

Cell senescence does not stop tumor growth

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Since cancer cells grow indefinitely, it is commonly believed that senescence could act as a barrier against tumor growth and potentially be used as a way to treat cancer. A collaboration between a cancer biologist from the University of Milano, Italy, and two physicists, from the National Research Council of Italy and from Cornell University, has shown that cell senescence occurs spontaneously in melanoma cells, but does not stop their growth, which is sustained by a small population of cancer stem cells. The results, published in the open-access journal *PLoS Computational Biology* on January 19 explain why it is difficult to treat cancer cells by inducing senescence alone.

The work explores the relationship between melanoma and senescence, the normal process where cells decline and eventually stop duplicating after reaching maturity. The investigators followed the long-term evolution of melanoma <u>cell populations</u>, monitoring the number of senescent cells. After three months, growth slowed and most of the cells turned senescent, however growth did not stop and eventually resumed its initial rate until the <u>senescent cells</u> had almost disappeared.

The authors mathematically modeled the experimental data using the cancer stem cell hypothesis, where a sub-group of <u>cancer cells</u> replicate indefinitely, and are thus unaffected by senescence. These cancer stem cells give rise to a larger population of cancer cells that can duplicate only a finite number of times. The model yielded an indirect confirmation of the presence of cancer stem cells in melanoma, an issue that is still controversial in the cancer research community.



Although a large fraction of cancer cells are susceptible to senescence, the researchers conclude that inducing senescence is unlikely to provide a successful therapeutic strategy because these cells are irrelevant for tumor growth. However, the indirect evidence of cancer stem cells in melanoma may enable the development of new methods to treat specific kinds of cancer. The challenge will be in the strong resistance to drug induced senescence that would be found in the cancer stem cells. Along this line of research, treatment of tumors would focus on targeting only these cancer stem cells, rather than every single cancerous cell.

More information: La Porta CAM, Zapperi S, Sethna JP (2012) Senescent Cells in Growing Tumors: Population Dynamics and Cancer Stem Cells. PLoS Comput Biol 8(1): e1002316. <u>doi:10.1371/journal.pcbi.1002316</u>

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