



Chugai Obtains Approval for Additional Indication of Rozlytrek for *ROS1* Fusion-Positive Non-Small Cell Lung Cancer

- Rozlytrek approved for *ROS1* fusion-positive non-small cell lung cancer following MHLW approval of Rozlytrek for *NTRK* fusion-positive solid tumors last year
- *ROS1* fusion gene is found in 1 to 2% of patients with non-small cell lung cancer and is one of the important cancer-driver genes in the treatment of lung cancer

TOKYO, February 21, 2020 -- [Chugai Pharmaceutical Co., Ltd.](https://www.chugai-pharm.co.jp/english/news/detail/20191226113000_678.html) (TOKYO: 4519) announced today that it obtained approval for an additional indication of an anticancer agent/tyrosine kinase inhibitor, Rozlytrek® (generic name: entrectinib) for the treatment of *ROS1* fusion-positive, unresectable, advanced or metastatic non-small cell lung cancer (NSCLC) from the Ministry of Health, Labour and Welfare (MHLW).

"We are very pleased to announce the approval of Rozlytrek for the treatment in adults with *ROS1* fusion-positive NSCLC. Rozlytrek was launched last year as Chugai's first product to embody advanced personalized healthcare for the treatment of patients with *NTRK* fusion-positive solid tumors regardless of their age or site of origin," said Dr. Osamu Okuda, Executive Vice President, Co-Head of Project & Lifecycle Management Unit. "*ROS1* is an important cancer-driver gene found in 1 to 2% of patients with NSCLC. Offering Rozlytrek as a new treatment option for *ROS1* fusion-positive NSCLC patients, we will continue making efforts to contribute to the progress in advanced personalized medicine."

This approval is based mainly on the results of an open-label, multicenter, global phase II STARTRK-2 study. Efficacy was evaluated in *ROS1* fusion-positive NSCLC cohort from the STARTRK-2 study while safety was evaluated in two overseas phase I studies (STARTRK-1 study and ALKA-372-001 study) in addition to STARTRK-2 study.

FoundationOne CDx® Cancer Genomic Profile is used as a companion diagnostic to identify people that could potentially benefit from Rozlytrek for the *ROS1* fusion-positive, unresectable, advanced or metastatic NSCLC. The MHLW granted the approval for the expanded use of FoundationOne CDx Cancer Genomic Profile as a companion diagnostic of Rozlytrek on December 25, 2019.

As a leading company in the field of oncology, Chugai is committed to contribute to patients and medical professionals through realization of advanced personalized healthcare.

[Reference information]

Chugai Obtains Approval for Expanded Use of FoundationOne CDx Cancer Genomic Profile as a Companion Diagnostic of Rozlytrek for *ROS1*-positive Lung Cancer (A press release issued by Chugai on December 26, 2019)

https://www.chugai-pharm.co.jp/english/news/detail/20191226113000_678.html

Roche's investigational medicine entrectinib showed a durable response of more than two years in people with a specific type of lung cancer (A press release issued by Roche on September 24, 2018)

<https://www.roche.com/media/releases/med-cor-2018-09-24c.htm>

Approval information The underlined part has been newly added.

Product name	Rozlytrek® Capsules 100 mg Rozlytrek® Capsules 200 mg
Generic name	entrectinib
Indications	<i>NTRK</i> fusion-positive advanced or metastatic solid tumors <u><i>ROS1</i> fusion gene positive, unresectable, advanced or metastatic non-small cell lung cancer</u>
Dosage and administration	<ul style="list-style-type: none"> • <u>In case of patients with <i>NTRK</i> fusion-positive advanced or metastatic solid tumors.</u> The usual adult dosage is 600 mg entrectinib administered orally once a day. Reduce the dose as necessary depending on the patient's condition. The usual pediatric dosage is 300 mg/m² (body surface area) entrectinib administered orally once a day. However, the dose should not exceed 600 mg. Reduce the dose as necessary depending on the patient's condition. • <u>In case of patients with <i>ROS1</i> fusion gene positive, unresectable, advanced or metastatic non-small cell lung cancer.</u> <u>The usual adult dosage is 600 mg entrectinib administered orally once a day. Reduce the dose as necessary depending on the patient's condition.</u>
Conditions for approval	<ol style="list-style-type: none"> 1. A risk management plan should be created and appropriately implemented. 2. Given that the number of patients in clinical studies in Japan was extremely limited, post-marketing drug use surveillance of all patients receiving Rozlytrek should be conducted until data for a certain number of patients have been accumulated, in order to understand background information on patients receiving Rozlytrek, collect early data on the safety and efficacy of Rozlytrek, and take necessary measures for appropriate use of Rozlytrek.

About Rozlytrek

Rozlytrek is an oral tyrosine kinase inhibitor that blocks *ROS1* (c-ros oncogene 1) and TRK (neurotrophin receptors) family strongly and selectively. It blocks *ROS1* and TRK kinase activity, and inhibits proliferation of cancer cells with *ROS1* or *NTRK* gene fusions. Rozlytrek was approved for the treatment of locally advanced or metastatic solid tumors that harbor *NTRK1/2/3* gene fusions on June 18, 2019 and was launched on September 4, 2019.

About *ROS1* fusion-positive NSCLC

ROS1 fusion gene is an abnormal gene that can be formed by fusing the *ROS1* gene and other genes (*CD74*, etc.) as a result of chromosomal translocation for some reason. The *ROS1* fusion kinase made from *ROS1* fusion gene is thought to promote cancer cell proliferation. *ROS1* fusion gene is found in about 1 to 2% of NSCLC, among which it is more expressed in adenocarcinoma.¹⁾

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[Reference]

1: Bergethon K, Shaw AT, Ou SH, et al. *ROS1* rearrangements define a unique molecular class of lung cancers. *J Clin Oncol* 2012;30:863-70.

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