

Why are some cells more cancer prone?

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Cells in the body wear down over time and die. In many organs, like the small intestine, adult stem cells play a vital role in maintaining function by replacing old cells with new ones. Learning about the nature of tissue stem cells can help scientists understand exactly how our organs are built, and why some organs generate cancer frequently, but others only rarely.

New work from Carnegie's Alexis Marianes and Allan Spradling used some of the most experimentally accessible tissue <u>stem cells</u>, the adult stem cells in the midsection of the fruit fly gut, with surprising results. Their findings are published by *eLife*.

Like the small intestine in mammals, the midgut of <u>fruit flies</u> is where most digestion takes place. Scientists had noticed a few regions in both the midgut and <u>small intestine</u> were specialized for certain tasks, such as absorbing iron, but had little understanding of the extent of these regional differences or how they were maintained.

Marianes and Spradling were able to demonstrate that there are 10 different major subregions within the fruit fly midgut. They occur in a specific order and each is responsible for different digestive and nutrient-storage processes, as evidenced by the expression of many specific genes. Most importantly, the <u>adult stem cells</u> in each region are specialized as well, and only support the types of cells found within it. Thus, during development, achieving the right spatial sequence of stem cells is probably critical to causing intestines to be built and maintained in order to function optimally.



The researchers also showed that tumors arise preferentially in specific regions of the midgut, a phenomenon well known in oncology. They showed the tumor-prone regions were specialized for lipid absorption, and stem cell function in them differed in small ways from stem cell function in other regions.

This work will motivate the search for fine-grained specialization in both tissue organization and in stem cells within many mammalian tissues. These subtle differences may explain the surprising results that are sometimes obtained following the removal or transplantation of human tissue. This must be considered carefully in ongoing attempts to utilize stem cells therapeutically.

Finally, it may be possible to learn what makes some stem cells more susceptible to cancer than others, and develop strategies to counteract this tendency.

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