Vepdegestrant (vep-DEG-eh-strent),

a PROTAC ER degrader, in combination with PF-07220060 in people living with ER+/HER2- advanced breast cancer

This summary contains information from the scientific poster:

TACTIVE-K: Phase 1b/2 Study of Vepdegestrant, a PROteolysis TArgeting Chimera (PROTAC) Estrogen Receptor Degrader, in Combination with PF-07220060, a Cyclin-Dependent Kinase 4 Inhibitor, in Estrogen Receptor+/Human Epidermal Growth Factor Receptor 2- Advanced Breast Cancer

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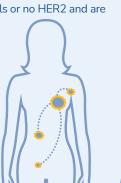
What is ER+/HER2- advanced breast cancer?

ER+/HER2- breast cancer is a specific type of breast cancer

- Certain types of breast cancer grow in response to estrogen, a hormone in the body. This is called estrogen receptor-positive (ER+) breast cancer
- Some types of breast cancer have a lot of a protein called human epidermal growth factor receptor 2 (HER2) and are called HER2-positive (HER2+). Other breast cancer types have low levels or no HER2 and are called HER2-negative

(HER2-)

Advanced breast cancer is cancer that has spread from the breast to nearby tissue (locally advanced cancer) or from the breast to more distant parts of the body (metastatic cancer)



What are some common treatments for ER+/HER2advanced breast cancer?

Some treatments, called **endocrine therapies**, work by either blocking the body's ability to produce estrogen, or blocking the activity of estrogen in cancer cells, which may slow or stop cancer growth

- Aromatase inhibitors, such as letrozole, anastrozole, or exemestane, are endocrine therapies that reduce the production of estrogen
- **Fulvestrant** is an endocrine therapy that attaches to estrogen receptors and blocks their activity, which reduces estrogen's effects on tumors

Chemotherapy is a treatment that damages cancer cells. Sometimes people get chemotherapy prior to surgery to shrink the size of their tumor, after surgery to kill remaining cancer cells, or if their cancer has spread beyond the breast
CDK4/6 inhibitors are another type of treatment and work by blocking CDK4 and CDK6 proteins that cause cancer cells to grow

What is vepdegestrant?

Vepdegestrant, also called **ARV-471**, is an investigational drug taken by mouth as a pill that researchers are testing for the treatment of ER+ breast cancer. It is a **PROteolysis TArgeting Chimera (PROTAC) estrogen receptor degrader**

- PROTAC protein degraders are designed to attach to specific proteins in cells that can cause disease, which causes those proteins to be marked for elimination by a natural protein disposal system in the body
- Vepdegestrant works by causing estrogen receptors to be eliminated, which blocks the activity of estrogen and may stop ER+ breast cancer tumors from growing or cause the tumors to shrink

Why is the combination of vepdegestrant plus PF-07220060 being tested in this study?

- PF-07220060 is an investigational drug that works by blocking the action of the CDK4 protein. This may help stop certain cancers from growing
- PF-07220060 is being studied as a treatment for ER+ breast cancer
- In laboratory research studies, vepdegestrant plus PF-07220060 worked better at stopping cancer from being able to grow than vepdegestrant or PF-07220060 alone

This summary describes a new clinical study to test the combination of vepdegestrant plus **PF-07220060** in people living with ER+/HER2- advanced breast cancer

The **main aims** of this study are to find out

This study will also look at

The side effects people who take vepdegestrant plus PF-07220060 may experience The best dose(s) of vepdegestrant plus PF-07220060 to use for future clinical studies

- How long people receiving these treatments live without their cancer getting worse
- The percentage of people whose cancer shrinks or disappears after treatment
- How well vepdegestrant and PF-07220060 are absorbed by the body and how long they last in the body

Study Design



In part 1 of the study, people living with ER+/HER2- advanced or metastatic breast cancer for whom surgery is not recommended who also

• Were previously treated with 1 or more standard of care therapies for advanced or metastatic breast cancer

In part 2 of the study, people living with ER+/HER2- advanced or metastatic breast cancer who also

- Were previously treated with 1 or 2 endocrine therapies for advanced or metastatic breast cancer
- Received their most recent endocrine therapy for at least 6 months before their disease got worse
- Were previously treated with a CDK4/6 inhibitor

WHAT IS THE TREATMENT?

People will receive different doses of vepdegestrant plus PF-07220060

- Vedegestrant will be taken as pills by mouth once daily
- PF-07220060 will be taken as pills by mouth twice daily

WHAT WILL BE MEASURED IN THE STUDY?

- The side effects experienced by people taking vepdegestrant plus PF-07220060
 - This includes any symptoms felt by the participants in the study, signs observed in the participants by the investigators, or **abnormalities** that are detected in the participants' blood samples or by electrocardiogram
- Tumor size will be measured by scans to evaluate if vepdegestrant plus PF-07220060 treatment has any effect on slowing tumor growth or shrinking tumors
- The amount of time that people survive without their cancer getting worse
- Levels of vepdegestrant and PF-07220060 in the blood will be measured to assess how the different treatments may influence each other

Who sponsored the study?

This study is sponsored by Pfizer, Inc. in collaboration with Arvinas Estrogen Receptor, Inc.

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Arvinas and Pfizer thank the people who are volunteering to participate in this study and their caregivers, as well as the investigators, researchers, and coordinators who are contributing to this study

Writing and editorial support for this summary was provided by Marita Chakhtoura, MS, PhD, of Red Nucleus, and was funded by Arvinas Operations, Inc.

Where can I find more information?

For more information on this study

WHO CANNOT PARTICIPATE IN THE

• People who are **at risk**

of life-threatening complications in the short

People who have

cancer

inflammatory breast

term due to their cancer

People whose cancer has

spread to the brain and has not been treated

STUDY?

VIEW CLINICAL TRIAL RECORD

For more information on clinical studies in general

VIEW INFORMATION