

Brain cancer vaccine effective in some patients

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Glioblastoma (histology slide). Credit: Wikipedia/CC BY-SA 3.0

Most people with the deadly brain cancer glioblastoma die less than 18 months after diagnosis. But a multicenter clinical trial of a personalized vaccine that targets the aggressive cancer has indicated improved survival rates for such patients.



The study appears May 29 in the Journal of Translational Medicine.

The phase three clinical trial included 331 patients at over 80 sites in four countries. Among the trial participants was Siteman Cancer Center at Barnes-Jewish Hospital and Washington University School of Medicine in St. Louis, which recruited one of the largest groups of patients in the trial. The patients were randomized to receive standard therapy plus the personalized vaccine, called DCVax-L, or standard therapy plus a placebo. Personalized vaccines are specifically tailored to individual patients.

According to the trial design, the vaccine group had twice the number of patients as the placebo group. Further, any patient in the trial was allowed to receive the vaccine if his or her cancer recurred or progressed after initial treatment. As such, almost 90 percent of all participants received the investigational vaccine.

The trial, which was funded by the vaccine maker, Northwest Biotherapeutics Inc., is ongoing to allow for continued study of patients who are living beyond what is expected of those diagnosed with glioblastoma.

As of the analysis detailed in the study, all 331 patients in the trial (including those who did and did not receive the vaccine) had a median survival of just over 23 months. One-hundred patients had an average overall survival of 40.5 months—more than three years—and were designated as "extended survivors." Investigators are particularly interested in the latter group's response to the vaccine, though they don't yet know whether everyone in this group received the vaccine since the trial is ongoing and remains blinded. The continued blinding means neither the patients nor physicians know which patients are receiving the vaccine. But since the researchers reported that almost 90 percent of all participants received the vaccine at some point, the chances that the



extended survivors are getting the vaccine, rather than placebo, are high.

"The overall patient population in the trial appears to live longer than we would typically see with current standard of care, and 30 percent of the patients have lived much longer than we would expect, given the typical course of this cancer," said oncologist Jian L. Campian, MD, Ph.D., one of the study's authors and a Washington University assistant professor of medicine who treats patients at Siteman Cancer Center. "In general, patients with this cancer live 15 to 17 months. The surprising part was that the 100 'extended survivors' don't appear to have the usual characteristics associated with a good prognosis. We are continuing to study these patients to understand why they have done so well."

The <u>personalized vaccine</u> used was specific to each patient. After surgery to remove as much of the tumor as possible, a small amount of tumor tissue is processed and then exposed to the same patient's own immune cells, called dendritic cells. Exposure to the tumor material essentially trains the dendritic cells to seek out and destroy tumor cells. These trained <u>dendritic cells</u> are returned to the patient as a vaccine injected in the arm. Relatively few vaccinations are needed: The first three are weeks apart; the next three are months apart; and then they continue with one vaccination every six months after the first year.

Campian said the vaccine had very few side effects, especially compared with standard treatment for this <u>cancer</u>, which includes surgery, radiation and chemotherapy. About 2 percent of participants (seven <u>patients</u>) experienced a serious adverse event—such as brain swelling or seizures—that may have been related to the <u>vaccine</u>, according to the researchers.

More information: Liau L, et al. First results on survival from a large phase 3 clinical trial of an autologous dendritic cell vaccine in newly diagnosed glioblastoma. *Journal of Translational Medicine*. May 29,



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