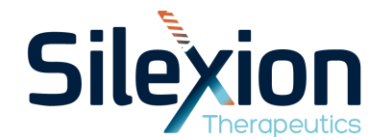




Silencing Oncogenes at the Level of Gene Expression

Corporate Presentation January 2025

Nasdaq: SLXN



Forward-Looking Statement

The statements contained in this presentation that are not purely historical are forward-looking statements. Our forward-looking statements include, but are not limited to, statements regarding our or our management team's expectations, hopes, beliefs, intentions or strategies regarding the future. In addition, any statements that refer to projections, forecasts or other characterizations of future events or circumstances, including any underlying assumptions, are forward-looking statements. The words "anticipate," "believe," "continue," "could," "estimate," "expect," "intends," "may," "might," "plan," "possible," "potential," "predict," "project," "should," "would" and similar expressions may identify forward-looking statements, but the absence of these words does not mean that a statement is not forward-looking. Forward-looking statements in this presentation may include, for example, statements about:

- the future performance of the Company, including Silexion's projected timeline for regulatory approvals of its product candidates; and
- the Company's future plans and opportunities.

The forward-looking statements contained in this presentation are based on our current expectations and beliefs concerning future developments and their potential effects on us. There can be no assurance that future developments affecting us will be those that we have anticipated. These forward-looking statements involve a number of risks, uncertainties (some of which are beyond our control) or other assumptions that may cause actual results or performance to be materially different from those expressed or implied by these forward-looking statements. These risks and uncertainties include, but are not limited to, the items in the following list:

- Silexion is a development-stage company and has a limited operating history on which to assess its business;
- Silexion has never generated any revenue from product sales and may never be profitable;
- The approach Silexion is taking to discover and develop novel RNAi therapeutics is unproven for oncology and may never lead to marketable products;
- Silexion does not have experience producing its product candidates at commercial levels, currently has no marketing and sales organization, has an uncertain market receptiveness to its product candidates, and is uncertain as to whether there will be insurance coverage and reimbursement for its potential products;
- Silexion may be unable to attract, develop and/or retain its key personnel or additional employees required for its development and future success;
- Additional factors relating to the business, operations and financial performance of Silexion.

Should one or more of these risks or uncertainties materialize, or should any of our assumptions prove incorrect, actual results may vary in material respects from those projected in these forward-looking statements. We undertake no obligation to update or revise any forward-looking statements, whether as a result of new information, future events or otherwise, except as may be required under applicable securities laws.

Company Overview

Clinical-stage company developing proprietary treatments for KRAS-driven cancers

KRAS-Focused RNA Interference Platform with Targeted Delivery

Silexion's siRNA platform technology is designed to silence oncogenes and prevent the production of the mutated KRAS proteins that drive cancer growth.
Inhibition of mutations important for pancreatic, colorectal and lung cancers

Promising Clinical Data in Locally Advanced Pancreatic Cancer

Phase 2 in Locally Advanced Pancreatic Cancer (LAPC) with first-generation siG12D-Loder siRNA with extended-release delivery showed trends for Overall Survival benefit (9.3 months) and Objective Response Rate in patients harboring KRAS^{G12D/V} mutations with Loder + chemo vs. chemo alone

Second generation, SIL204, siRNA optimized and administered with Integrated Treatment Regimen to enter Phase 2/3 trial

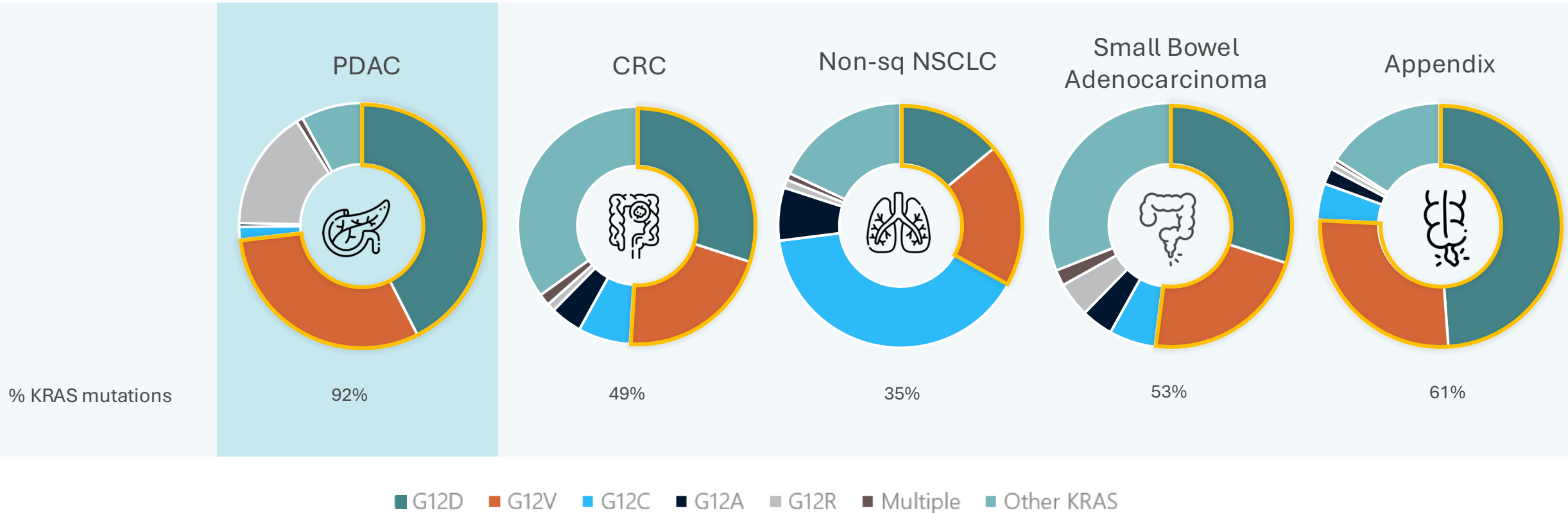
Compelling investment proposition

Listed on Nasdaq on August 2024 ("SLXN")

Late-Stage Ready Asset with Regulatory Path Forward

KRAS Oncogene is a Validated Target for Numerous Cancers

Prevalence of The Most Common Types of KRAS Mutations Across Cancers



KRAS is the most common oncogenic gene driver in human cancers with gastrointestinal cancers having high percentages of KRAS G12D/V mutations

Pancreatic Cancer Has One of the Highest Mortality Rates of All Major Cancers

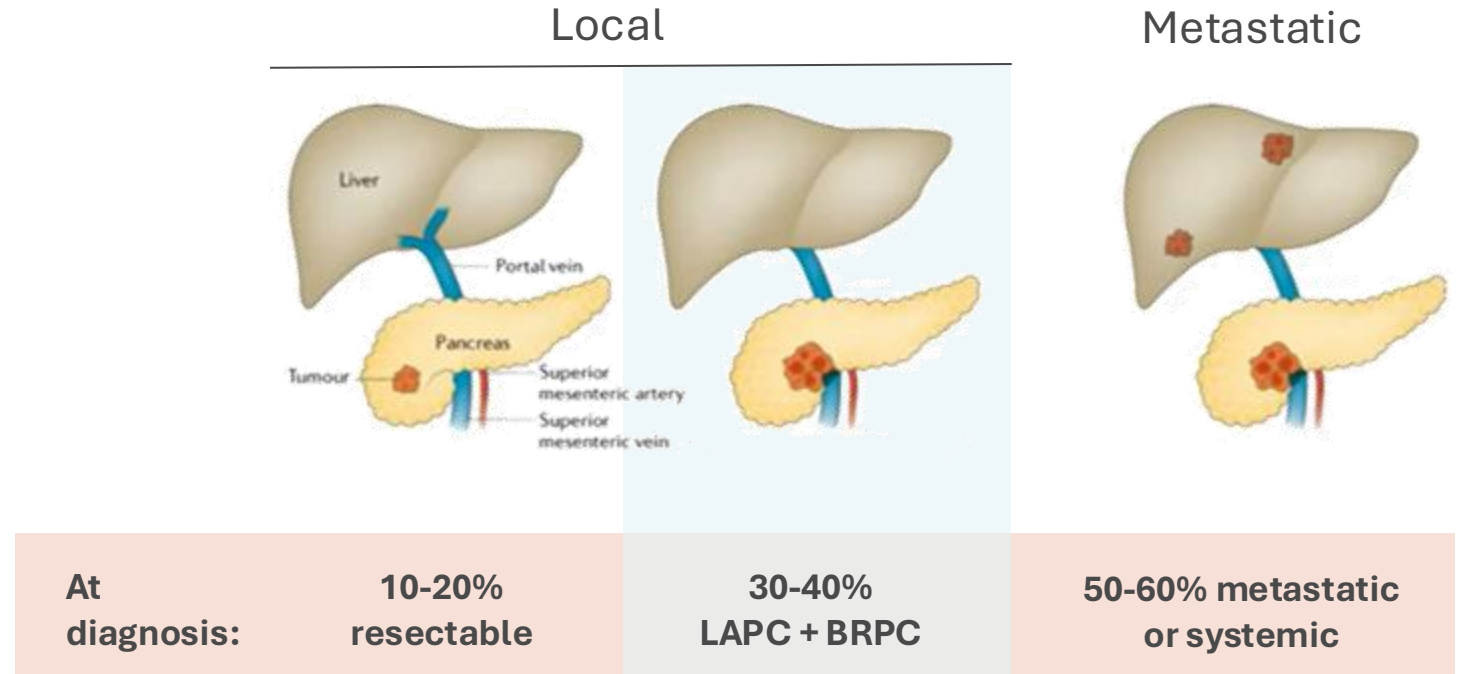
3rd leading cause today in the U.S.²

2nd leading cause by 2030²

12.8% 5-year relative survival (2014-2020) is one of the poorest in the U.S.³

Median overall survival for non-resectable PC populations is 14-17 months⁴

Types and Prevalence of Pancreatic Cancer^{4,5}



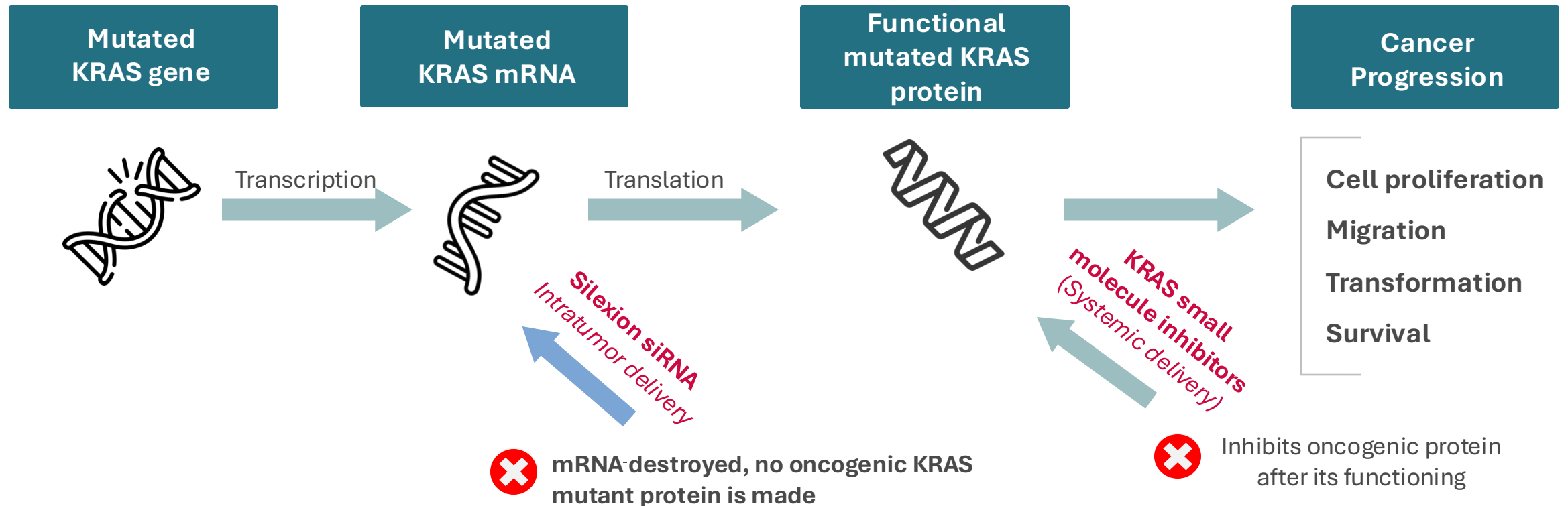
There are no effective treatment options for our intended indication LAPC

BRPC=borderline resectable pancreatic cancer; LAPC = locally advanced pancreatic cancer.

1. Bray F, et al. *CA Cancer J Clin.* 2024;74(3):229-263. 2. Hirshberg Foundation for Pancreatic Cancer Research. Pancreatic cancer Facts. <https://pancreatic.org/pancreatic-cancer/pancreatic-cancer-facts>.

3. National Cancer Institute. Cancer Stat Facts: Pancreatic Cancer. <https://seer.cancer.gov/statfacts/html/pancreas.html>. 4. Gemenetzis G, et al. *Ann Surg.* 2019;270(2):340-347. 5. Kleeff J, et al. *Nat Rev Dis Primers.* 2016;2:16022.

Innovatively Treating the Cancer-Driving KRAS at the Source and Site of Action



Moving closer to treating the mechanism of the cancer more efficiently and with a greater chance to overcome treatment-resistance

**Silexion
Innovative
Oncological
Approach May
Lead to
Significant
Improvement in
Clinical
Outcomes Over
KRAS Inhibitors**

**Limitation of currently approved and investigational
small molecule KRAS inhibitors:**

- Treatment resistance
- Low tolerability with adverse events such as rashes and GI side effects that require special monitoring
- Limited overall response rate and progression-free survival

Silexion's Approach

Inhibit oncogenic
KRAS synthesis
before it is active

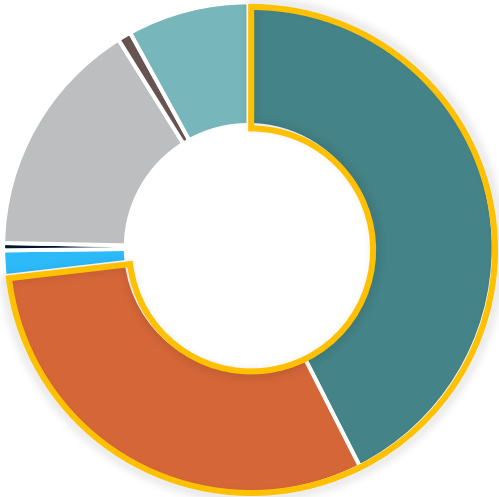
Integrated treatment
regimen to more
effectively treat both
the primary tumor
and micrometastases

Optimized siRNA to
enhanced stability
and broader activity
while maintaining
good safety

SIL204 is the Most Advanced siRNA for LAPC With a Significant Market Opportunity

KRAS mutations are present in ~92% pancreatic cancer cases¹

SIL204 covers > 74% of KRAS mutations in PDAC² while currently available KRAS G12C treatment are treating ~1.5%



■ G12D ■ G12V ■ G12C ■ G12A ■ G12R ■ Multiple ■ Other KRAS

Total Addressable Market in LAPC



| | U.S. | E.U. |
|-------------------------------------|---------------------|----------------------|
| Annual PC cases | 66,400 ⁴ | 146,477 ³ |
| KRAS-G12D/V mutated LAPC incidence* | ~16,000 | ~35,000 |

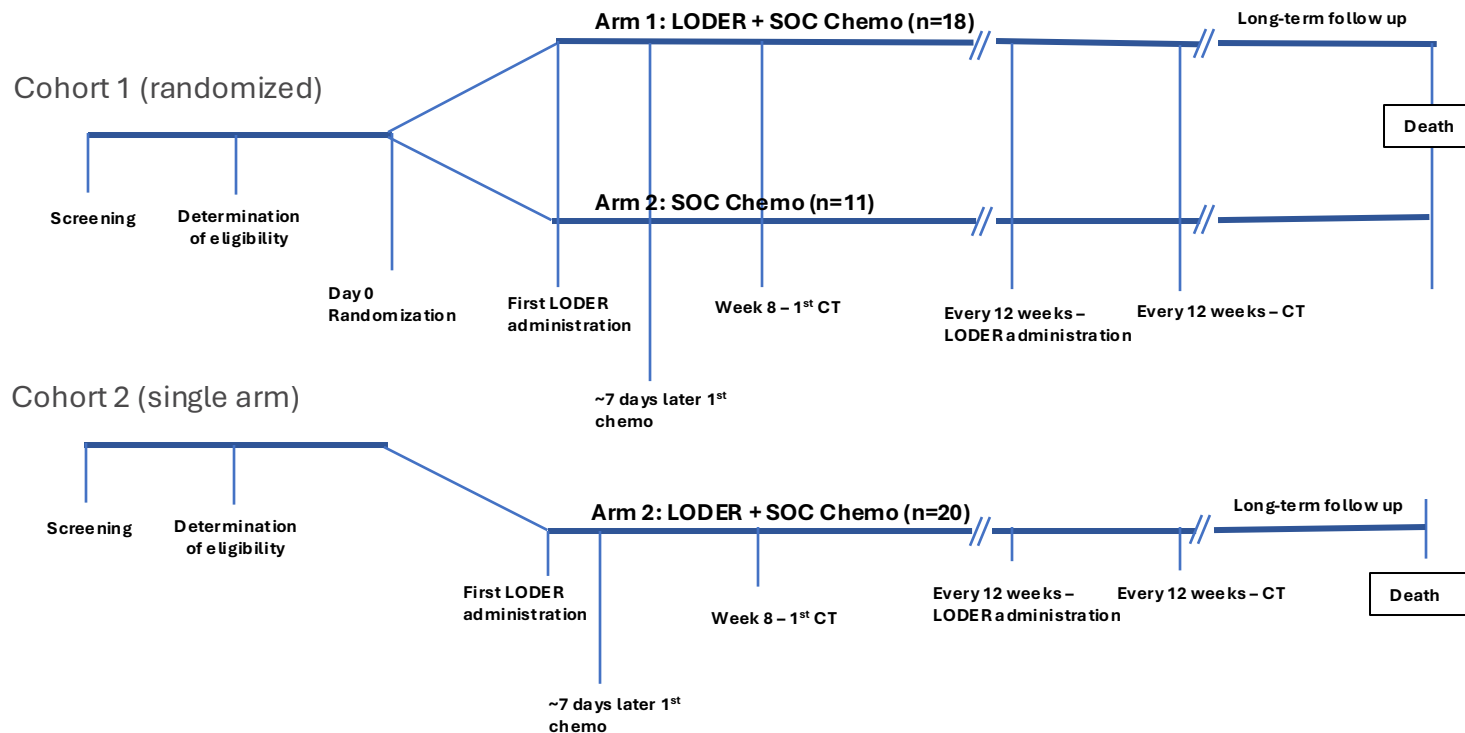
LAPC = localized advanced pancreatic cancer; ROW=rest of the world.
 *Number of KRAS G12D/V mutated LAPC were calculated based on KRAS mutations being present in 92% of pancreatic cancer patients, 70-75% with KRAS G12D and G12V mutations and 30-35% of cases being LAPC. Potential for cancers with pan KRAS G12x activity
 1. Lee, J.K. et al. *NPJ Precis Oncol.* 2022;6(1):91. 2. Yousef, A. et al. *NPJ Precis Oncol.* 2022;4(1):27. 3. Global Cancer Observatory. Pancreatic Cancer. 2022. <https://gco.iarc.who.int/media/globocan/factsheets/cancers/13-pancreas-fact-sheet.pdf>. 4. National Cancer Institute. Cancer Stat Facts: Pancreatic Cancer. 2023. <https://seer.cancer.gov/statfacts/html/pancreas.html>.

LODER

Phase 2 Trial Data

Phase 2 Trial of Loder Completed in 2023 – a Proof-of-Concept

Two-part, open label, study of LODER + SoC chemotherapy vs SoC chemotherapy alone across the U.S. and Israel in patients with non-resectable pancreatic cancer



Key inclusion criteria

Non-resectable without signs of metastasis

ECOG Status ≤ 1

Both cohorts all patients meeting inclusion/exclusion criteria randomized without checking for KRAS mutation status

Endpoints

Overall survival (OS)

Response rate (RR, RECIST v1.1)

Safety

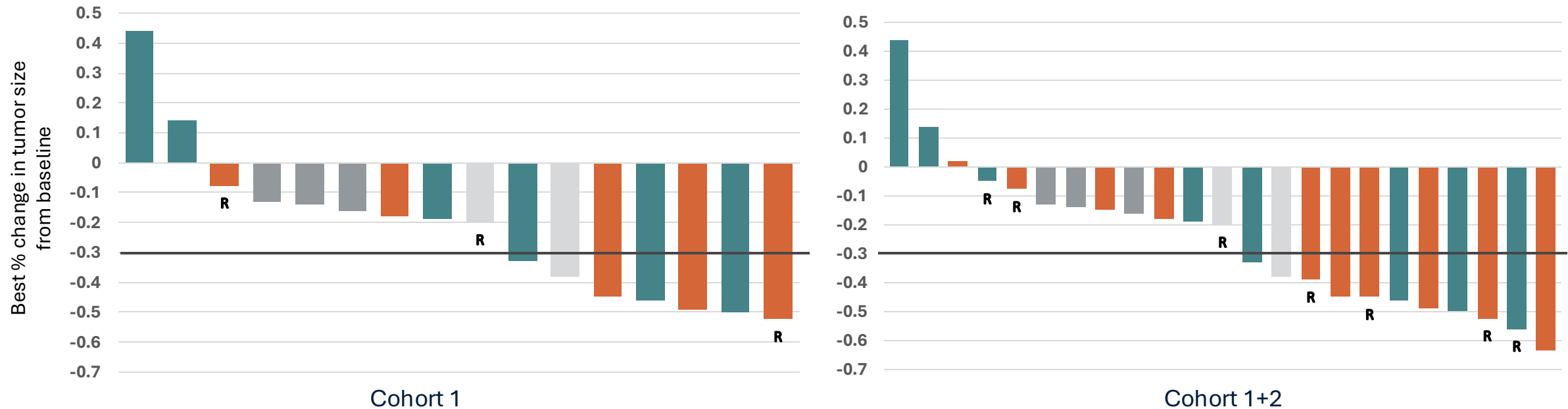
Tolerability

Baseline Characteristics and Cohorts Information

| | Cohort 1 (n=29) | Cohort 2 (n=20) |
|----------------------------------|--|---|
| Design/Arms | Randomized, controlled (SoC) | Single arm |
| Population | Locally advanced PC (LAPC) | Non-resectable (BRPC+ LAPC) |
| Nationality | 62% U.S. (4 sites) , 38% Israel (5 sites) | |
| Male/ Female % | 42% male; 58% female | |
| Median age (years) | 69.7 | 64.9 |
| KRAS Mutations | G12D/V*: Loder 11/12, Control 5/10 G12R*: Loder: 1/12, Control 5/10 | G12D/V*: Loder 7/9 G12R*: Loder: 2/9 |
| Avg Loder cycles | 2.8 | 2.1 |
| Total number of Loder injections | 370 | |
| SoC chemotherapy | gemcitabine/nab-paclitaxel (GnP) | (modified) FOLFIRINOX ((m)FFX) |

Due to results of a clinical trial indicating FOLFIRINOX's advantage over GnP as SoC chemotherapy, cohort 2's SoC chemotherapy was changed from GnP (used in cohort 1) to FOLFIRINOX.

Loder Treatment Led to Robust Objective Response Rate in Patients with LAPC Harboring G12D/V Mutations*



G12D G12V
 Chemo:
 Loder:
 R = Non-resectable tumor becomes resectable

| | Cohort 1 LODER+Chemo | Cohort 1+2 LODER+Chemo | Chemo |
|---------------------------------|-------------------------|---------------------------|----------|
| % Response | 55 (6/11) | 56 (10/18) | 20 (1/5) |
| % Response+ becoming resectable | 64 | 67 | 40 |

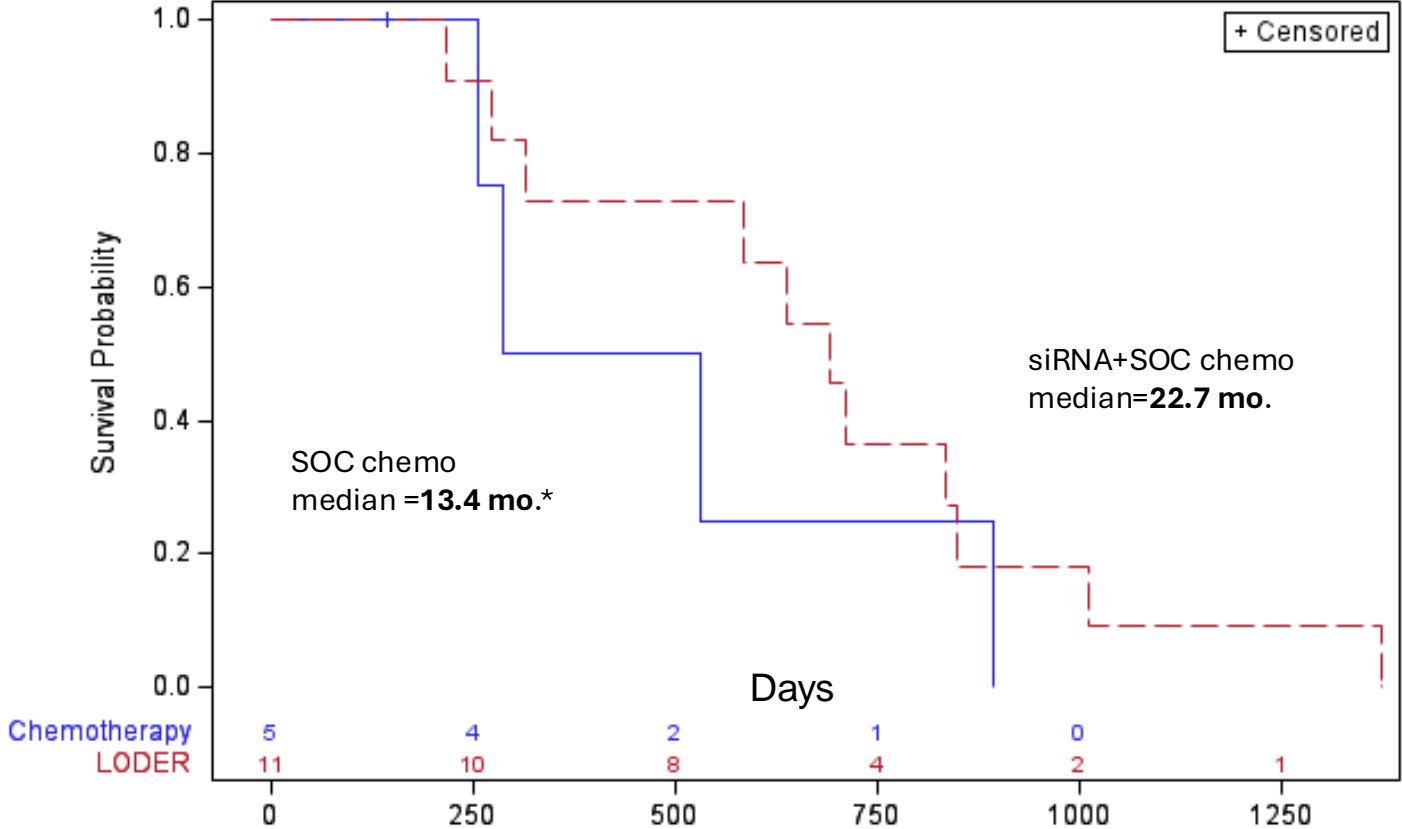
LAPC=locally advanced pancreatic cancer.

*Overall response rate was confirmed by RECIST 1.1 of the target tumor, as analyzed by sites.

Bar curves below the solid black line starting at y-axis -0.3 indicates criteria for positive RECIST response.

Cohort 1 Patients Treated with Loder Had 9.3 Months Improvement in Overall Survival

Overall Survival in Cohort 1



Hazard ratio (HR)=0.59, (95% CI, 0.18, 1.96, p=0.39)

Time to death is slower, 41% reduction in the rate of mortality.

Patients living longer with Loder+SOC vs. SOC

13 * SoC (Control) OS consistent with recent trials for LAPC (Gemenetis G, et al. *Ann Surg.* 2019;270(2):340-347).



Phase 2 Safety Results

Serious Adverse Events (SAEs) Related to Treatment in Patients with LAPC who Received LODER + Chemotherapy (treatment plus EUS-endoscopy administration procedure)

| SAE | LODER + SOC chemo (n=38) | |
|--|--------------------------|---------------------|
| | All grades n (%) | Grades 3-4 n (%) |
| Gastrointestinal disorders | 3 (8%) | 2 (5%) |
| Hematemesis | 1 (3%) | 0 (0%) |
| STOMACH ACUTE PAIN | 1 (3%) | 1 (3%) |
| Gastric hemorrhage | 1 (3%) | 1 (3%) |
| General disorders and administration site conditions | 2 (5%) | 0 (0%) |
| Fever | 2 (5%) | 0 (0%) |
| Hepatobiliary disorders | 3 (8%) | 2 (5%) |
| Cholangitis | 2 (5%) | 1 (3%) |
| Obstructive Hyperbilirubinemia | 1 (3%) | 1 (3%) |
| Infections and infestations | 2 (5%) | 2 (5%) |
| Sepsis | 1 (3%) | 1 (3%) |
| Pancreas infection | 1 (3%) | 1 (3%) |
| Depression | 1 (3%) | 1 (3%) |
| Injury, poisoning and procedural complications | 1 (3%) | 1 (3%) |
| procedural hemorrhage | 1 (3%) | 1 (3%) |
| Nervous system disorders | 1 (3%) | 1 (3%) |
| Presyncope | 1 (3%) | 1 (3%) |

Loder Was Overall Well Tolerated

- The Phase 2 PoC clinical trial investigators reported that Loder treatment was well tolerated; Safety events were primarily related to procedure
 - Intratumor administration of extended-release siRNA via endoscopy (EUS) is safe
- No Treatment Emergent Adverse Events (TEAEs) leading to study discontinuation related to Loder treatment
- No meaningful observations in any vital sign parameter nor any physical examination findings in the study
- Independent Drug Safety Monitoring Board (DSMB) Reviews had no safety concerns nor safety restrictions
- In a subset analysis, no measurable amount of Loder was detected (<BLQ) in any plasma samples suggesting low systemic levels

Building upon the Loder results, we optimized:

siRNA:

- Enhanced stability
- Broadening activity

Integrated Treatment Regimen
for primary tumor and
micrometastases

SIL204

KRAS G12D/V siRNA with
potential for pan-KRAS^{G12x} activity

Leveraging Loder Clinical Data to Further Improve SIL204 Potential Efficacy and Safety

| | LODER | SIL204 |
|--|--|--|
| siRNA target | KRAS G12D/V+ KRAS amplify | KRAS G12D/V+ KRAS amplify, potential pan KRAS ^{G12x} |
| Stability in human serum (HS) and rat pharmacokinetics | HS <1 hr | HS > 48 hrs, Remains at substantial levels for >56 Days in plasma and tissues after single subcutaneous administration to rats |
| Access to tumor cell site of action | No hydrophobic lead | Added hydrophobic lead to increase siRNA access into cell |
| Route and Ease of administration | EUS-endoscopy* with larger needle; Required loading device | S.C. administration and EUS Endoscopy* with smaller and more flexible needle; No loading device needed |
| Improvement in OS with chemo vs. chemo alone | + 9.3 months HR=0.59 Reduction in rate of death by 41% compared to chemotherapy alone | TBD in Phase 2/3 trial with expected improvement in OS |
| Safety | Generally well tolerated; Safety events were primarily related to procedure | Expectations for good safety profile |

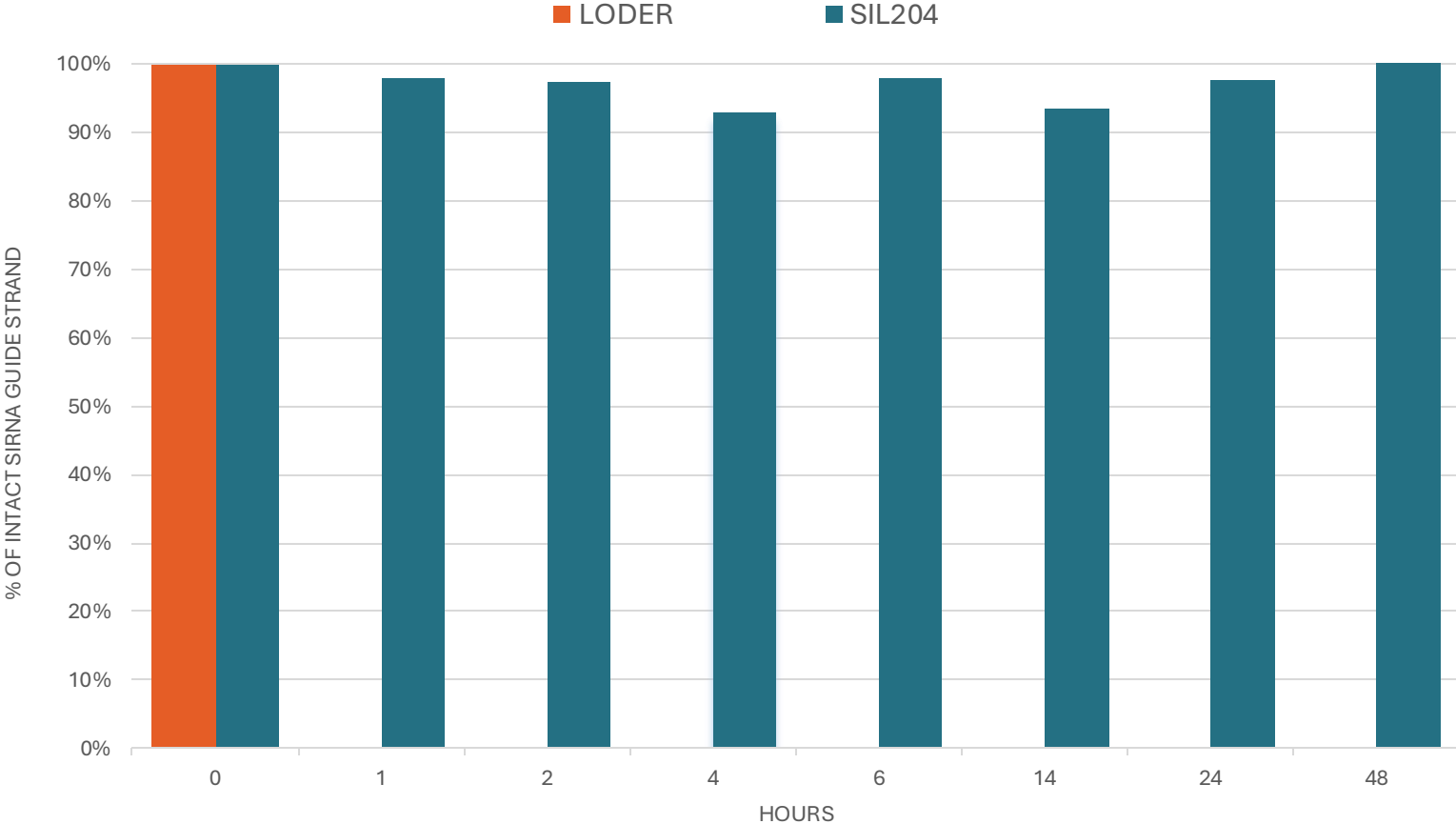
HR=Hazard Ratio.

18 *EUS endoscopy is a standard procedure used to obtain ultrasound guided biopsies once every 3 months.

SIL204 is Stable In Vitro for Over 48 Hours in Human Serum

siRNA strand placed in human serum and tested for stability

Stability of siRNA Strand in Human Serum



Potentially longer effectiveness of siRNA

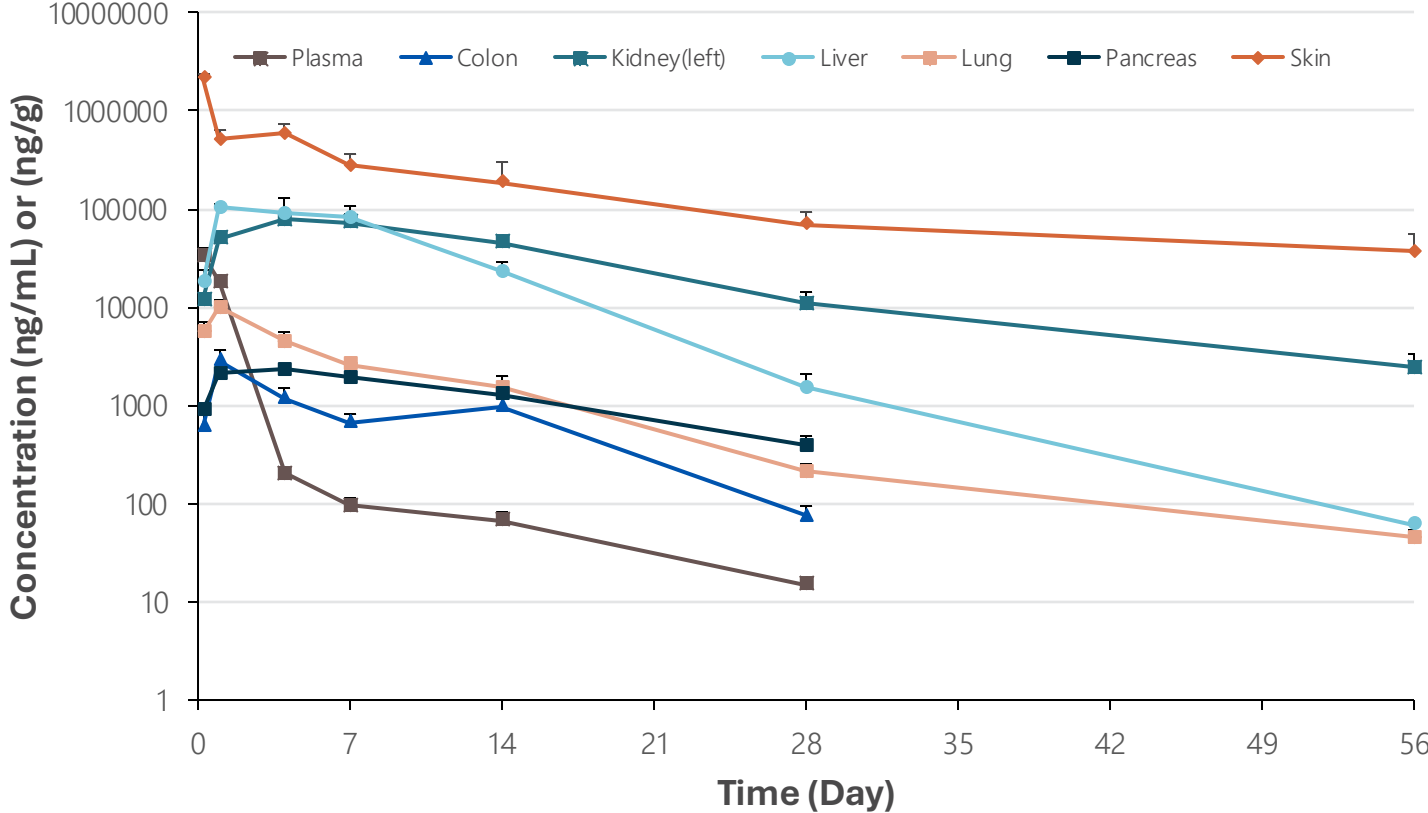
Greater ability to diffuse throughout the fibrous tumor environment

Potential in other indications

Previous studies have shown siG12D (Loder) half-life to be 5 min in human serum.

SIL204 Remains at Substantial Levels for >56 Days in Plasma and Tissues

Single subcutaneous administration SIL204 solution (10mg, not formulated) to Sprague Dawley rats



Potential for longer effectiveness of siRNA for treating micrometastases with dosing on monthly basis

SIL204 Shows Broad Inhibition Across Human KRAS Mutations at Sub-Nanomolar Concentrations

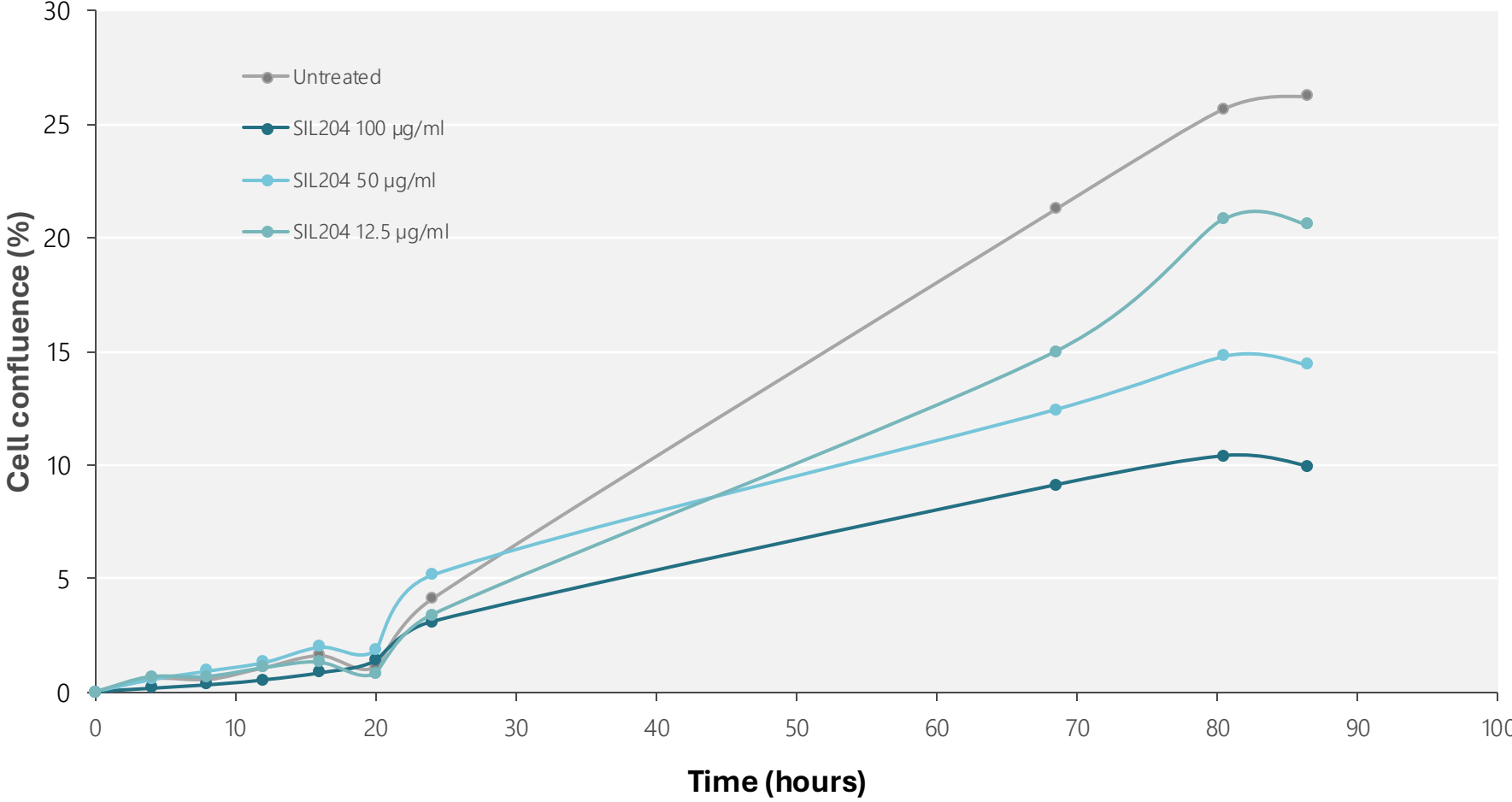
SIL204 maintains and expands the silencing activity of the Loder

Model is a co-transfection setup in mouse Hepa1-6 cells with Dual-Glo reporter plasmids.

| Mutation | Negative siRNA Control | WT KRAS | KRAS G12D | KRAS G12V | KRAS G12C | KRAS G12R | KRAS Q61H* | KRAS G13D* |
|-----------------------|------------------------|---------|-----------|-----------|-----------|-----------|------------|------------|
| IC ₅₀ (nM) | | 0.16 | 0.19 | 0.44 | 0.47 | 0.59 | 0.24 | 0.37 |
| MAX Inhibition (%) | 0-7 | 91 | 90 | 80 | 73 | 71 | 88 | 88 |

IC₅₀=half-maximal inhibitory concentration.

SIL204 Robustly Inhibits Growth of Human Pancreatic Tumor Cell Line (Panc-1) in Dose-Dependent Manner



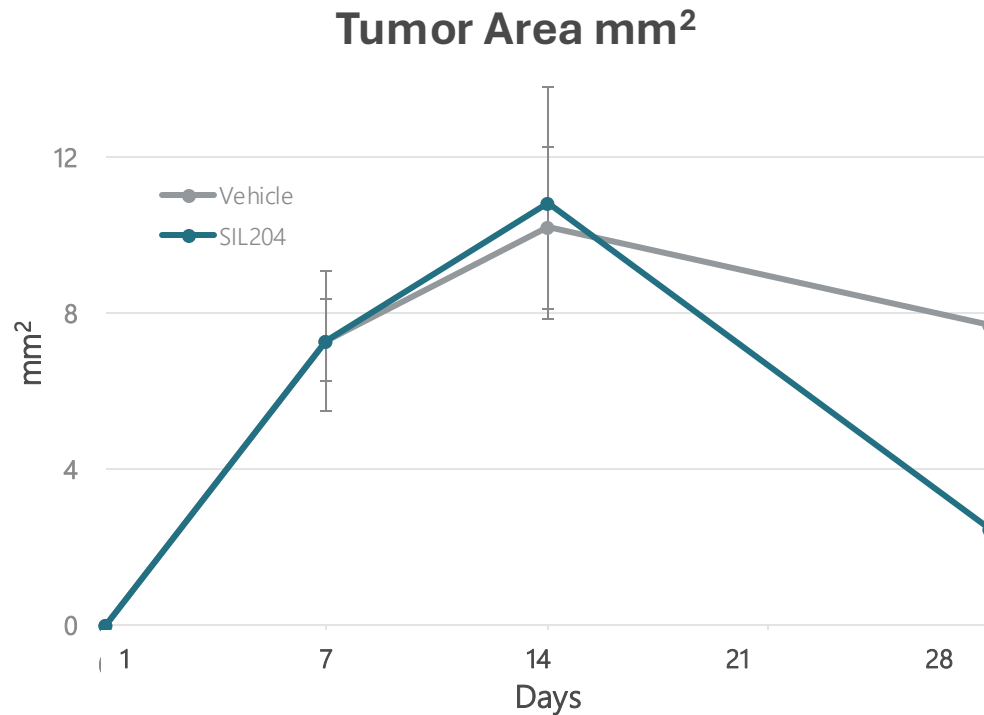
SIL204-MP Inhibited Human Pancreatic Cancer Xenograft Growth in Mice

SIL204 significantly reduced tumor area and increased tumor necrosis

~Day -7: Panc-1 (G12D mutation) cells were xenografted to mice (s.c.) and grown to ~200mm³

Day 1: SIL204-MPs or PBS (phosphate buffer Saline) was injected sub-cutaneous into animals

Day 30: Tumors were dissected, and histological slices from the center of the tumor taken and analyzed for necrosis



After 4 weeks 50% of Panc-1 tumors showed complete necrosis



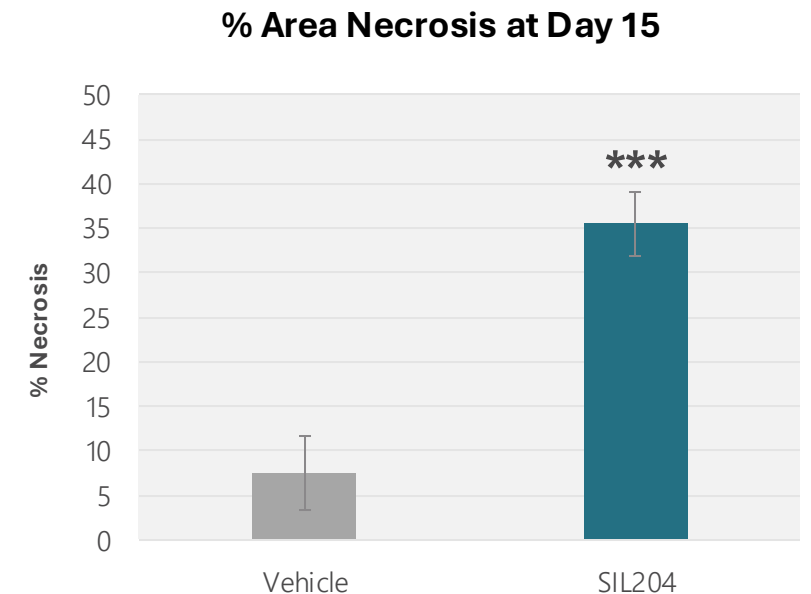
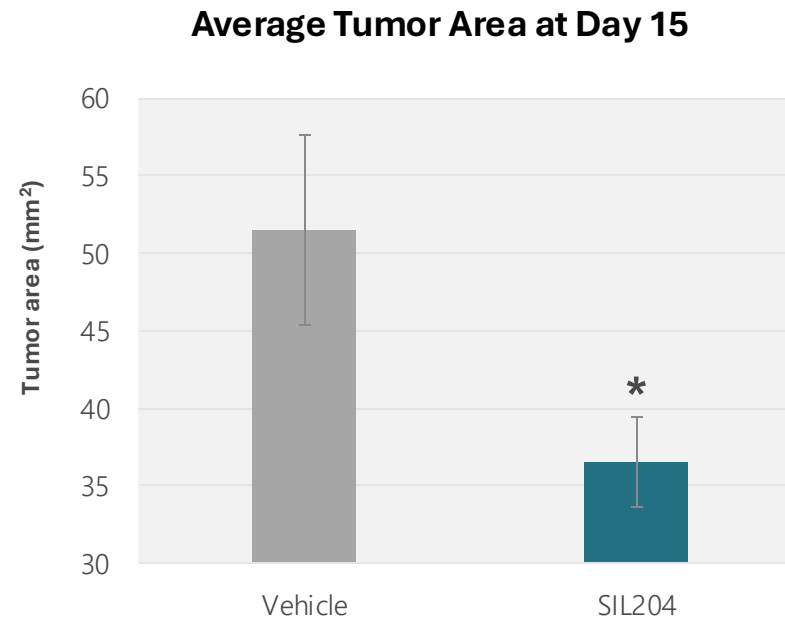
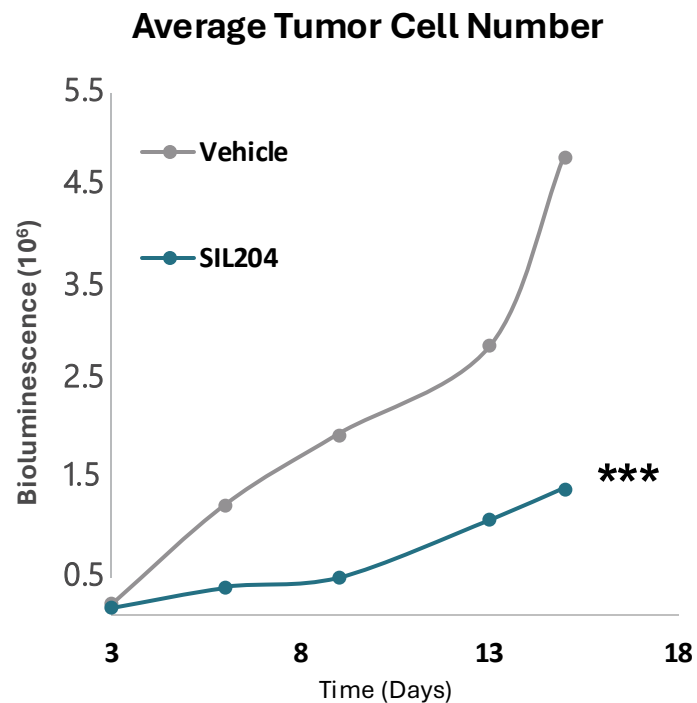
SIL204 Inhibited Human Pancreatic Cancer Xenograft Growth in Mice

SIL204 significantly reduced tumor volume and growth while increasing the necrosis (cell death) within the tumor

Day 1: Capan-1 (G12V) luciferase cells were xenografted to mice (s.c.) concurrently with SIL204 formulated in extended-release microparticles

Days 3-15: mice were evaluated for bioluminescence to evaluate relative tumor cell counts

Day 15: tumors were removed, area determined and analyzed by histology for % necrosis from tumor center slice



SIL204 Development Strategy in LAPC

| 2023 | 2024 | H1 2025 | H2 2025 | H1 2026 | H2 2026 |
|---|--|--|--|--|--|
| <p>✓</p> <p>Clinical proof of concept for Loder in LAPC in an approvable endpoint for FDA</p> | <p>✓</p> <p>Optimization of siRNA on various fronts; selection of SIL204</p> <p>Received guidance on trial design from the German Federal Institute for Drugs and Medical Devices (BfArM), intratumor administration</p> | <p>✓</p> <ul style="list-style-type: none"> Initiate toxicology studies SIL204 GMP production API (SIL204) <p>Meeting with German authorities for regulatory buy-in/scientific advice (BfArM) on integrated regimen, and plans to proceed to Phase 2/3 trial</p> | <p>GMP production injectable formulation</p> | <p>Initiate Phase 2/3, LAPC Germany/Israel</p> <p>Submit CTA in E.U. for Phase 2/3</p> | <ul style="list-style-type: none"> Leverage safety clinical data from first trial segment Phase 2/3 Pre-IND meeting FDA Submit IND to FDA Expand Phase 2/3 to USA\additional EU, etc |

✓ Indicates completed activity. Unmarked activities to be performed.

Focused Pipeline to Address KRAS-driven Solid Tumor Localized Cancers

| Program | Indication | Setting | Discovery | Preclinical | Phase 1 | Phase 2 | Phase 3 | Status/ Anticipated Milestone |
|--|------------------------------------|-------------------------|-----------|-------------|---------|-------------------|---------|---|
| LODER siG12D + KRAS amplify with extended release PLGA delivery system | Locally advanced pancreatic cancer | Adjunct to chemotherapy | | | | Phase 2 Completed | | Phase 2 completed: observed 9.3 months improvement with LODER over SOC. Continue development of SIL204. |

Current Focus: Optimized siRNA formulation with Integrated Treatment Regimen

| | | | | | | | | |
|---|------------------------------------|-------------------------|--|--|--|--|--|--|
| SIL204 (Integrated treatment regimen) KRAS G12D/V (primary), pan-KRAS (secondary) | Locally advanced pancreatic cancer | Adjunct to chemotherapy | | | | | | H1 2026: CTA submission in E.U. for Phase 2/3 1H 2026: Initiate Phase 2/3 |
| | Colorectal cancer | Adjunct to chemotherapy | | | | | | H2 2025: Preclinical |

Strong Intellectual Property Protection

IP Portfolio

10 patents issued worldwide | 6 patents pending worldwide

| Technology | Term |
|--|---|
| Compositions for inhibition of KRAS expression and treatment regimens therewith | 2045* (provision patent) |
| Inhibition of KRAS expression and methods of use thereof | 2043* (International patent application) |
| siRNA against KRAS G12x for regional perineural invasion or pain associated with a solid tumor | Pending US/EU, expected term till 2040 |
| Nanoparticles | 2016-2032 (E.U.) |
| G12D siRNA Sequences | 2014-2032 (U.S.) |
| Loder | 2015-2030 (U.S.) 2014-2029 (E.U.) |

* Exclusivity can be extended under country-specific regulatory-based extension rules.

World-Renowned Expert Scientific Advisory Board



Eileen M. O'Reilly, MD

Memorial Sloan Kettering, NY, NY

Winthrop Rockefeller Endowed Chair of Medical Oncology; Co-Director, Medical Initiatives, David M. Rubenstein Center for Pancreatic Cancer Research; Section Head, Hepatopancreatobi



Thomas Seufferlein, MD

University Hospital Ulm, German

Director of Internal Medicine University Hospital Ulm, President German Cancer Society



Milind Javle, MD

The University of Texas & MD Anderson Cancer Center, Houston, TX

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Matthew Katz, MD

The University of Texas & MD Anderson Cancer Center, Houston, TX

Department Chair, Department of Surgical Oncology, Division of Surgery and Professor.



Philip A. Philip, MD

Henry Ford Health, Detroit, MI

Director, Gastrointestinal Oncology; Co-Director, Pancreatic Cancer Center; Medical Director, Research and Clinical Care Integration, Henry Ford Cancer Institute



Andrew M. Lowy, MD

UC San Diego, San Diego, CA

Chief, Division of Surgical Oncology; Professor of Surgery



Talia Golan, MD

Sheba Tel Hashomer Hospital,, Israel

Head, Sheba Pancreatic Cancer Center - SPCC



Mark A. Schattner, MD

Memorial Sloan Kettering, NY, NY

Chief, Gastroenterology, Hepatology and Nutrition Service



Hana Algul, MD

Technical University of Munich, Germany

chair for tumor metabolism; Director of the Comprehensive Cancer Center Munich, Germany at the Klinikum rechts der Isar, and Mildred-Scheel-professor and

Highly Experienced Leadership Team



Ilan Hadar, MBA Chairman and Chief Executive Officer

> 25 years of multinational managerial and corporate experience with pharmaceutical and high-tech companies



Mitchell Shirvan, PhD, MBA Chief Scientific and Development Officer

> 25 years of experience in R&D, innovation and discovery in biotech companies



Mirit Horenshtein Hadar, CPA Chief Financial Officer

> 15 years of corporate finance experience in senior financial positions of public companies and privately held companies, in the pharmaceutical and high-tech industries



Ilan Levin, Director

Former Chairman & Chief Executive Officer of Moringa Acquisition Corp with 25 years of experience as an executive and venture capital/private equity investor in high-tech, Israel-related ventures



Investment Highlights

Advanced RNA
therapeutic candidate in
oncology

- Clinical-stage company with proprietary oncogene siRNA platform
- Integrated Treatment Regimen (To more effectively treat both the primary tumor and micrometastases)
- Phase 2 clinical trial with first generation Loder in LAPC showed 9.3 months improvement in FDA approvable endpoint of overall survival
- Lead Candidate SIL204 with enhanced siRNA stability and broader activity

Late-Stage Ready Asset
with Regulatory Path
Forward

- Guidance received from German Federal Institute for Drugs and Medical Devices (BfArM) on Phase 2/3 trial
- Submit CTA in E.U. in 1H 2026 and initiate Phase 2/3 trial of SIL204 in 1H 2026
- Plan for U.S. IND submission with clinical safety data from limited number of patients the trial in E.U.

Strong Partnerships with
Solid IP Portfolio

- Established partnerships for GMP production of siRNA and formulations
- PCT submitted with favorable international review for claims for siRNA composition of matter and use, IP exclusivity through December 2043 plus extension

Thank You

Ilan Hadar

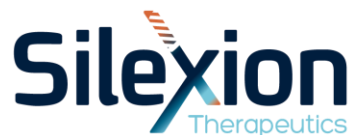
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