



Silence Therapeutics

*Collaboration with AstraZeneca
and R&D update*

March 25 2020

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Agenda



- Opening remarks
Iain Ross, Executive Chairman
- Collaboration with AstraZeneca
Iain Ross, Executive Chairman
- R&D Update
Dr Giles Campion, Head R&D, Chief Medical Officer
- Q&A
All participants

Iain Ross



Giles Campion



Rob Quinn



John Strafford



Collaboration with AstraZeneca for the Treatment of Cardiovascular, Renal, Metabolic and Respiratory diseases



- **Research, Collaboration, Option and Licence agreement to discover and develop RNA interference therapeutics for the treatment of serious diseases.**
- > Draws on Silence's extensive experience in the development of siRNA therapeutics, together with AstraZeneca's industry leading expertise in disease biology and target identification, to develop first-in-class and differentiated therapeutics to address significant unmet need.
- > Five targets anticipated to be started within three years with AstraZeneca having the option to extend the collaboration to a further five targets.
- > Silence's GalNAc-siRNA platform to be harnessed for liver expressed gene targets. Both parties to collaborate to achieve targeted delivery of siRNA molecules to other tissues including heart, kidney and lung.
- > Silence will be responsible for designing siRNA molecules against gene targets selected by AstraZeneca, and for manufacturing of material to support GLP toxicology studies and Phase I clinical studies.
- > Silence will have the option to negotiate for co-development of two programs of their choice starting from Phase II.

- > \$60 million cash upfront and \$20 million equity investment.
- > Total deal value over \$4 billion
- > Option fee of \$10 million per target at the point of candidate nomination. Up to \$140 million in development milestones and up to \$250 million in commercialisation milestones. Total milestones of up to \$400 million per target.
- > High single digit to low double-digit royalties on net sales for each target.

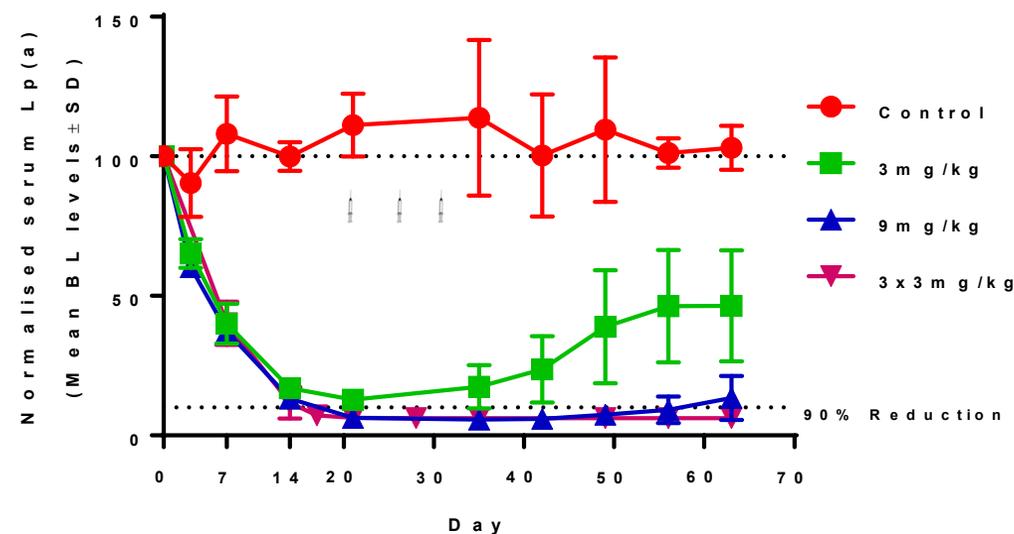


R&D Update



- Re-prioritisation of R&D pipeline due to strengthened balance sheet
- **SLN360** brought forward as priority asset
- LP(a) targeting siRNA for cardiovascular disease
- High unmet medical need and excellent preclinical profile
- IND to be filed in H2 2020, interim Phase I data mid 2021
- **SLN124** – granted Rare Pediatric Disease designation by the FDA for β Thalassemia
- Screening for patients stopped as precautionary measure
- New protocol with broader patient population to be submitted
- First data H1 2021
- **Covid-19**: Necessary steps taken to maintain Company operations and health and wellbeing of employees

Preclinical profile of SLN360 in NHP

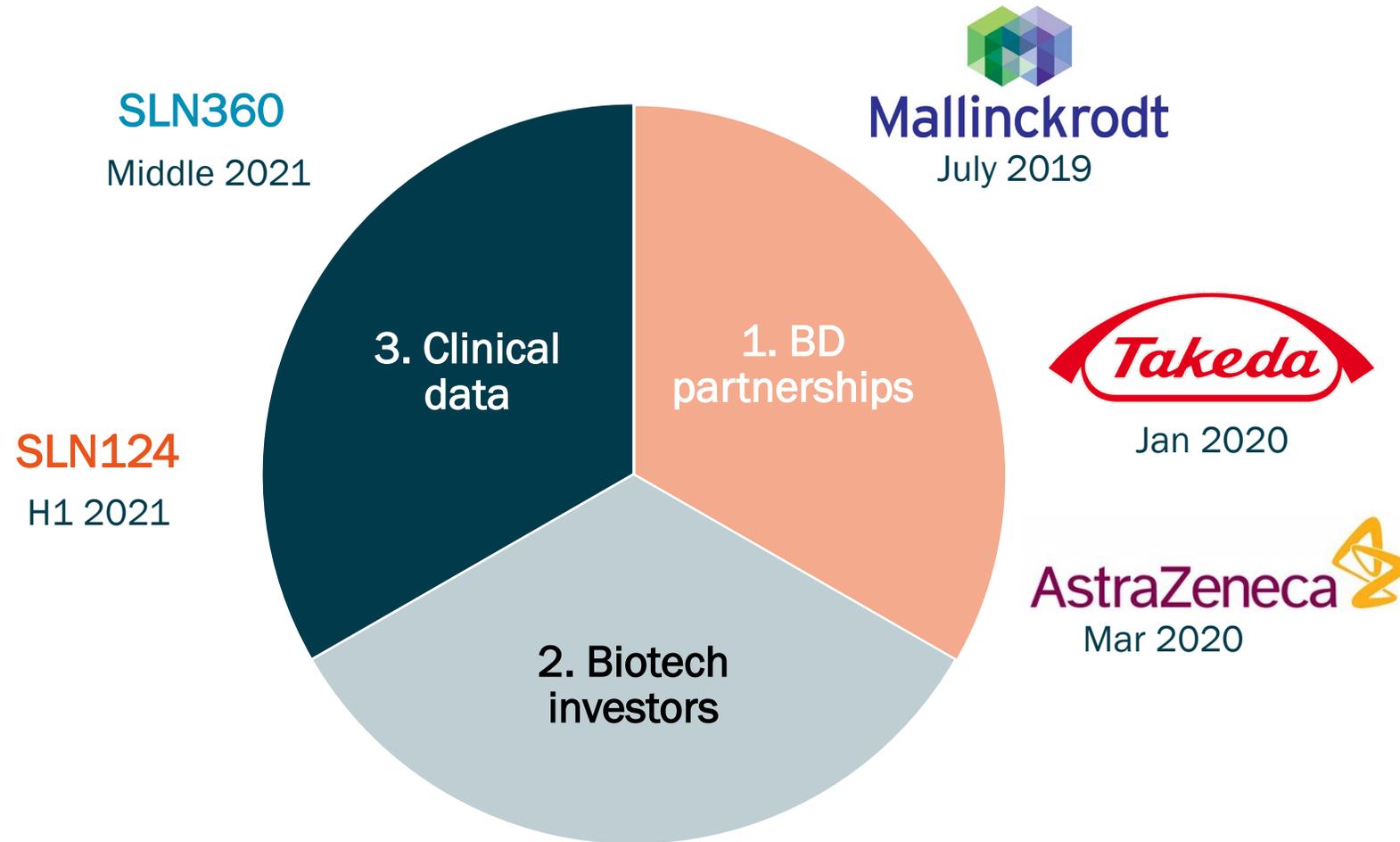




Other Recent Highlights from Q1

- CEO recruitment progressing well with announcement possible as soon as mid-2020
- Technology Evaluation Agreement with Takeda signed in January 2020
- Scientific Advisory Board formed, to be led by Professor Sir Gordon Duff
 - World leading scientists and clinicians to guide Silence's ground-breaking siRNA research programmes
 - Dr. Annemieke Aartsma-Rus, Dr. Art Levin, Dr. Henry Ginsberg, Dr. John Porter
- Silence Therapeutics, Inc. established, as well as a New York office, and 1st US-based employee (Dr. Barbara Ruskin, SVP, General Counsel and Chief Patent Officer)
- Expansion of US Investor Relations activity with presentations or attendance at several investor conferences

Platform validation: putting the pieces in place to unlock value



Silence Therapeutics - Summary



Valuable Platform	<ul style="list-style-type: none">> Reproducible, proprietary gene silencing (RNAi) therapeutics platform, rapidly generating internal pipeline and out-licensing options. Platform validated through collaborations with AstraZeneca, Mallinckrodt and Takeda
Growing Clinical Pipeline	<ul style="list-style-type: none">> SLN360 (CVD with high LP(a)) - IND/CTA in H2 2020> SLN124 (β-Thalassemia and MDS¹) - Phase Ib trial underway> SLN500 (C3) - IND/CTA in 2021
Strong Experienced Team	<ul style="list-style-type: none">> Pioneers in siRNA for over 18 years, growing clinical team, and experienced biopharma Board and Management team> New CEO recruitment ongoing
Target Selection	<ul style="list-style-type: none">> Focused on targeting indications in rare diseases and large population targets, including new medicines for cardiovascular disease and complement-mediated diseases
Strong Financial Position	<ul style="list-style-type: none">> Strong financial position with a cash runway extending beyond key clinical milestones such as SLN360 and SLN124 Phase I trial readouts

HQ in London



R&D in Berlin



+ New office in New York

Approx. 45 employees
across all sites

Notes:

¹ MDS = Myelodysplastic syndrome



Q&A