

Researchers dig up new molecular details on 'the other type' of stem cells

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Nuclei of Drosophila trachea cells during the transition from larvae to an adult fly. The cells in the Tr2 segment are dividing; they are the facultative stem cells. Credit: N.J. Djabrayan, IRBBarcelona

Scientists at IRB Barcelona and CSIC have discovered that the combination of two molecular signals determines which cells that have already differentiated can regain their stem cell properties.



Their studies on fruit flies advance the field of regenerative medicine and confer a better understanding of processes involved in cancer.

In a study published this week in *PLos Genetics*, scientists Nareg J. Djabrayan and Jordi Casanova from the Institute for Research in Biomedicine (IRB Barcelona) and CSIC, have identified two <u>molecular</u> signals and the pathway of events that allows cells in a tissue that are already specialized to undifferentiate into <u>stem cells</u>. The study offers new information about how cells become differentiated and how this type of facultative stem cell is activated, which is of particular interest in cell reprogramming, <u>regenerative medicine</u>, and in understanding cancer.

Facultative stem cells are increasingly being identified in human tissues and organs, but little is known about them compared to typical stem cells, which have distinct morphological traits. "The existence of facultative stem cells indicates that many differentiatied cells continue to have great plasticity and are able to respond in situations of stress, such as in wounds or in the regeneration of worn out tissues," explains Jordi Casanova, head of the Development and Morphogenesis in Drosophila group at IRB Barcelona.





Drosophila trachea fragment. Externally, there is no difference between the Tr2 segment, where facultative stem cells are found, and Tr3, which indicates the rest of the cells in the tissue. Credit: N.J. Djabrayan, IRBBarcelona

"This plasticity is good if it is well regulated. If any of the mechanisms involved become unbalanced, it becomes dangerous," adds Casanova. "In fact, cellular plasticity in tissues that regenerate often are associated with organs that have a high incidence of cancer, such as the colon or blood," he explains.

A combination of signals in space and time



The researchers looked at the moment when a Drosophila larva changes into an adult fly. "This model is a good one to identify and study facultative stem cells because there are some cells in the larva that will give rise to the adult fly and they get activated during the switch from larva to adult, a transition which could be considered a condition of stress," explains Casanova.

The researchers studied cells with facultative stem cell traits in Drosophila trachea. They showed that only some cells in the tissue receive a spatial signal that marks them. Then, when the transition occurs, a second signal is triggered by hormones that prompts the reactivation of the stem cell programme. This signal reaches all cells in the tissue, but only those that have been marked previously with the spatial signal are reactivated as stem cells.

"Understanding these signal regulation mechanisms is fundamental to controling the reprogramming of cells," indicates Casanova. Using the same model, the scientists now want to investigate which mechanisms allow <u>differentiated cells</u> to maintain their plasticity and how they go from being a differentiated cell to a stem cell and vice versa.

More information: Nareg J.-V. Djabrayan et al. Snoo and Dpp Act as Spatial and Temporal Regulators Respectively of Adult Progenitor Cells in the Drosophila Trachea, *PLOS Genetics* (2016). <u>DOI:</u> <u>10.1371/journal.pgen.1005909</u>

Provided by Institute for Research in Biomedicine (IRB Barcelona)

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