NEWS RELEASE

Horizon Therapeutics plc Announces New UPLIZNA® (inebilizumab-cdon) Data in Neuromyelitis Optica Spectrum Disorder (NMOSD) to be presented at ECTRIMS2023

10/2/2023

-- Presentations will feature data from the pivotal Phase 3 study of UPLIZNA, including new biomarker analyses --

DUBLIN--(BUSINESS WIRE)-- Horizon Therapeutics plc (Nasdaq: HZNP) today announced that new UPLIZNA analyses will be presented at the 39th Congress of the European Committee for Treatment and Research in Multiple Sclerosis (ECTRIMS) 2023, Oct. 11-13. UPLIZNA is the first and only targeted CD19+ B-cell-depleting therapy approved by the U.S. Food and Drug Administration, the European Commission and the Brazilian Health Regulatory Agency (ANVISA) for the treatment of NMOSD in adults who are anti-aquaporin-4 immunoglobulin G seropositive (AQP4-IgG+).

Presentation Details:

- **P015:** Association of Cytokine Proteins with Disease Activity in NMOSD Participants Receiving Inebilizumab Treatment (S. Pittock)
    - Poster Session: Clinical aspects of MS - NMOSD
    - Date: Oct. 11, 2023, 8-8:30 a.m. CEST
- **P409:** Long-Term Comparative Efficacy of Inebilizumab in the AQP4+ Subpopulation from the N-MOmentum Open-Label Extension Versus Azathioprine and Immunosuppressive Therapies and Versus Placebo in Patients with NMOSD (B. Cree)
    - Poster Session: Clinical aspects of MS - NMOSD

    - Date: Oct. 11, 2023, 8-8:30 a.m. CEST

- **P011:** Matching-Adjusted Indirect Comparison of Current Treatments for NMOSD and Evaluation of Long-Term Effectiveness (F. Paul)
    - Poster Session: Clinical aspects of MS - NMOSD
    - Date: Oct. 11, 2023, 8-8:30 a.m. CEST

Horizon will host a symposium Thursday, Oct. 12 from 8:45 to 9:45 a.m. CEST, "Looking for unrecognized disease activity in NMOSD to optimize treatment choice and prevent disability," chaired by Massimo Filippi, M.D., Ph.D., featuring presentations from Maria Rocca, M.D., Orhan Aktas, M.D. and Jeffrey Bennett, M.D., Ph.D.

About Neuromyelitis Optica Spectrum Disorder (NMOSD)

NMOSD is a unifying term for neuromyelitis optica (NMO) and related syndromes. NMOSD is a rare, severe, relapsing, neuroinflammatory autoimmune disease that attacks the optic nerve, spinal cord, brain and brain stem.1,2 Approximately 80% of all patients with NMOSD test positive for anti-AQP4 antibodies.3 AQP4-IgG binds primarily to astrocytes in the central nervous system and triggers an escalating immune response that results in lesion formation and astrocyte death.4

Anti-AQP4 autoantibodies are produced by plasmablasts and some plasma cells. These B-cell populations are central to NMOSD disease pathogenesis, and a large proportion of these cells express CD19.5 Depletion of these CD19+ B-cells is thought to remove an important contributor to inflammation, lesion formation and astrocyte damage. Clinically, this damage presents as an NMOSD attack, which can involve the optic nerve, spinal cord and brain.4,6 Loss of vision, paralysis, loss of sensation, bladder and bowel dysfunction, nerve pain and respiratory failure can all be manifestations of the disease.7 Each NMOSD attack can lead to further cumulative damage and disability.8,9 NMOSD occurs more commonly in women and may be more common in individuals of African and Asian descent.10,11

About UPLIZNA

INDICATION

UPLIZNA is indicated for the treatment of neuromyelitis optica spectrum disorder (NMOSD) in adult patients who are anti-aquaporin-4 (AQP4) antibody positive.

IMPORTANT SAFETY INFORMATION

UPLIZNA is contraindicated in patients with:

- A history of life-threatening infusion reaction to UPLIZNA
- Active hepatitis B infection
- Active or untreated latent tuberculosis

WARNINGS AND PRECAUTIONS

Infusion Reactions: UPLIZNA can cause infusion reactions, which can include headache, nausea, somnolence, dyspnea, fever, myalgia, rash or other symptoms. Infusion reactions were most common with the first infusion but were also observed during subsequent infusions. Administer pre-medication with a corticosteroid, an antihistamine and an anti-pyretic.

Infections: The most common infections reported by UPLIZNA-treated patients in the randomized and open-label periods included urinary tract infection (20%), nasopharyngitis (13%), upper respiratory tract infection (8%) and influenza (7%). Delay UPLIZNA administration in patients with an active infection until the infection is resolved.

Increased immunosuppressive effects are possible if combining UPLIZNA with another immunosuppressive therapy.

The risk of Hepatitis B Virus (HBV) reactivation has been observed with other B-cell-depleting antibodies. Perform HBV screening in all patients before initiation of treatment with UPLIZNA. Do not administer to patients with active hepatitis.

Although no confirmed cases of Progressive Multifocal Leukoencephalopathy (PML) were identified in UPLIZNA clinical trials, JC virus infection resulting in PML has been observed in patients treated with other B-cell-depleting antibodies and other therapies that affect immune competence. At the first sign or symptom suggestive of PML, withhold UPLIZNA and perform an appropriate diagnostic evaluation. Patients should be evaluated for tuberculosis risk factors and tested for latent infection prior to initiating UPLIZNA.

Vaccination with live-attenuated or live vaccines is not recommended during treatment and after discontinuation, until B-cell repletion.

Reduction in Immunoglobulins: There may be a progressive and prolonged hypogammaglobulinemia or decline in the levels of total and individual immunoglobulins such as immunoglobulins G and M (IgG and IgM) with continued UPLIZNA treatment. Monitor the level of immunoglobulins at the beginning, during, and after discontinuation of treatment with UPLIZNA until B-cell repletion especially in patients with opportunistic or recurrent infections.

Fetal Risk: May cause fetal harm based on animal data. Advise females of reproductive potential of the potential risk to a fetus and to use an effective method of contraception during treatment and for 6 months after stopping UPLIZNA.

Adverse Reactions: The most common adverse reactions (at least 10% of patients treated with UPLIZNA and greater than placebo) were urinary tract infection and arthralgia.

For additional information on UPLIZNA, please see the Full Prescribing Information at **www.UPLIZNA.com**.

About Horizon

Horizon is a global biotechnology company focused on the discovery, development and commercialization of medicines that address critical needs for people impacted by rare, autoimmune and severe inflammatory diseases. Our pipeline is purposeful: We apply scientific expertise and courage to bring clinically meaningful therapies to patients. We believe science and compassion must work together to transform lives. For more information on how we go to incredible lengths to impact lives, visit **www.horizontherapeutics.com** and follow us on **Twitter**, **LinkedIn**, **Instagram** and **Facebook**.

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