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Datroway approved in the EU as only TROP2-directed medicine with overall survival benefit for the 1st-line treatment of patients with metastatic TNBC who are not candidates for immunotherapy

Approval based on TROPION-Breast02 Phase III trial results where AstraZeneca and Daiichi Sankyo's Datroway showed a statistically significant and clinically meaningful improvement for the dual primary endpoints of overall survival and progression-free survival

Datroway now approved for two breast cancer indications in the EU

AstraZeneca and Daiichi Sankyo's *Datroway* (datopotamab deruxtecan) has been approved in the European Union (EU) as monotherapy for the 1st-line treatment of adult patients with unresectable or metastatic triple-negative breast cancer (TNBC) who are not candidates for PD-1/PD-L1 inhibitor therapy.

The approval by the European Commission follows the positive opinion of the Committee for Medicinal Products for Human Use of the European Medicines Agency and is based on results from the TROPION-Breast02 Phase III trial which were presented at the 2025 European Society for Medical Oncology Congress and subsequently published in *Annals of Oncology*.

Giuseppe Curigliano, MD, PhD, Director of the Early Drug Development Division, European Institute of Oncology, Professor of Medical Oncology, University of Milan, Italy and investigator for the TROPION-Breast02 trial, said: "For people living with metastatic triple-negative breast cancer, every new treatment option matters. Despite recent advances, more than two thirds of patients are not candidates for immunotherapy and have had limited options beyond chemotherapy. In my practice, I see firsthand the devastating impact this aggressive disease has on patients and their families. This approval of datopotamab deruxtecan provides a new treatment option for eligible patients and represents meaningful progress."

Dave Fredrickson, Executive Vice President, Oncology Haematology Business Unit, AstraZeneca, said: "Every year, more than 80,000 people in Europe are diagnosed with triple-negative breast cancer, a disease that often affects younger women and has limited treatment options in the metastatic setting. Today's approval of *Datroway* brings an antibody drug conjugate with a differentiated clinical profile underpinned by a strong survival benefit to people with this aggressive disease."

Ken Keller, Global Head of Oncology Business, and President and CEO, Daiichi Sankyo, Inc, said: "With this approval, *Datroway* is the only TROP2-directed antibody drug conjugate approved in the EU that has demonstrated an overall survival benefit in the 1st-line setting for the treatment of patients with metastatic triple-negative breast cancer. We look forward to bringing *Datroway* to patients in the EU as an additional treatment option with the potential to extend survival, reflecting our commitment to advancing innovative medicines that address unmet needs for people living with cancer."

In the trial, which included patients with metastatic TNBC who experienced early relapse following prior treatment, *Datroway* demonstrated a statistically significant and clinically meaningful 5.0-month improvement in median overall survival (OS) (hazard ratio [HR] 0.79; 95% confidence interval [CI] 0.64-0.98; p=0.0291) compared to chemotherapy as 1st-line treatment in this patient population. Median OS was 23.7 months for patients treated with *Datroway* versus 18.7 months for those treated with chemotherapy. *Datroway* reduced the risk of disease progression or death by 43% compared to chemotherapy (HR 0.57; 95% CI 0.47-0.69; p<0.0001) as assessed by blinded independent central review (BICR). *Datroway* was also associated with more robust treatment responses, including an objective response rate (ORR) of 62.5% compared to an ORR of 29.3% with chemotherapy.[1]

The safety profile of *Datroway* in TROPION-Breast02 was consistent with previous clinical trials of *Datroway* in breast cancer.

Based on the results of TROPION-Breast02, *Datroway* has been included in the ESMO Clinical Practice Guidelines as a Category IA 1st-line treatment option for patients with metastatic TNBC who are not candidates for immunotherapy, and it is the preferred option for patients who have relapsed within six months of completing adjuvant therapy.[2] In addition, *Datroway* received a score of 4 out of 5 on the ESMO Magnitude of Clinical Benefit Scale (ESMO-MCBS), recognising the clinically meaningful benefit demonstrated in TROPION-Breast02.[3]

Datroway was approved in the US in May 2026 for the same indication. Additional reviews are underway in China and Japan, as well as Australia, Canada, Singapore and Switzerland as part of Project Orbis.

Datroway is a specifically engineered TROP2-directed DXd antibody drug conjugate discovered by Daiichi Sankyo and being jointly developed and commercialised by AstraZeneca and Daiichi Sankyo.

Notes

Triple-negative breast cancer
TNBC accounts for approximately 15% of all breast cancer cases, with an estimated 365,000 diagnoses globally each year.[4,5] In Europe, there are an estimated 81,000 diagnoses of TNBC each year.[4,6] TNBC is diagnosed more frequently in younger and premenopausal women, and is more prevalent in Black and Hispanic women.[7-9] Metastatic TNBC is the most aggressive type of breast cancer and has one of the worst prognoses, with median OS of just 12 to 18 months and only about 15% of patients living five years following diagnosis.[7,10,11]

While some breast cancers may test positive for oestrogen receptors, progesterone receptors or overexpression of HER2, TNBC tests negative for all three.[7] Due to its aggressive nature and absence of common breast cancer receptors, TNBC is characteristically difficult to treat.[7] For patients with metastatic disease with PD-L1 expressing tumours, the addition of immunotherapy to chemotherapy has improved outcomes in the 1st-line setting.[12,13] However, for approximately 70% of patients with metastatic TNBC who are not candidates for immunotherapy, chemotherapy was the standard 1st-line treatment.[14]

TROP2 is a protein broadly expressed in several solid tumours, including TNBC.[15] TROP2 is associated with increased tumour progression and poor survival in patients with breast cancer.[16,17]

TROPION-Breast02

TROPION-Breast02 is a global, multicentre, randomised, open-label Phase III trial evaluating the efficacy and safety of *Datroway* versus investigator's choice of chemotherapy (paclitaxel, nab-paclitaxel, capecitabine, carboplatin or eribulin) in patients with previously untreated locally recurrent inoperable or metastatic TNBC for whom immunotherapy was not an option. This included patients whose tumours did not express PD-L1 as well as patients with PD-L1 expressing tumours who could not receive immunotherapy due to prior exposure in early-stage disease, comorbidities or immunotherapy not being accessible in their geography. Enrolment included patients with de novo or recurrent disease, regardless of disease-free interval, and those with poor prognostic factors such as stable brain metastases.

The dual primary endpoints of TROPION-Breast02 are OS and progression-free survival (PFS) as assessed by blinded independent central review. Secondary endpoints include PFS as assessed by investigator, ORR, duration of response, disease control rate, pharmacokinetics and safety.

TROPION-Breast02 enrolled 644 patients at sites in Africa, Asia, Europe, North America and South America. For more information, visit ClinicalTrials.gov.

Datroway
Datroway (datopotamab deruxtecan; datopotamab deruxtecan-dlnk in the US only) is a TROP2-directed ADC. Designed using Daiichi Sankyo's proprietary DXd ADC Technology, *Datroway* is one of seven DXd ADCs in the oncology pipeline of Daiichi Sankyo, and one of the most advanced programmes in AstraZeneca's ADC scientific platform. *Datroway* is comprised of a humanised anti-TROP2 IgG1 monoclonal antibody, developed in collaboration with Sapporo Medical University, attached to a number of topoisomerase I inhibitor payloads (an exatecan derivative, DXd) via tetrapeptide-based cleavable linkers.

Datroway is approved in more than 45 countries/regions worldwide for the treatment of adult patients with unresectable or metastatic HR-positive, HER2-negative (IHC 0, IHC 1+ or IHC 2+/ISH-) breast cancer who have received prior endocrine-based therapy and chemotherapy for unresectable or metastatic disease based on results from the TROPION-Breast01 trial.

Datroway is approved in more than 30 countries/regions worldwide for the treatment of adult patients with unresectable or metastatic TNBC who are not candidates for PD-1/PD-L1 inhibitor therapy based on the results from the TROPION-Breast02 trial.

Datroway is available in the US under accelerated approval for the treatment of adult patients with locally advanced or metastatic *EGFR*-mutated non-small cell lung cancer (NSCLC) who have received prior *EGFR*-directed therapy and platinum-based chemotherapy based on results from the TROPION-Lung05 and TROPION-Lung01 trials. Continued approval for this indication in the US may be contingent upon verification and description of clinical benefit in a confirmatory trial.

***Datroway* clinical development programme**
A comprehensive global clinical development programme is underway with more than 20 trials evaluating the efficacy and safety of *Datroway* across multiple cancers, including NSCLC, TNBC and urothelial cancer. The programme includes eight Phase III trials in lung cancer, five Phase III trials in breast cancer, and one Phase II/III trial in urothelial cancer evaluating *Datroway* as a monotherapy and in combination with other cancer treatments in various settings.

Daiichi Sankyo collaboration
AstraZeneca and Daiichi Sankyo entered into a global collaboration to jointly develop and commercialise *Enhertu* (trastuzumab deruxtecan) in March 2019 and *Datroway* in July 2020, except in Japan where Daiichi Sankyo maintains exclusive rights for each ADC. Daiichi Sankyo is responsible for the manufacturing and supply of *Enhertu* and *Datroway*.

AstraZeneca in breast cancer
Driven by a growing understanding of breast cancer biology, AstraZeneca is challenging, and redefining, the current clinical paradigm for how breast cancer is classified and treated to deliver even more effective treatments to patients in need - with the bold ambition to one day eliminate breast cancer as a cause of death.

AstraZeneca has a comprehensive portfolio of approved and promising compounds in development that leverage different mechanisms of action to address the biologically diverse breast cancer tumour environment.

With *Enhertu*, AstraZeneca and Daiichi Sankyo are aiming to improve outcomes in previously treated HER2-positive, HER2-low and HER2-ultralow metastatic breast cancer, and expanding its potential in earlier lines of treatment and in new breast cancer settings.

In HR-positive breast cancer, AstraZeneca continues to improve outcomes with foundational medicines *Faslodex* (fulvestrant) and *Zoladex* (goserelin) and aims to reshape the HR-positive space with first-in-class AKT inhibitor, *Truqap* (capivasertib), the TROP2-directed ADC, *Datroway*, and next-generation oral SERD, *Etcamah*.

PARP inhibitor *Lynparza* (olaparib) is a targeted treatment option that has been studied in early and metastatic breast cancer patients with an inherited *BRCA* mutation. AstraZeneca with MSD (Merck & Co., Inc. in the US and Canada) continue to research *Lynparza* in these settings. AstraZeneca is also exploring the potential of saruparib, a potent and selective inhibitor of PARP1, in combination with camizestrant in *BRCA*-mutated, HR-positive, HER2-negative advanced breast cancer.

To bring much-needed treatment options to patients with triple-negative breast cancer, an aggressive form of breast cancer, AstraZeneca is collaborating with Daiichi Sankyo to evaluate the potential of *Datroway* alone and in combination with immunotherapy *Imfinzi* (durvalumab).

AstraZeneca in oncology
AstraZeneca is leading a revolution in oncology with the ambition to provide cures for cancer in every form, following the science to understand cancer and all its complexities to discover, develop and deliver life-changing medicines to patients.

The Company's focus is on some of the most challenging cancers. It is through persistent innovation that AstraZeneca has built one of the most diverse portfolios and pipelines in the industry, with the potential to catalyse changes in the practice of medicine and transform the patient experience.

AstraZeneca has the vision to redefine cancer care and, one day, eliminate cancer as a cause of death.

AstraZeneca
AstraZeneca (LSE/STO/NYSE: AZN) is a global, science-led biopharmaceutical company that focuses on the discovery, development, and commercialisation of prescription medicines in Oncology, Rare Disease, and BioPharmaceuticals, including Cardiovascular, Renal & Metabolism, and Respiratory & Immunology. Based in Cambridge, UK, AstraZeneca's innovative medicines are sold in more than 125 countries and used by millions of patients worldwide. Please visit astrazeneca.com and follow the Company on Social Media @AstraZeneca.

Contacts
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