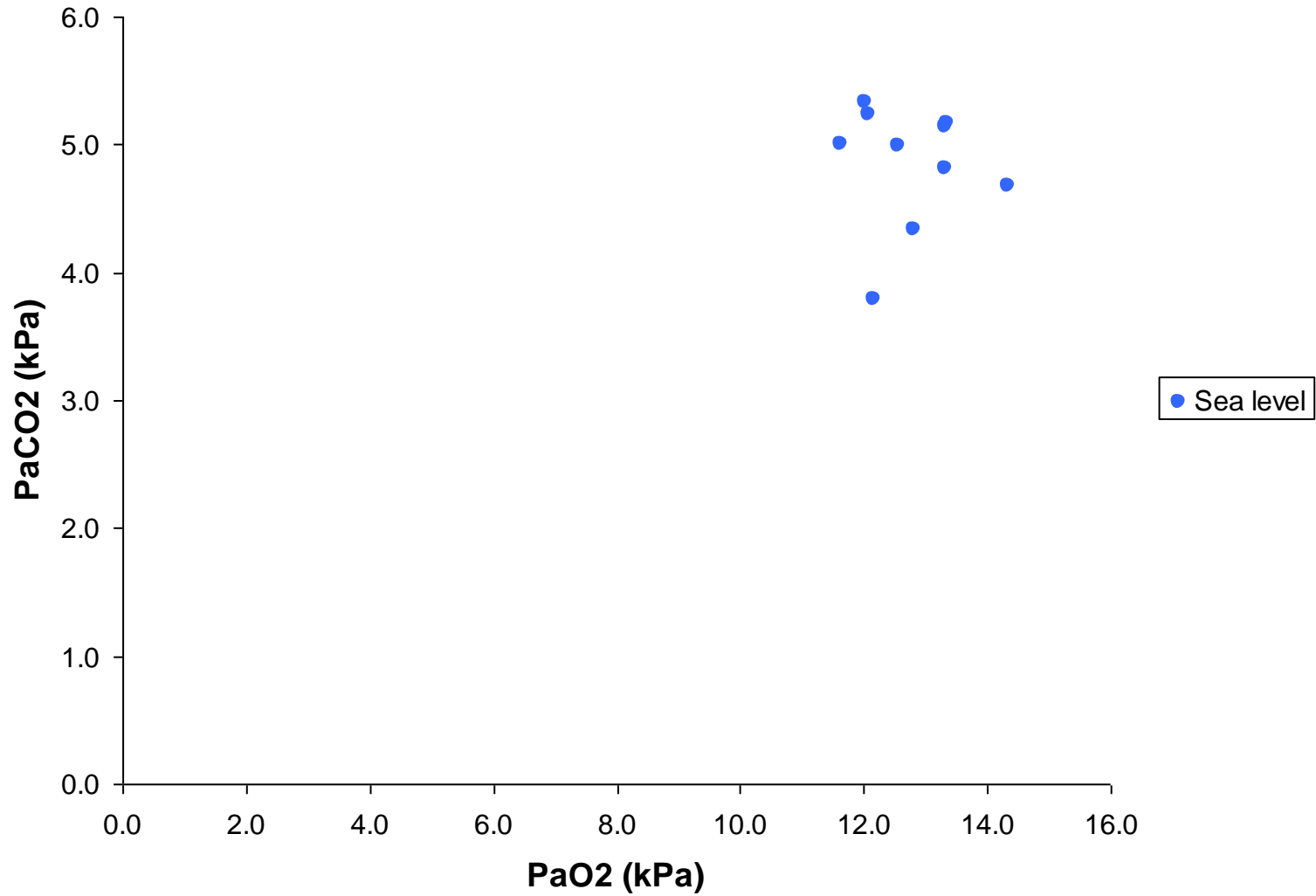


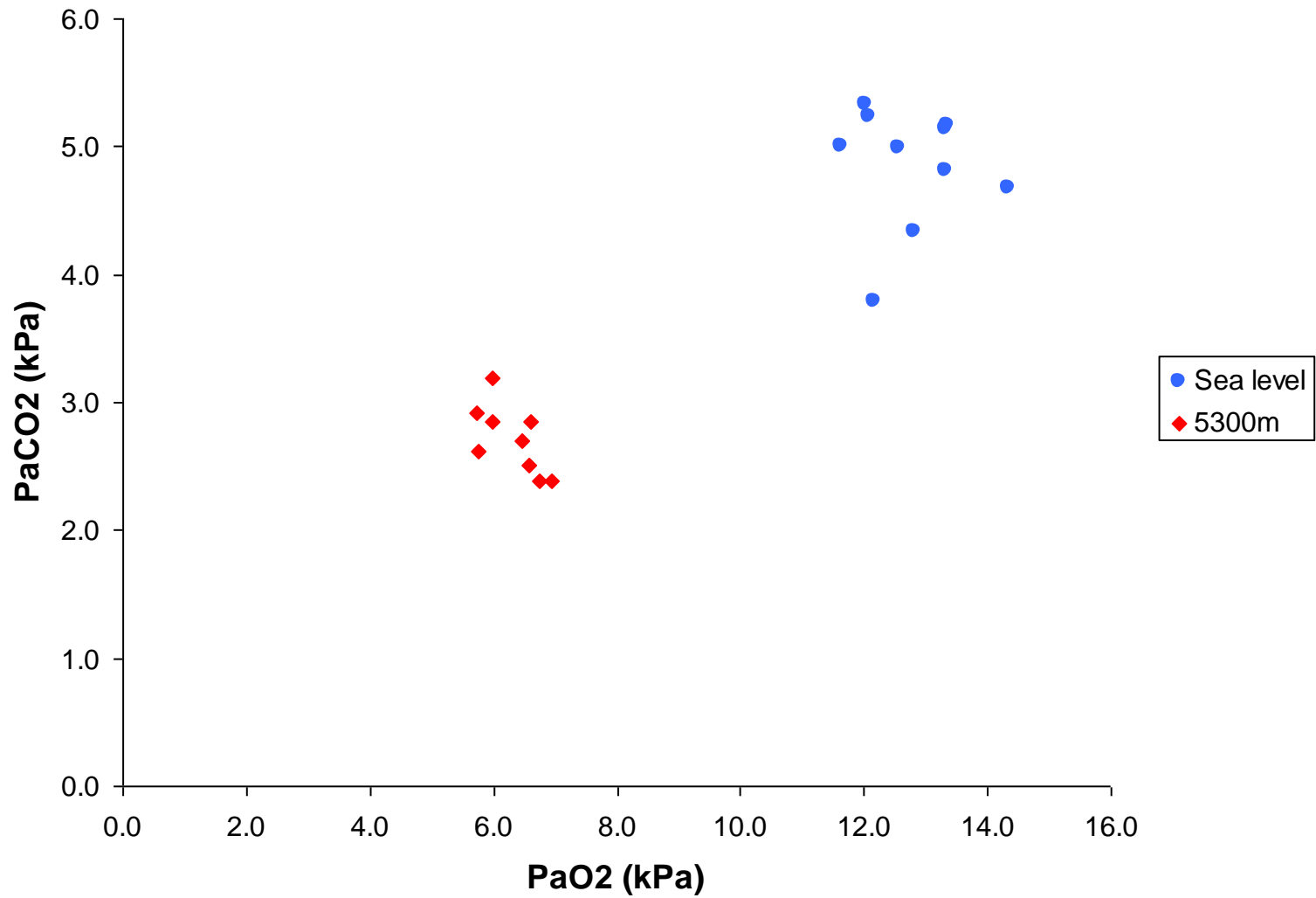




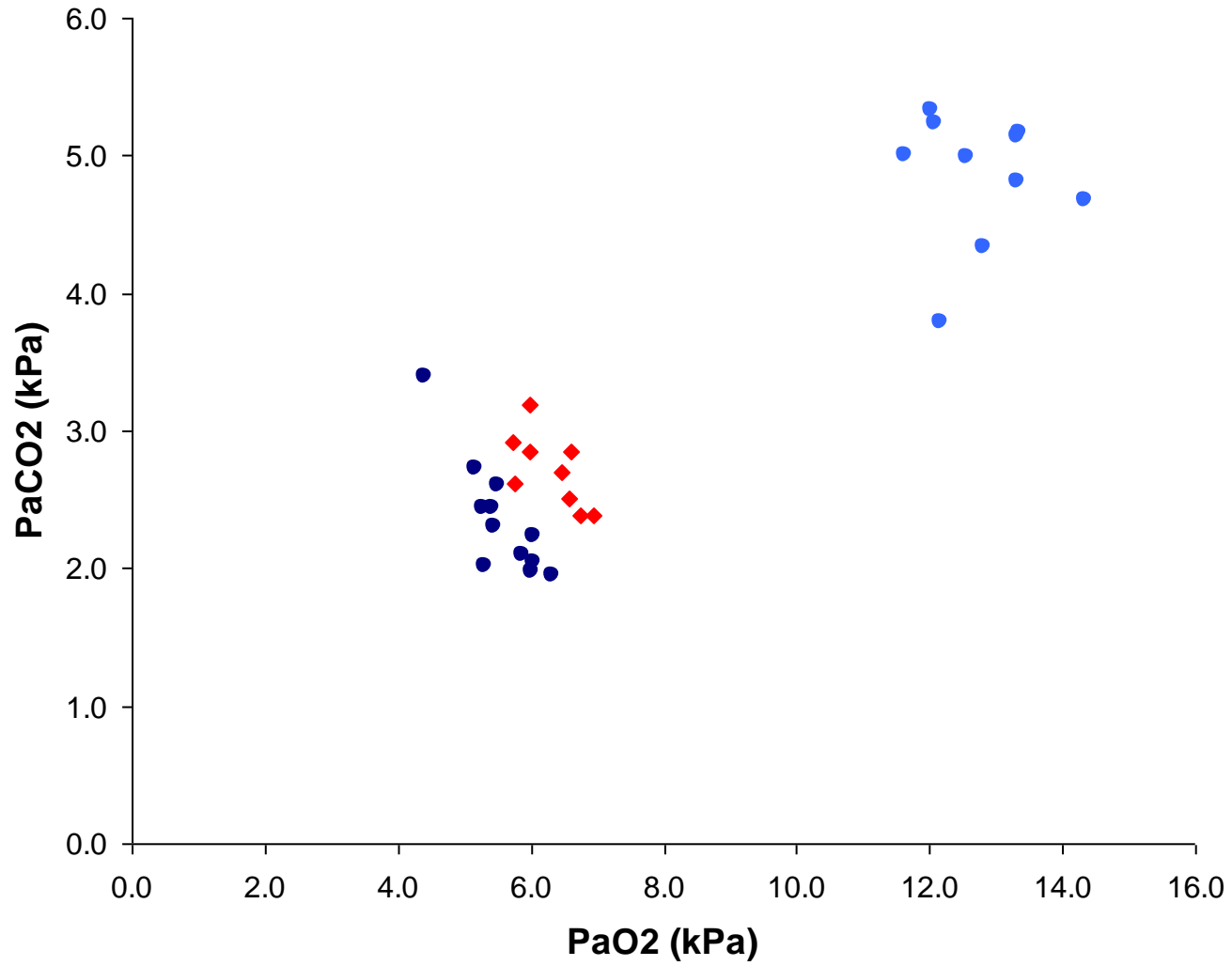


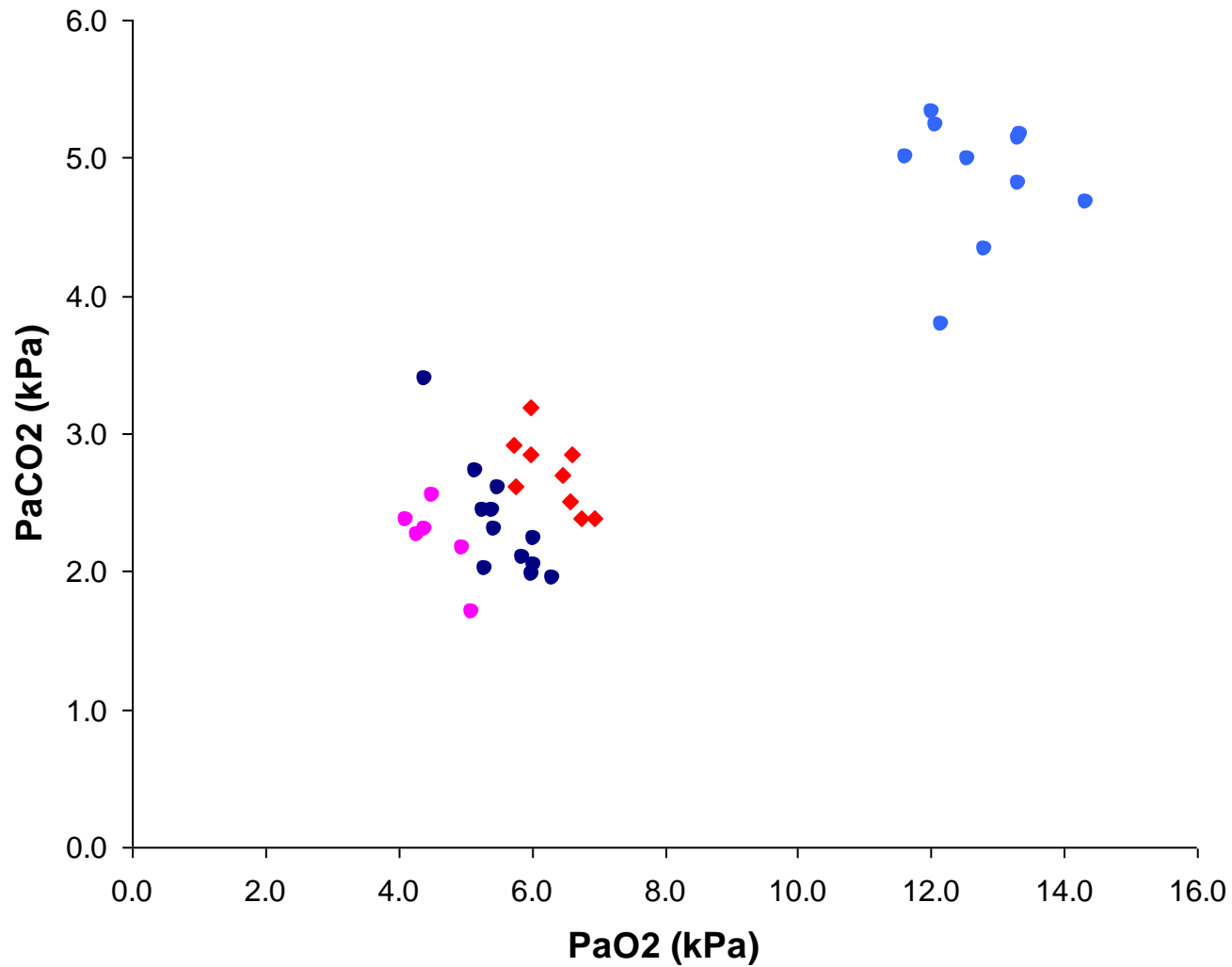


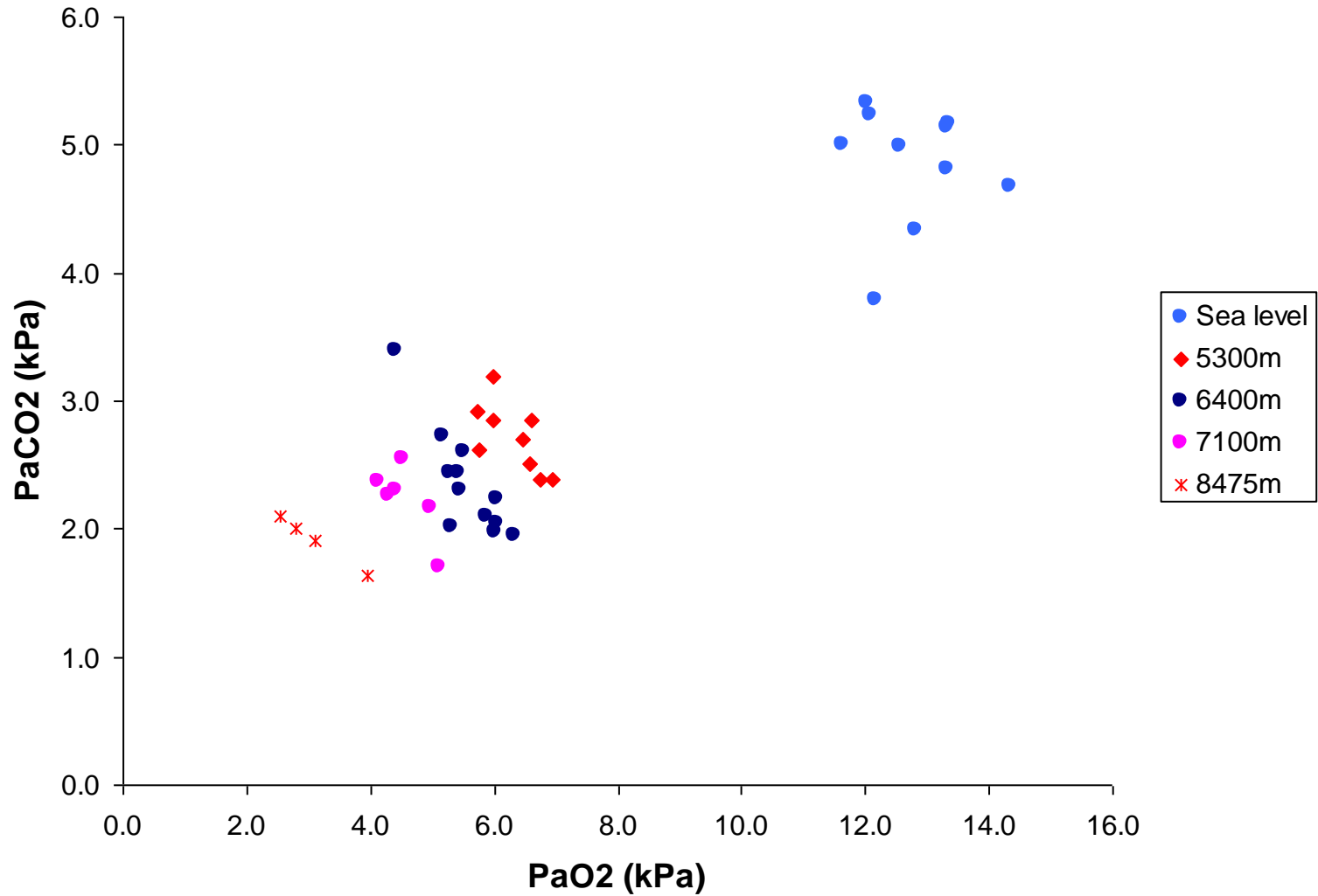




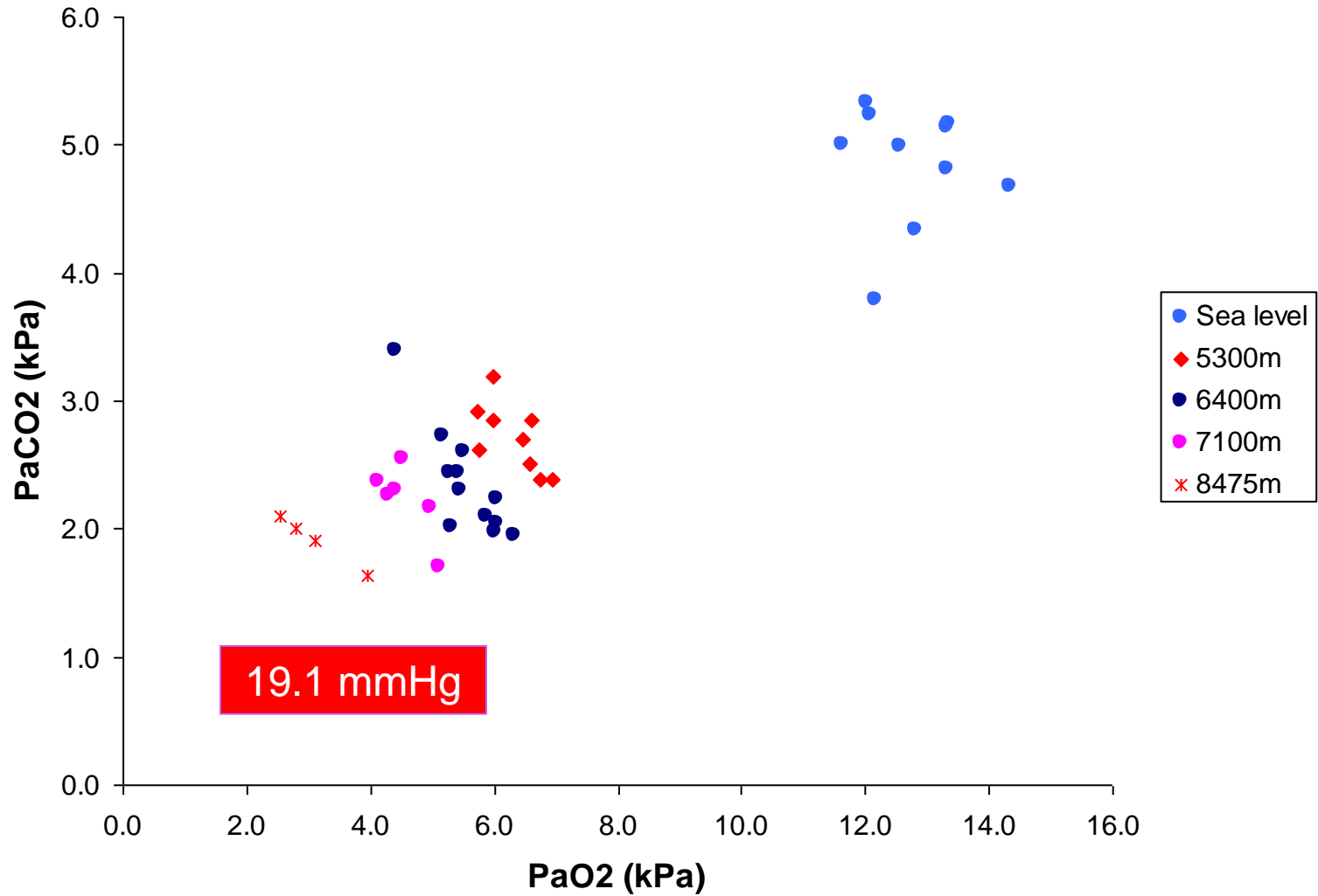












“Everest *in utero*”

$\text{PaO}_2 \sim 2.5\text{-}3.0 \text{ kPa}$

- Foetus *in utero*
- Diving Seals
- Everest Climber



**So what?**



# Implications for practice

- We can measure gross oxygen delivery
- More difficult to measure oxygen delivery at cellular level
- The two do not necessarily correlate
- Beware hyperoxia
- Good saturations do not always suggest adequate availability of oxygen at cellular level

Provide feedback  
(external link)

NB: The information displayed below does not replace the protocol. The latest protocol version should always be consulted before making clinical decisions.

Data cut:  
06 March 2017

Search for a  
study

<b>Tissue metabolism and blood flow in critically ill patients</b> An observational study in critically ill patients of the relationship between tissue blood flow and metabolism and their severity of disease and outcomes	<b>IRAS ID</b> 159977
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<b>Current Status</b>	Open, With Recruitment	<b>CPMS ID</b>	20087	<b>Geographical Scope</b>	Single Site
<b>Planned Duration</b>	04/03/2016 - 01/07/2017	<b>ISRCTN</b>	-	<b>Lead Country</b>	England
<b>Managing Specialty</b>	Critical care	<b>EudraCT</b>	-	<b>Study Setting</b>	Secondary Care
<b>All Specialties</b>	Critical care	<b>MREC No.</b>	-	<b>Portfolio eligibility route</b>	Standard
<b>Managing Specialty Primary Subspecialty</b>	Critical Care	<b>Funder(s)</b>	The Intensive Care Society, The Royal Free Charity		
<b>All SubSpecialties</b>	Critical Care	<b>Sponsor(s)</b>	ROYAL FREE LONDON NHS FOUNDATION TRUST		

### Research Summary

There remains much to be learnt regarding what determines favourable survival in critically ill patients. Traditional teaching of maintaining adequate delivery of oxygen to organs and tissues becomes ever less important as disease progresses. As our understanding of tissue metabolism and the interaction of complex cellular systems improves a concept has developed that may explain why measures of oxygen delivery frequently fail to differentiate those patients with good outcomes from those who ultimately fail to recover. The aim of our observational study of critically ill patients admitted to intensive care is to characterise three key physiological characteristics that we believe are likely to be closely interrelated and may have a significant influence on tissue oxygen balance. These mechanisms have been selected from our previous studies in which healthy volunteers were exposed to low levels of oxygen at high altitude. We aim to quantify: i) Nitric oxide activity a naturally occurring molecule that plays an important

<b>Study Type</b>	Observational	<b>England Sample Size</b>	33	<b>Chief Investigator</b>	Dr Daniel Martin
<b>Intervention Detail</b>	n/a	<b>England Recruitment to Date</b>	19	<b>Contact Details</b>	Dr Helen McKenna htmckenna@gmail.com
<b>Phase(s)</b>	N/A	<b>UK Sample Size</b>	33		
<b>Open to New Sites</b>	No	<b>UK recruitment to Date</b>	19		

### Main Inclusion Criteria

i) Patients aged 18 years and over ii) Unplanned admissions to ICU who are expected to remain there for > 5 days

### Main Exclusion Criteria

i) Severe coagulopathy ii) Therapeutic immunosuppression iii) Primary neuromuscular pathology iv) Disseminated cancer vi) Lack of permission from patient's Consultant

# TOXYC

- RfPB funded study
- Two sites recruiting
- Royal Free Hospital, London
- University Hospital Southampton
- Tight oxygen control





**HIGHLAND SHERPA RESORT**  
Pangboche 3985 mt. 13085 ft.  
**Facilities**

- Attached bathroom
- Common room
- Western woodfire stove
- Warm Dining & Great View
- Organic & Hygienic Food
- Camping Ground
- Horse Rent
- Heli Rescue
- Oxygen Service
- Wifi Service

exodus

24 HOUR  
((WiFi))  
ZONE





# Sponsors and supporters

## **Caudwell Xtreme Everest**

- BOC Medical
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- Intensive Care Foundation
- Sir Halley Stewart Trust
- NIHR UCLH BRC

## **Xtreme Everest 2**

- Smiths Medical
- UCLH Charity
- Deltex Medical
- Royal Free Charity
- London Clinic
- University Hospital Southampton
- Rhinology and Laryngology Fund
- Atlantic Customer Solutions
- Physiological society

**All grants were unrestricted**



# Ultimate team collaboration: Xtreme Everest 10 years on

- No single person is able to contribute all the necessary expertise to solve increasingly complex problems
- Research teams need a shared mission, a good organisational structure, and plenty of interpersonal interactions
- Collaborations with an element of fun are most productive

“The Scientist”, Culotta 1993



**10TH ANNIVERSARY**

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