



Better Health, Brighter Future

Takeda Announces Positive Results from Two Pivotal Phase 3 Studies of Oveporexton (TAK-861) in Narcolepsy Type 1

- Both Phase 3 Studies Met all Primary and Secondary Endpoints Demonstrating Statistically Significant Improvements Across Symptoms at All Doses, Building Upon Phase 2b Results*
- Oveporexton was Generally Well-Tolerated in Phase 3 Safety Profile*
- Takeda is Rapidly Advancing Regulatory Submissions and Launch Preparedness with the Aim to Bring Oveporexton to People Living with Narcolepsy Type 1 as Quickly as Possible*
- These Results Mark a Major Advancement Toward Transforming the Standard of Care by Addressing the Underlying Cause of Narcolepsy Type 1*

OSAKA, Japan and CAMBRIDGE, Massachusetts, July 14, 2025 – Takeda ([TSE:4502/NYSE:TAK](#)) today announced that all primary and secondary endpoints were met in two Phase 3 randomized, double-blind, placebo-controlled studies of oveporexton (TAK-861), a potential first-in-class investigational oral orexin receptor 2 (OX2R)-selective agonist, in narcolepsy type 1 (NT1). NT1 is caused by the loss of orexin-producing neurons in the brain. Orexin agonists are designed to address this underlying orexin deficiency. For the first time, this mechanism of action has been validated in Phase 3 studies demonstrating significant improvement across a broad range of symptoms. These results reinforce the potential of oveporexton to transform the standard of care.

“We are thrilled to reach this pivotal milestone for the oveporexton program. Oveporexton is a testament to Takeda’s strength in discovering and developing a potential new class of medicines for difficult to treat diseases such as narcolepsy type 1,” said Christophe Weber, president and chief executive officer at Takeda. “Our leadership in orexin biology and building a multi-asset orexin franchise with transformative potential will position Takeda for long-term future growth.”

The FirstLight (TAK-861-3001) and RadiantLight (TAK-861-3002) studies were two large, global Phase 3 studies conducted in 19 countries. Both studies achieved statistically significant improvement compared to placebo with p-values of <0.001 for all primary and secondary endpoints across all doses at week 12. The primary and secondary endpoints measuring objective and patient reported improvements in wakefulness, excessive daytime sleepiness, cataplexy, ability to maintain attention, overall quality of life and daily life functions demonstrate statistically significant and clinically meaningful improvements achieving near normal ranges across the broad range of symptoms investigated.

Oveporexton was generally well-tolerated with a safety profile from the Phase 3 studies overall consistent with oveporexton studies to date including the Phase 2b study. No serious treatment-related adverse events were reported. The most common adverse events were insomnia, urinary urgency and frequency. More than 95 percent of the participants who completed the studies enrolled in the ongoing long-term extension (LTE) study.

“We are grateful to the patients who took part in these clinical studies and to their families, the investigators and clinical staff. The studies were accelerated at an unprecedented pace with the aim to

bring this potential treatment to people living with narcolepsy type 1 as quickly as possible,” said Andy Plump, M.D., Ph.D., president of R&D at Takeda. “The comprehensive assessments from our Phase 3 studies build on the transformative results we saw with our Phase 2b study with most participants reaching normative ranges and reporting clinically meaningful improvement across a broad range of symptoms at the end of the 12-week treatment period. The positive results also reinforce the continued momentum for our late-stage pipeline, which we believe will deliver value to the patients we serve around the world.”

Takeda intends to present the results at upcoming medical congresses and plans to submit a New Drug Application with the United States Food and Drug Administration and additional global regulatory authorities in fiscal year 2025.

Results from the Phase 3 studies have no significant impact on the full year consolidated forecast for the fiscal year ending March 31, 2026.

About Narcolepsy Type 1 (NT1) and Orexin Science

NT1 is a chronic, rare neurological disease that results in a range of debilitating symptoms including excessive daytime sleepiness (EDS), cataplexy, disrupted nighttime sleep, sleep paralysis and hallucinations upon falling asleep or waking. Additionally, individuals living with NT1 often report cognitive symptoms, including difficulty thinking clearly, remembering, concentrating and paying attention. NT1 is caused by loss of the orexin-producing neurons in the brain, which regulate wakefulness and sleep, and is also believed to be essential to other functions such as attention through activation of orexin receptors. Currently, the standard of care is limited to symptomatic therapies that may only partially address some of the symptoms people face.

About Oveporexton (TAK-861)

Oveporexton (TAK-861) is an investigational orexin receptor 2 (OX2R)-selective agonist, which selectively stimulates the OX2R to restore signaling and address the underlying orexin deficiency that causes narcolepsy type 1 (NT1). By activating OX2Rs, oveporexton is designed to promote wakefulness and reduce abnormal rapid eye movement (REM)-sleep like phenomena, including cataplexy, to address the broad spectrum of daytime and nighttime symptoms.

About the FirstLight and RadiantLight Phase 3 Studies

FirstLight (TAK-861-3001; [NCT06470828](#)) and RadiantLight (TAK-861-3002; [NCT06505031](#)) are global, multicenter, placebo-controlled studies to evaluate the efficacy, safety and tolerability of oveporexton compared to placebo in patients with narcolepsy type 1 (NT1) over 12 weeks. The studies were conducted in 19 countries with enrollment completed within six months. The FirstLight study enrolled 168 participants randomized to one of three dosing arms (high dose, low dose and placebo). The RadiantLight study enrolled 105 participants randomized to two dosing arms (high dose and placebo). The primary endpoint in both studies was improvement in excessive daytime sleepiness (EDS) as measured by the Maintenance of Wakefulness Test (MWT), a standard measure of wakefulness. Key secondary endpoints included improvement in EDS as measured by the Epworth Sleepiness Scale (ESS) and in the Weekly Cataplexy Rate (WCR), a measure evaluating cataplexy. The studies also evaluated the effect of oveporexton on participants' ability to maintain attention, participants' overall quality of life, the spectrum of narcolepsy symptoms and daily life functions, as well as the safety and tolerability of oveporexton.

About Takeda's Orexin Agonists for Sleep-Wake Disorders

Takeda is leading the field of orexin science with a multi-asset franchise. Orexin is a key regulator of sleep and wake patterns and contributes to other essential functions including attention, mood, metabolism and respiration. Oveporexton (TAK-861) is the lead investigational orexin receptor 2 (OX2R) agonist asset in Takeda's orexin franchise and received Breakthrough Therapy designation for the treatment of excessive daytime sleepiness in narcolepsy type 1 (NT1) from the U.S. Food and Drug Administration and the Center for Drug Evaluation of China's National Medical Products Administration. The company is also investigating other orexin agonists in populations with orexin levels in the normal range, including TAK-360, an oral OX2R agonist initially being investigated for the treatment of narcolepsy type 2 (NT2), idiopathic hypersomnia (IH), and other potential indications where orexin signaling is implicated.

About Takeda

Takeda is focused on creating better health for people and a brighter future for the world. We aim to discover and deliver life-transforming treatments in our core therapeutic and business areas, including gastrointestinal and inflammation, rare diseases, plasma-derived therapies, oncology, neuroscience and vaccines. Together with our partners, we aim to improve the patient experience and advance a new frontier of treatment options through our dynamic and diverse pipeline. As a leading values-based, R&D-driven biopharmaceutical company headquartered in Japan, we are guided by our commitment to patients, our people and the planet. Our employees in approximately 80 countries and regions are driven by our purpose and are grounded in the values that have defined us for more than two centuries. For more information, visit www.takeda.com.

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