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Amylyx Pharmaceuticals Receives CHMP Negative Opinion on its Conditional Marketing Authorisation Application for AMX0035 for the Treatment of ALS in the European Union

Amylyx will seek re-examination of its Conditional Marketing Authorisation Application

Amylyx announced that the Committee for Medicinal Products for Human Use (CHMP) of the European Medicines Agency (EMA) has adopted a negative opinion on the application for conditional marketing authorisation of AMX0035 (sodium phenylbutyrate and ursodoxicoltaurine [also known as taurursodiol]), under the trade name ALBRIOZA®, for the treatment of adults with amyotrophic lateral sclerosis (ALS) in the European Union (EU).

The update follows the Company's May 2023 announcement that the CHMP was trending toward a negative opinion.

"We are confident in the strength of our CENTAUR trial data, which we believe meets the criteria for conditional approval. These data were also the basis of the full approval received from the U.S. Food and Drug Administration and the approval with conditions from Health Canada," said Tammy Sarnelli, Global Head, Regulatory Affairs and Clinical Compliance at Amylyx.

"We disagree with the CHMP's opinion and will request a formal re-examination procedure of the current Marketing Authorisation Application (MAA)."

CENTAUR met its prespecified primary outcome, and AMX0035 is the first ALS therapy to demonstrate, in the same trial, both a statistically significant benefit in function, as well as an observed benefit on overall survival in a longer-term post hoc analysis. AMX0035 demonstrated a generally well-tolerated safety profile in the CENTAUR trial, with similar reported rates of adverse events and discontinuations in AMX0035 and placebo groups during the 24-week randomized phase; however, gastrointestinal events occurred with greater frequency (≥2%) in the AMX0035 group.

The CENTAUR data were published in peer-reviewed medical journals, including the *New England Journal of Medicine*, *Muscle & Nerve*, and the *Journal of Neurology, Neurosurgery, and Psychiatry*.
"We will continue to engage with CHMP and EMA through the re-examination process with the goal of making ALBRIOZA available in Europe as it is in the United States and Canada. We know how precious time is for people with ALS and their families," said Joshua Cohen and Justin Klee, Co-CEOs of Amylyx.

The re-examination procedure is an approximately four-month process, which includes the appointment of a different rapporteur and co-rapporteur from the initial evaluation. At the end of the re-examination, the CHMP will adopt a final opinion.

"While our MAA continues to be under review, we will also work towards completing our global Phase 3 PHOENIX study, with topline results anticipated in mid-2024, which will provide important additional data on the efficacy and safety profile of ALBRIOZA," said Stéphanie Hoffmann-Gendebien, General Manager and Head of EMEA at Amylyx. "We remain committed to exploring all potential paths forward. There have been no new innovations approved in Europe for this devastating disease in over 25 years, and we recognize the urgent need of the ALS community in Europe to access new treatment options."
ALS affects approximately 29,000 people in the U.S. and more than 30,000 people are estimated to be living with ALS in Europe (EU and United Kingdom).

About RELYVRIO®/ ALBRIOZA™ / ALBRIOZA® / AMX0035

RELYVRIO®, an oral, fixed-dose combination of sodium phenylbutyrate and taurursodiol (known as ursodoxicoltaurine outside of the U.S.), is approved to treat amyotrophic lateral sclerosis (ALS) in adults in the U.S. and approved with conditions as ALBRIOZA™ for the treatment of ALS in Canada. The Committee for Medicinal Products for Human Use (CHMP) of the European Medicines Agency (EMA) has adopted a negative opinion on the application for conditional marketing authorisation of AMX0035, under the trade name ALBRIOZA®, for the treatment of ALS in the European Union, and Amylyx will request a formal re-examination procedure of the current Marketing Authorisation Application. AMX0035 is being explored for the potential treatment of other neurodegenerative diseases. The formulation of RELYVRIO, ALBRIOZA, and AMX0035 are identical.

About the CENTAUR Trial

CENTAUR was a multicenter Phase 2 clinical trial in 137 participants with ALS encompassing a 6-month randomized placebo-controlled phase and an open-label extension (OLE) long-term follow-up phase. The trial met its primary efficacy endpoint.
Detailed safety and functional efficacy data from CENTAUR were published in the *New England Journal of Medicine*. Data from additional analyses from the CENTAUR trial were published in *Muscle & Nerve* in 2020 and 2022, and the *Journal of Neurology, Neurosurgery, and Psychiatry* in 2022.

Source: **Amylyx.com**