Researchers pinpoint hierarchy of breast cancer cells as potential cause for treatment resistance
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"Our recent findings emphasize the need to account for the specific cell states that are present within a tumor," says Yeo, who is a member in the lab of Jun-Lin Guan, Ph.D., the Francis Brunning Endowed Chair and professor of cancer biology. "This could potentially help determine the combination of drugs that are required to eliminate all the cell states that are present to eliminate treatment resistance."

Yeo says that when it comes to breast cancers, it is known that cells within a tumor are varied.

"This diversity poses a problem to treating patients because particular subsets of tumor cells may be drug resistant and eventually lead to disease recurrence," he says. "One of the factors contributing to this diversity is the fact that tumor cells can exist in different cellular states, ranging from more stem-like cells that can become other cell types to more differentiated cells that have been coded to serve a purpose, or do a certain 'job' within the system.

"Cancer cells with stem-like properties are known to cause drug resistance, and they are generally seen as being at the top of the tumor hierarchy, like the king or queen of the village, with more differentiated tumor cells towards the bottom of the hierarchy, like the common townspeople."

In this study, researchers used breast cancer animal models to determine tumor hierarchies beyond 'ruler' and 'common people' cells, Yeo says. They identified and categorized singular cells which helped them understand each, individual cell's purpose. Yeo adds that bulk tumor cell analysis would have masked the cellular details.

"We were able to find a complex spectrum of cell states between different tumor types that can range
from stem-cells to the 'beginner cells' to more
differentiated cells," he says. "In our village
scenario, these would be the governors and
mayors, followed by the common townspeople.
Furthermore, depending on the lineage of the
tumor, some may show a spectrum of cell states
that are higher up in the hierarchy and vice versa.

"These findings are important because they show
we need to know more about how these specific
cell states contribute to tumor growth so we can
target them with combination drug therapies,
potentially helping more people who may otherwise
experience drug resistance."

More information: Syn Kok Yeo et al, Single-cell
RNA-sequencing reveals distinct patterns of cell
state heterogeneity in mouse models of breast

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