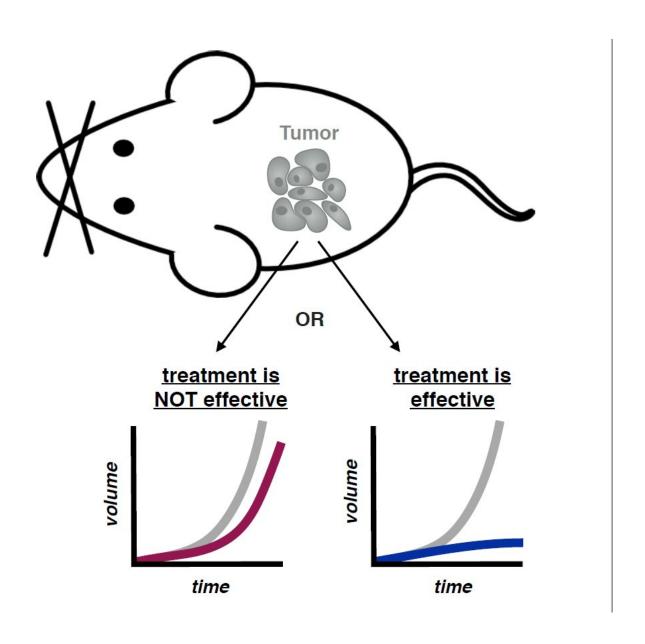


Tumor growth parameters predict response to anti-angiogenic therapy in mice

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Properties related to how a tumor grows can be used to predict the response to



anti-angiogenic treatment that targets VEGF. Gray lines, tumor growth with no treatment; Red line, treatment yields poor response; blue line, treatment effectively reduces tumor growth. Credit: Stacey D. Finley, Ph.D.

Using a mathematical modeling approach, scientists have found that certain parameters of tumor growth in mice can predict the effectiveness of drugs that block formation of tumor-nourishing blood vessels. The findings are published in *PLOS Computational Biology*.

Even within the same cancer type, different tumors can respond differently to cancer drugs, and some tumors do not respond at all. Therefore, it is useful to identify <u>tumor</u> features that can serve as "predictive biomarkers" to forecast how a person's tumor might respond to a particular treatment. However, before drugs are tested in humans in a clinical setting, they are usually explored in mice.

In efforts to enhance pre-clinical drug testing, Thomas Gaddy of the University of Southern California, Los Angeles, and colleagues in the research group of Professor Stacey Finley investigated a potential predictive biomarker for a type of cancer drug known as anti-angiogenic therapy. This treatment inhibits the formation of new <u>blood vessels</u> that can help a tumor grow.

The researchers had previously built a mathematical model that simulates the activity of a blood-vessel-growth-promoting protein called VEGF in a mouse tumor. For the new study, they added a component to the model that simulates the tumor's response to drugs that target VEGF and inhibit its activity. They trained and validated the model using real-world data on tumor growth in mice.

Using their new model, the researchers found that certain parameters



about the way a tumor grows could successfully and accurately predict its response to anti-angiogenic treatment that targets VEGF activity. Using those characteristics, they were able to predict how effective the treatment would be (i.e., whether tumor growth would slow), even before the start of treatment.

If these results are confirmed in live mouse studies, they could eventually lead to new strategies to predict <u>tumor response</u> to antiangiogenic therapy in people. Meanwhile, the research team is using their model to simulate such a study.

"In our virtual mouse study, we will simulate tumor growth with and without treatment and determine if the growth parameters can indeed be used to distinguish between mice whose tumors will respond to therapy or not," Finley says. "We are also scouring published literature to find more experimental data to further validate our <u>model</u> predictions."

More information: Gaddy TD, Wu Q, Arnheim AD, Finley SD (2017) Mechanistic modeling quantifies the influence of tumor growth kinetics on the response to anti angiogenic treatment. *PLoS Comput Biol* 13 (12): e1005874. doi.org/10.1371/journal.pcbi.1005874

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