

Himalayan powerhouses: How Sherpas have evolved superhuman energy efficiency

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Credit: Xtreme Everest

Sherpas have evolved to become superhuman mountain climbers, extremely efficient at producing the energy to power their bodies even when oxygen is scarce, suggests new research published today in the



Proceedings of the National Academy of Sciences (PNAS).

The findings could help scientists develop new ways of treating hypoxia - lack of <u>oxygen</u> - in patients. A significant proportion of patients in intensive care units (ICUs) experience potentially life-threatening hypoxia, a complication associated with conditions from haemorrhage to sepsis.

When oxygen is scarce, the body is forced to work harder to ensure that the brain and muscles receive enough of this essential nutrient. One of the most commonly observed ways the body has of compensating for a lack of oxygen is to produce more red blood cells, which are responsible for carrying blood around the body to our organs. This makes the blood thicker, however, so it flows more slowly and is more likely to clog up blood vessels.

Mountain climbers are often exposed to low levels of oxygen, particularly at high altitudes. This is why they often have to take time during long ascents to acclimatise to their surroundings, giving the body enough time to adapt itself and prevent <u>altitude</u> sickness. In addition, they may take oxygen supplies to supplement the thin air.

Scientists have known for some time that people have different responses to high altitudes. While most climbers require additional oxygen to scale Mount Everest, whose peak is 8,848m above sea level, a handful of climbers have managed to do so without. Most notably, Sherpas, an ethnic group from the mountain regions of Nepal, are able to live at high altitude with no apparent consequences to their health - as a result, many act as guides to support expeditions in the Himalayas, and two Sherpas are known to have reached the summit of Everest an incredible 21 times.

Previous studies have suggested differences between Sherpas and people



living in non-high altitude areas, known collectively as 'lowlanders', including fewer <u>red blood cells</u> in Sherpas at altitude, but higher levels of nitric oxide, a chemical that opens up blood vessels and keeps blood flowing.

Evidence suggests that the first humans were present on the Tibetan Plateau around 30,000 years ago, with the first permanent settlers appearing between 6,000-9,000 years ago. This raises the possibility that they have evolved to adapt to the extreme environment. This is supported by recent DNA studies, which have found clear genetic differences between Sherpa and Tibetan populations on the one hand and lowlanders on the other. Some of these differences were in their mitochondrial DNA - the genetic code that programmes mitochondria, the body's 'batteries' that generate our energy.

To understand the biological differences between the Sherpas and lowlanders, a team of researchers led by scientists at the University of Cambridge followed two groups as they made a gradual ascent up to Everest Base Camp at an elevation of 5,300m.

The study was part of Xtreme Everest, a project that aims to improve outcomes for people who become critically ill by understanding how our bodies respond to the extreme altitude on the world's highest mountain. This year marks 10 years since the group's first expedition to Everest.

The lowlanders group comprised 10 investigators selected to operate the Everest Base Camp laboratory, where the mitochondrial studies were carried out by James Horscroft and Aleks Kotwica, two PhD students at the University of Cambridge. They took samples, including blood and muscle biopsies, in London to give a baseline measurement, then again when they first arrived at Base Camp and a third time after two months at Base Camp. These samples were compared with those taken from 15 Sherpas, all of whom were living in relatively low-lying areas, rather



than being the 'elite' high altitude climbers. The Sherpas' baseline measurements were taken at Kathmandu, Nepal.

The researchers found that even at baseline, the Sherpas' mitochondria were more efficient at using oxygen to produce ATP, the energy that powers our bodies.

As predicted from genetic differences, they also found lower levels of fat oxidation in the Sherpas. Muscles have two ways to get energy - from sugars, such as glucose, or from burning fat (fat oxidation). The majority of the time we get our energy from the latter source; however, this is inefficient, so at times of physical stress, such as when exercising, we take our energy from sugars. The low levels of <u>fat oxidation</u> again suggest that the Sherpas are more efficient at generating energy.

The measurements taken at altitude rarely changed from the baseline measurement in the Sherpas, suggesting that they were born with such differences. However, for lowlanders, measurements tended to change after time spent at altitude, suggesting that their bodies were acclimatising and beginning to mimic the Sherpas'.

One of the key differences, however, was in phosphocreatine levels. Phosphocreatine is an energy reserve that acts as a buffer to help muscles contract when no ATP is present. In lowlanders, after two months at high altitude, phosphocreatine levels crash, whereas in Sherpas levels actually increase.

In addition, the team found that while levels of free radicals increase rapidly at high altitude, at least initially, levels in Sherpas are very low. Free radicals are molecules created by a lack of oxygen that can be potentially damaging to cells and tissue.

"Sherpas have spent thousands of years living at high altitudes, so it



should be unsurprising that they have adapted to become more efficient at using oxygen and generating energy," says Dr Andrew Murray from the University of Cambridge, the study's senior author. "When those of us from lower-lying countries spend time at high altitude, our bodies adapt to some extent to become more 'Sherpa-like', but we are no match for their efficiency."

The team say the findings could provide valuable insights to explain why some people suffering from hypoxia fare much worse in emergency situations that others.

"Although lack of oxygen might be viewed as an occupational hazard for mountain climbers, for people in intensive care units it can be life threatening," explains Professor Mike Grocott, Chair of Xtreme Everest from the University of Southampton. "One in five people admitted to intensive care in the UK each year die and even those that survive might never regain their previous quality of life.

"By understanding how Sherpas are able to survive with low levels of oxygen, we can get clues to help us identify those at greatest risk in ICUs and inform the development of better treatments to help in their recovery."

The 10th anniversary of the original Caudwell Xtreme Everest expedition will be marked this month by a conference at the Royal Society of Medicine, and an event open to the public on the evening of 23rd May at the Royal Geographical Society entitled A Celebration of Six Decades of Medicine on Everest.

More information: James A. Horscroft el al., "Metabolic basis to Sherpa altitude adaptation," *PNAS* (2017). <u>www.pnas.org/cgi/doi/10.1073/pnas.1700527114</u>



Provided by University of Cambridge

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