

Astellas to Present Findings from Phase 3 Long-Term Safety Study of Fezolinetant in Oral Session at The North American Menopause Society 2022 Annual Meeting

SKYLIGHT 4™ study results demonstrate the 52-week safety and tolerability of fezolinetant 30 mg and 45 mg once daily

Pooled analyses will also be presented from SKYLIGHT 1™ and SKYLIGHT 2™ studies of fezolinetant, Astellas' investigational nonhormonal treatment for vasomotor symptoms (VMS) associated with menopause

TOKYO, October 12, 2022 – Astellas Pharma Inc. (TSE: 4503, President and CEO: Kenji Yasukawa, Ph.D., “Astellas”) will present 52-week results from the Phase 3 SKYLIGHT 4™ clinical study evaluating the safety and tolerability of fezolinetant, an investigational oral, nonhormonal compound being studied for the treatment of moderate to severe vasomotor symptoms (VMS) associated with menopause. The results will be featured in an oral presentation October 13 at The North American Menopause Society Annual Meeting. VMS, characterized by hot flashes and/or night sweats, are common symptoms of menopause.^{1,2}

SKYLIGHT 4 results demonstrate the 52-week safety and tolerability of fezolinetant 30 mg and 45 mg once daily. Safety analyses demonstrated that both endometrial hyperplasia and endometrial malignancy were within pre-specified limits for fezolinetant-treated patients; reported treatment-emergent adverse events (TEAEs) were generally mild or moderate in severity; and headache and COVID-19 were the most common TEAEs, with similar incidences for fezolinetant and placebo. The frequency of elevated liver enzymes was low across groups, and elevations were generally asymptomatic, isolated, transient and resolved on treatment or soon after study drug discontinuation.

“There is an unmet need for safe and effective nonhormonal treatment options for VMS associated with menopause, which can adversely impact daily quality of life,” said Genevieve Neal-Perry, M.D., Ph.D., Chair, UNC School of Medicine Department of Obstetrics and Gynecology. “The SKYLIGHT 4 study results demonstrate the long-term safety and tolerability of fezolinetant, providing further support for its potential use as a treatment for VMS.”

In addition, pooled efficacy data from SKYLIGHT 1™ and SKYLIGHT 2™ evaluating early response to fezolinetant, its impact on sleep and treatment response analyzed by race will be presented.

“Results of the SKYLIGHT 4 study and the pooled analyses from the SKYLIGHT 1 and 2 studies provide further insights into the safety and effectiveness of fezolinetant,” said Ahsan Arozullah, M.D., M.P.H., Senior Vice President and Head of Development Therapeutic Areas, Astellas. “Through our clinical development program, we are excited to further

characterize the clinical profile of fezolinetant for women who experience hot flashes as part of menopause."

Fezolinetant is an investigational selective neurokinin 3 (NK3) receptor antagonist and is not approved for use anywhere in the world. New Drug and Marketing Authorization applications for fezolinetant are under review in the U.S. and Europe, respectively. If approved, fezolinetant would be a first-in-class, nonhormonal treatment option to reduce the frequency and severity of moderate to severe VMS associated with menopause.

Astellas will also present three abstracts examining the association between VMS and weight gain, sleep quality and work productivity.

Following are all of Astellas' presentations at NAMS 2022:

SKYLIGHT Data at NAMS

Abstract	Presentation Details
A phase 3, randomized, placebo-controlled, double-blind study to investigate the long-term safety and tolerability of fezolinetant in women seeking treatment for vasomotor symptoms associated with menopause (SKYLIGHT 4) – Abstract S-11	Genevieve Neal-Perry, M.D., Ph.D. Oral Presentation October 13, 5:30-5:45 p.m. EDT
Early response with fezolinetant treatment of moderate-to-severe vasomotor symptoms associated with menopause: pooled data from two randomized Phase 3 studies – Abstract P-73	Marla Shapiro, M.D.C.M, M.H.Sc. Poster Session October 13, 6:15-7:15 p.m. EDT
Effect of fezolinetant treatment on patient-reported sleep disturbance: pooled data from two Phase 3 studies in women with moderate-to-severe vasomotor symptoms associated with menopause – Abstract P-74	Marla Shapiro, M.D.C.M, M.H.Sc. Poster Session October 13, 6:15-7:15 p.m. EDT
Fezolinetant for treatment of moderate-to-severe vasomotor symptoms associated with menopause: efficacy in women stratified by race using pooled data from two Phase 3 studies – Abstract P-53	Genevieve Neal-Perry, M.D., Ph.D. Poster Session October 13, 6:15-7:15 p.m. EDT

Astellas VMS Data at NAMS

Presentation Title	Presentation Details
Association between vasomotor symptom frequency and weight gain in women enrolled in the study of women's health across the nation (SWAN) – Abstract S-24	Carolyn Gibson, Ph.D., M.P.H. Oral Presentation October 14, 4:45-5 p.m. EDT
Association between vasomotor symptom severity and sleep outcomes in a survey of US women with symptoms of menopause – Abstract P-16	Barbara DePree, M.D. Poster Session October 13, 6:15-7:15 p.m. EDT
Association between severity of vasomotor symptoms of menopause and work productivity in a survey of US women – Abstract P-15	Barbara DePree, M.D. Poster Session October 13, 6:15-7:15 p.m. EDT

About the BRIGHT SKY™ Phase 3 Program

The BRIGHT SKY pivotal trials, SKYLIGHT 1™ (NCT04003155) and SKYLIGHT 2™ (NCT04003142), enrolled over 1,000 women with moderate to severe VMS. The trials are double-blinded, placebo-controlled for the first 12 weeks followed by a 40-week treatment extension period. Women were enrolled at over 180 sites within the U.S., Canada and Europe. SKYLIGHT 4™ (NCT04003389) is a 52-week double-blinded, placebo-controlled study designed to investigate the long-term safety of fezolinetant. For SKYLIGHT 4, over 1,800 women with VMS were enrolled at over 180 sites within the U.S., Canada and Europe.

About VMS Associated with Menopause

VMS, characterized by hot flashes (also called hot flushes) and/or night sweats, are common symptoms of menopause.^{1,2} In the U.S., about 60% to 80% of women experience these symptoms during or after the menopausal transition and, worldwide, more than half of women 40 to 64 years of age experience VMS.^{3,4,5,6} VMS can have a disruptive impact on women's daily activities and overall quality of life.¹

About Fezolinetant

Fezolinetant is an investigational oral, nonhormonal therapy in clinical development for the treatment of moderate to severe VMS associated with menopause. Fezolinetant works by blocking neurokinin B (NKB) binding on the kisspeptin/neurokinin/dynorphin (KNDy) neuron to moderate neuronal activity in the thermoregulatory center of the brain (the hypothalamus) to reduce the frequency and severity of moderate to severe VMS associated with menopause.^{7,8,9} The safety and efficacy of fezolinetant are under investigation and have not been established. There is no guarantee the agent will receive regulatory approval or become commercially available for the uses being investigated.

About Astellas

Astellas Pharma Inc. is a pharmaceutical company conducting business in more than 70 countries around the world. We are promoting the Focus Area Approach that is designed to identify opportunities for the continuous creation of new drugs to address diseases with high unmet medical needs by focusing on Biology and Modality. Furthermore, we are also looking beyond our foundational Rx focus to create Rx+[®] healthcare solutions that combine our expertise and knowledge with cutting-edge technology in different fields of external partners. Through these efforts, Astellas stands on the forefront of healthcare change to turn innovative science into value for patients. For more information, please visit our website at <https://www.astellas.com/en>.

Cautionary Notes

In this press release, statements made with respect to current plans, estimates, strategies and beliefs and other statements that are not historical facts are forward-looking statements about the future performance of Astellas. These statements are based on management's current assumptions and beliefs in light of the information currently available to it and involve known and unknown risks and uncertainties. A number of factors could cause actual results to differ materially from those discussed in the forward-looking statements. Such factors include, but are not limited to: (i) changes in general economic conditions and in laws and regulations, relating to pharmaceutical markets, (ii) currency exchange rate fluctuations, (iii) delays in new product launches, (iv) the inability of Astellas to market existing and new products effectively, (v) the inability of Astellas to continue to effectively research and develop products accepted by customers in highly competitive markets, and (vi) infringements of Astellas' intellectual property rights by third parties. Information about pharmaceutical products (including products currently in development) which is included in this press release is not intended to constitute an advertisement or medical advice.

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