

To have or not to have ribs (a vertebrate story)

April 27 2010



Mice have a ribcage with 12 pairs of ribs (image on the left). When the gene Hox6 is activated in ribless regions of the embryo, extra ribs are formed, extending from the ribcage to the tail (image on the right). Credit: Image provided by Moises Mallo.

Like all vertebrates, snakes, mice and humans have in common a skeleton made of segments, the vertebrae. But a snake has between 200-400 ribs extending from all vertebrae, from the neck to the tail-end, whereas mice have only 13 pairs of ribs, and humans have 12 pairs, in both cases making up the ribcage. In the latest issue of *Developmental Cell*, researchers from the Instituto Gulbenkian de Ciencia, in Portugal, reveal that, contrary to what was thought, making ribs is not the default state for vertebrates, but is actually an active process of balancing the activities of a remarkable class of genes - the Hox genes.



It was thought that the rib less region of the mouse embryo was the result of a rib-inhibiting programme, driven by Hox10 genes. Indeed, previous studies, in which Hox10 genes were inactivated in the embryo, generated mice with extra ribs. However, by forcing another class of Hox genes (Hox6) to be activated in future rib-less regions of the mouse embryo, Moises Mallo and his team bred mice that also have extra ribs, both in the neck area, and from just after the rib cage, all the way down to the tail, resembling a snake-like <u>skeleton</u>.

'It was an extraordinary, and clear-cut result', says Mallo, 'suggesting that these two groups of Hox genes balance each other out: one actively promotes rib formation to produce the thoracic region, while the other blocks this activity in the lumbar region. Our results have unveiled this balance.'

The researchers went on to unpick the genes involved in this process, and came up with yet another surprising finding: that the whole process relies on first hitting so-called muscle genes in the embryo, which then provide signals to switch on the 'rib' genes to make both ribs and muscle, in a coordinated process.

According to Mallo, 'Our findings reveal a more complicated process than we would have imagined, but one that makes perfect sense, from a functional and evolutionary point of view: it is no good to make ribs without muscle, so, in the embryo, the production of both <u>ribs</u> and their associated muscles is under the control of a single and coordinated mechanism.

Provided by Instituto Gulbenkian de Ciencia

Citation: To have or not to have ribs (a vertebrate story) (2010, April 27) retrieved 6 May 2024 from <u>https://medicalxpress.com/news/2010-04-ribs-vertebrate-story.html</u>



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