

Lead Asset: Serplulimab

- Evaluated across different stages (**Extended Stage** and **Limited Stage**) of SCLC in **two phase 3 trials in the US**. Both trials are active and currently recruiting
- **SCLC** continues to be an **area of high unmet need with limited innovation**
- **Orphan drug designation by FDA and EC**
- Potential for lifecycle management through **additional indications** based on trials in China
- EMA validates Marketing Authorization application for first-line for treatment of ES-SCLC (Mar 2023)
- **~Early 2026 launch**

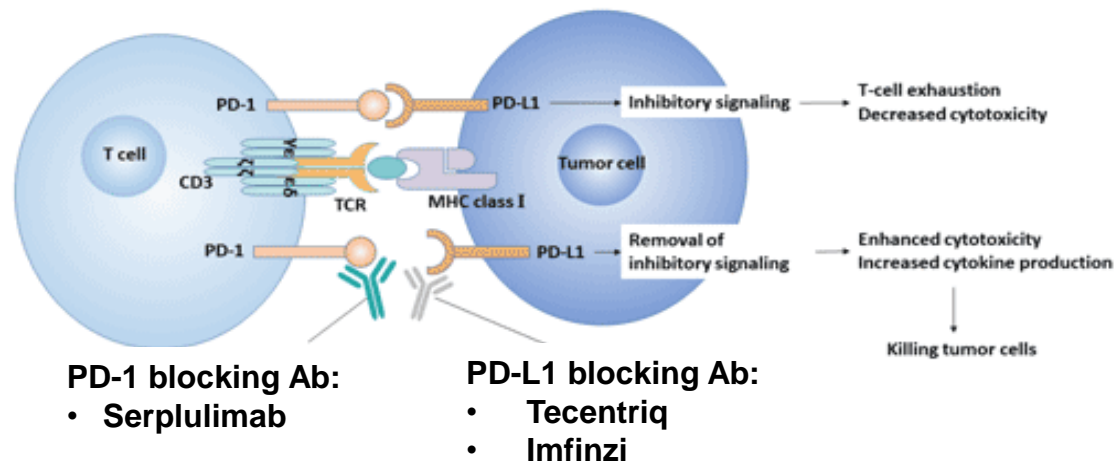
Serplulimab (Potential Best-in-Class 2nd Generation PD-1)

External Innovation / Licensing	Fosun Pharma Group / Henlius Pipeline Access
<p><i>Seek external assets that are:</i></p> <ul style="list-style-type: none"> • Synergistic with commercial capabilities • First/best in class oncology assets in suitable markets • PD-1 Combinable 	<p><i>Expand Henlius portfolio partnership:</i></p> <ul style="list-style-type: none"> • HLX22 (Ph. 3 ready global pivotal trial in HER2-positive gastric cancer) <ul style="list-style-type: none"> ○ Additional indications to be explored Henlius pipeline combination opportunities with Serplulimab

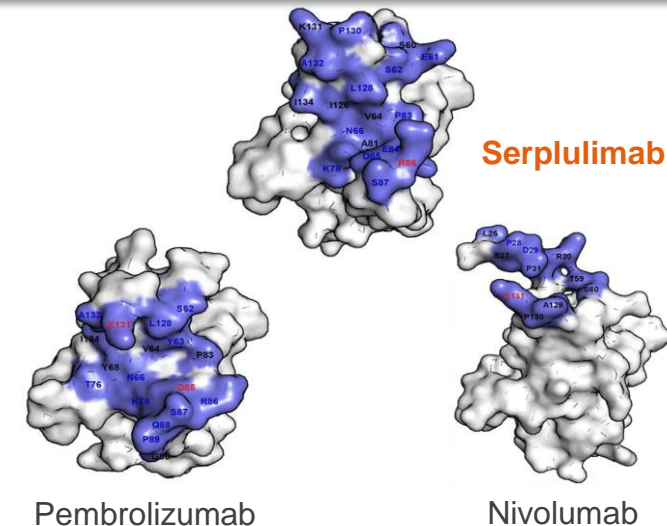
Serplulimab's Unique Mode of Recognition Leads to Differentiation

- Serplulimab distinguished itself by **binding to different epitopes on PD-1** in comparison to nivolumab (Opdivo) and pembrolizumab (Keytruda). Epitope analysis revealed that Serplulimab possessed a **unique mode of recognition** when contrasted with the clinically approved PD-1 antibodies (Ab) pembrolizumab and nivolumab, which had the capability to block both PD-L1 and PD-L2 binding.
 - In preclinical studies, Serplulimab demonstrated a **higher affinity for human PD-1** than nivolumab and pembrolizumab¹. It also exhibited **greater potency in blocking PD-L1 and PD-L2** signaling compared to nivolumab¹.
- **Tecentriq (atezolizumab) and Imfinzi (durvalumab) are PD-L1 inhibitors** that block PD-L1 on tumor cells. **Tumors are heterogeneous; some tumors with low or no PD-L1 expression do not respond to PD-L1 inhibitors.** Additionally, tumors expressing PD-L2 cannot be blocked by PD-L1 inhibitors, which still has the potential to inhibit T cell activation signaling.

PD-1 Immunoinhibitory signaling on T cells



Comparison of Binding Epitope Regions (Blue) of Serplulimab With Pembrolizumab and Nivolumab



Competitive Landscape for PD-(L)1 in ES-SCLC

Key Assumptions

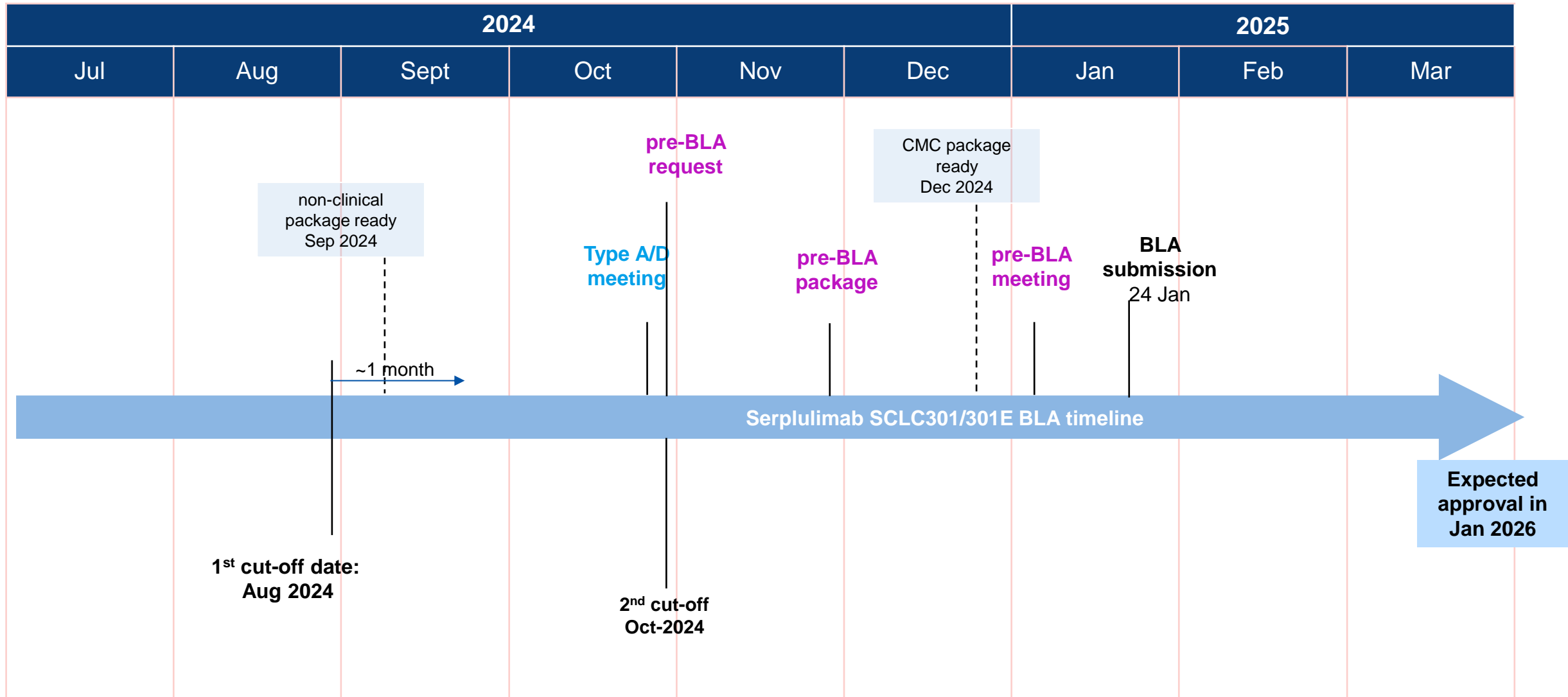
- Serplulimab could be 1st PD-1 and 3rd PD-(L)1 approved/ marketed in US in 1L ES-SCLC, following Tecentriq and Imfinzi
- Serplulimab expected to launch in US in early 2026

Competitive Pipeline – PD-(L)1s in Phase 3 for 1L ES-SCLC

Manufacturer	Product	MOA	Estimated Primary Completion Date	Trial Location	Bridging Study Requirement	Bridging Study in Place
 HENGRUI	SHR-1316	PD-L1	10/2022	China	✓	✗
 sorrento THERAPEUTICS	Socazolimab	PD-L1	5/2023	China	✓	✗
 BeiGene	Tislelizumab	PD-1	10/2023	China	✓	✗

Source(s): Evaluate Pharma SCLC Market Landscape; Clinicaltrials.gov; Pharmaprojects; GlobalData; Company websites; CRA analysis

Serplulimab ES- SCLC US BLA Timeline



Key Assumptions

- SCLC accounts for ~14% of all lung cancer cases¹
- Cigarette smoking is the major risk factor for lung cancer, accounting for more than 95% of SCLC cases²
- SCLC is a rapidly progressing tumor with a 5-year survival of only 8%³
- There are expected to be ~29,000 eligible patients with Extensive stage SCLC (ES-SCLC) at launch:
 - ~21,000 de-novo diagnosed patients
 - ~8,000 limited stage patients who progress to extensive stage

81% of 1L ES-SCLC are actively treated

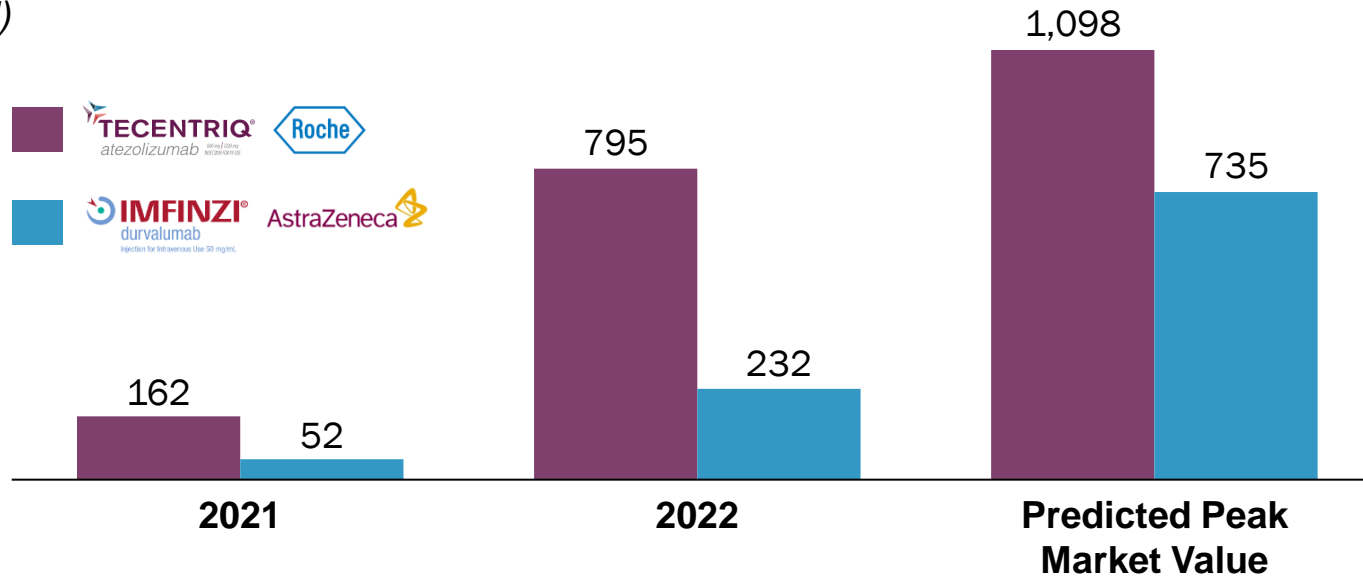
Driven by performance status and ability to tolerate chemo

90% of actively treated 1L ES-SCLC patients are eligible for IO therapy

Eligibility for immunotherapy is contingent upon the presence and severity of comorbidities, such as severe uncontrolled autoimmune disorders

1. Ganti AK, et al. JAMA Oncol. 2021;7(12):1824–1832. 2. Govindan R, et al. J Clin Oncol. 2006;24(28):4539-4544. 3. National Cancer Institute. SEER Incidence Data, November 2022 Submission (1975-2020). <https://seer.cancer.gov/statistics-network/explorer/>..SEER data used to understand patient staging. CRA Quant PMR (N=63 Oncs); CRA Qual PMR (N=16 Oncs); CRA analysis

US sales of Tecentriq® (atezolizumab) vs. Imfinzi® (durvalumab) in ES-SCLC Indication (\$M)



US SCLC Market



Key Drivers for SCLC Market Growth

- Biomarkers:** SCLC could become more similar to NSCLC with an increasing use of biomarkers driving market growth
- New Entrants:** Several SCLC PD-1/PD-L1 IOs are currently in the late-stage development pipeline and will soon crowd the market
- Duration of Therapy:** Extending duration of therapy from 3 to 4 cycles will increase use and therefore the size of the market
- Line of Therapy:** 1L treatment is vital in SCLC given late-stage diagnosis and low survival rates

*Atezo has an LOE date of 2032, and durva's LOE is in 2031; Global SCLC market size in 2028: ~\$4B

Sources: Citeline [article](#), Credit Suisse Research [Bulletin](#); Future Market Insights [\[link\]](#); Dawkins et al., Nature Reviews, 2020 [\[link\]](#); iHealthAnalyst [\[link\]](#) Evaluate