



Next-Generation Ionizable Lipid Nanoparticles: A Modular Platform for Therapeutic Payloads

Alex López, PhD

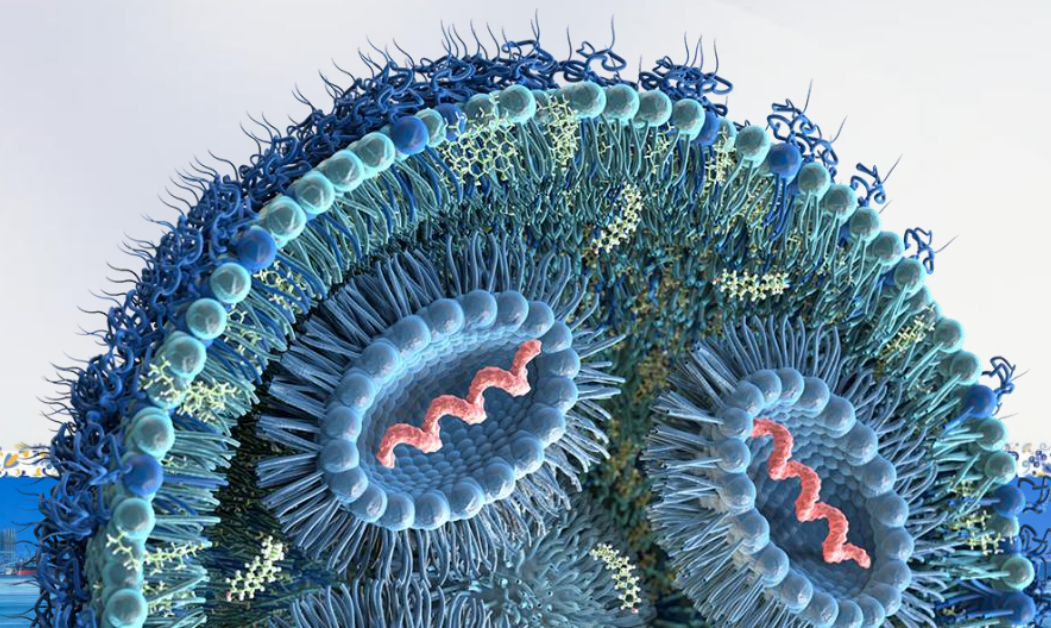
Senior Researcher at Certest Pharma

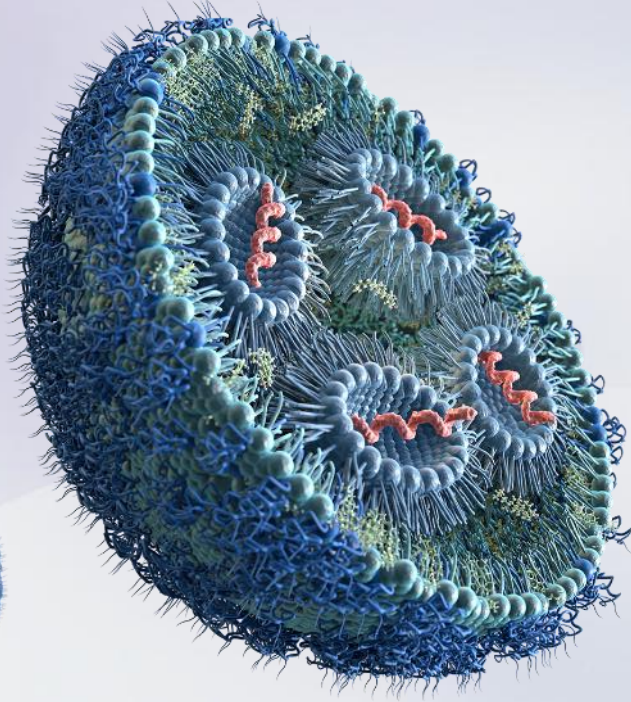


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08/07/2026

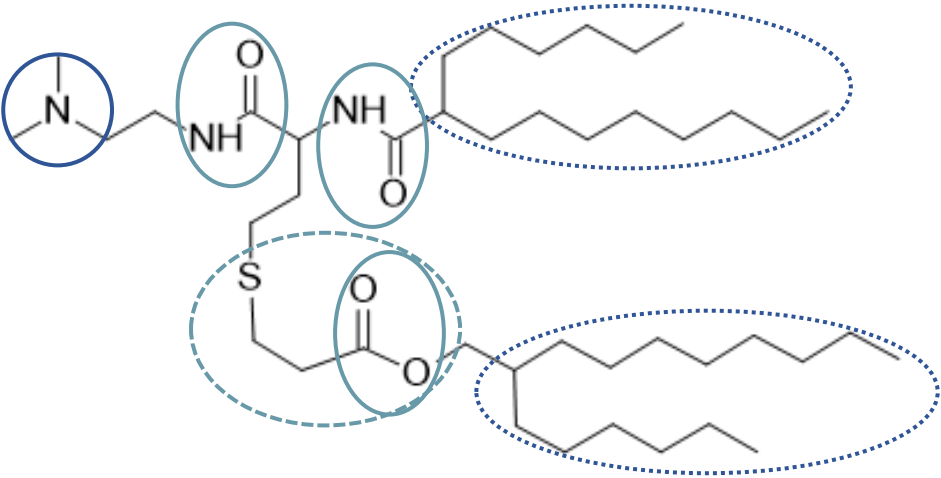
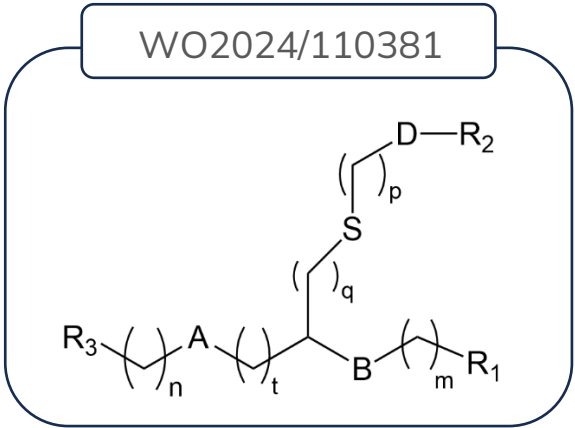


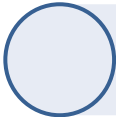


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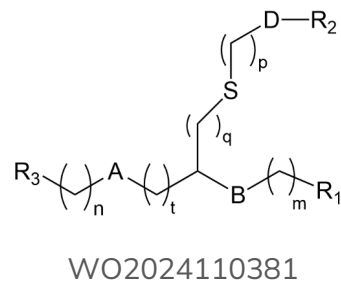
Sequential Thiolactone Amine Acrylate Reaction



-  Ionizable head
-  Branched tails
-  Biodegradable groups
-  Superior biodegradability

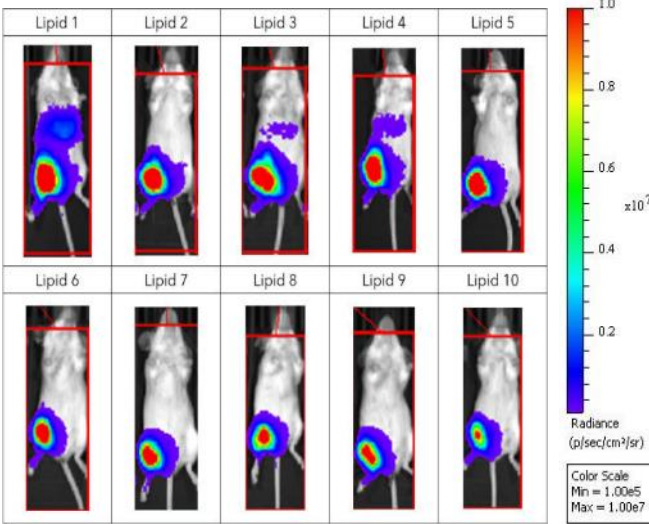
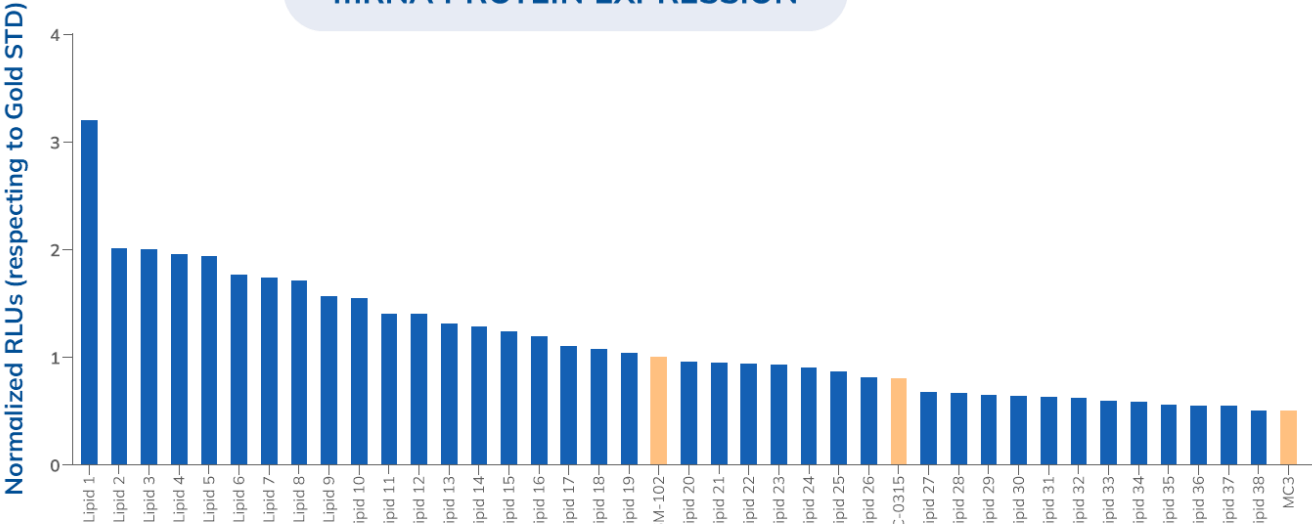


Clinical-Ready ILs

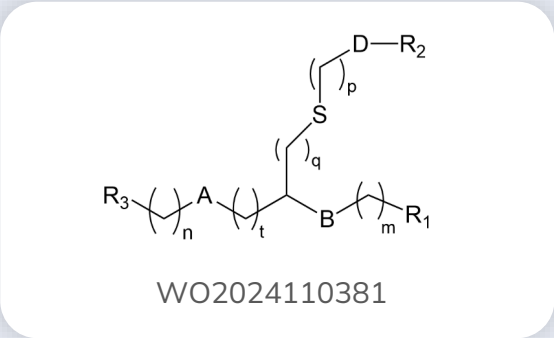


- Large-scale in vivo screening
- Potency and tolerability selected
- Broad compatibility across multiple cargos
- Validated in non-human primates
- Two Phase 1 clinical programs

mRNA PROTEIN EXPRESSION

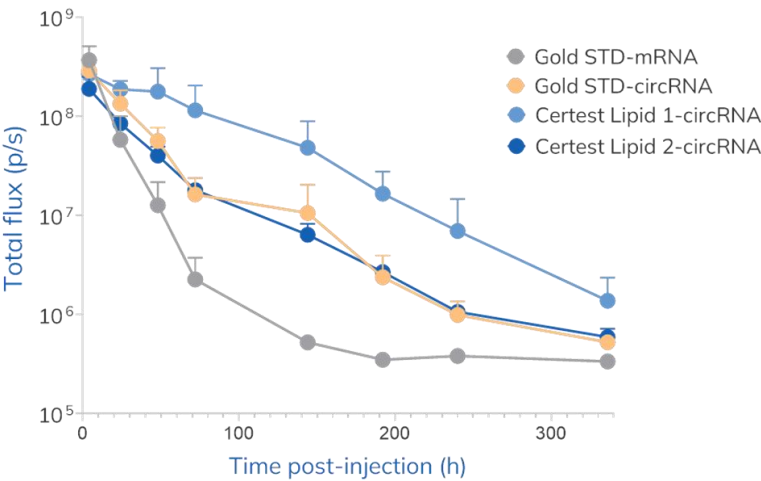


Clinical-Ready ILs

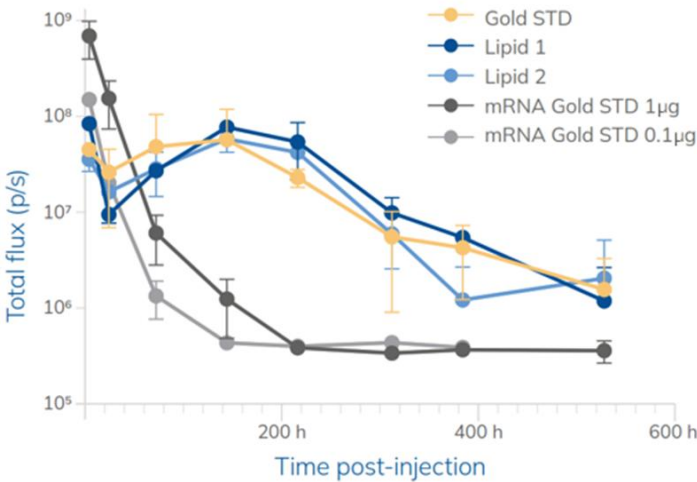


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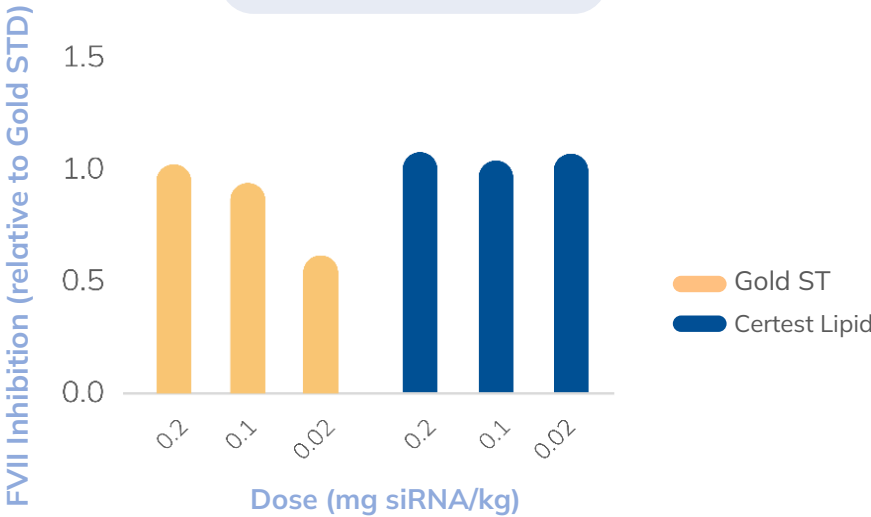
circRNA



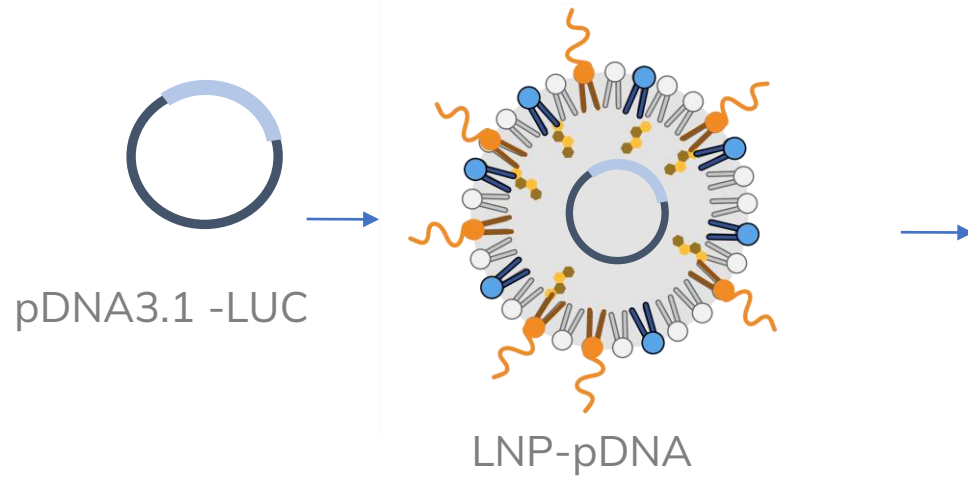
saRNA



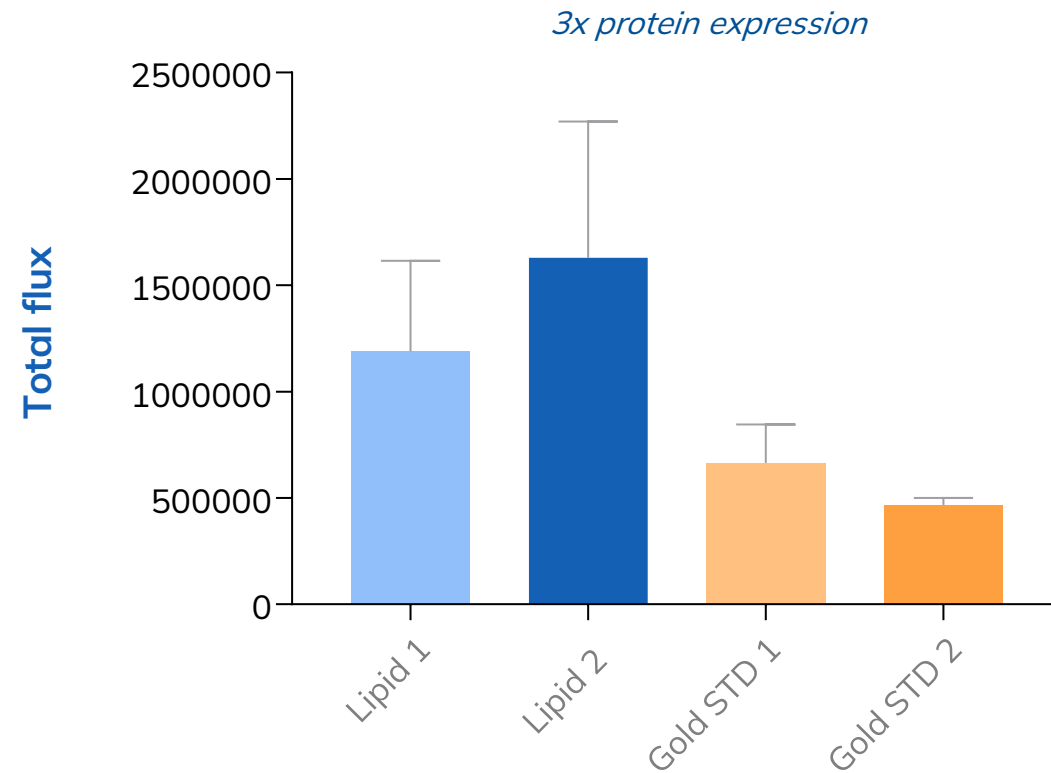
siRNA vaccines



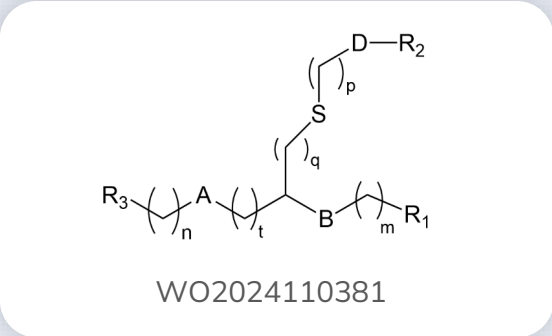
Delivery of pDNA with Certest lipids



Luciferase expression vs. gold-standard ionizable lipids

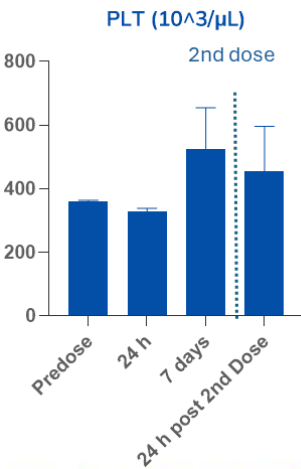
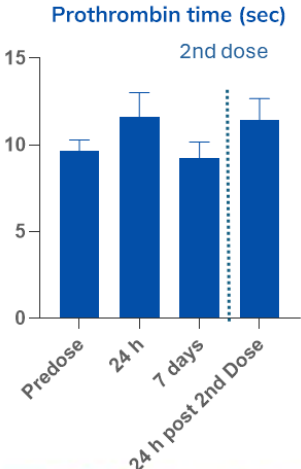
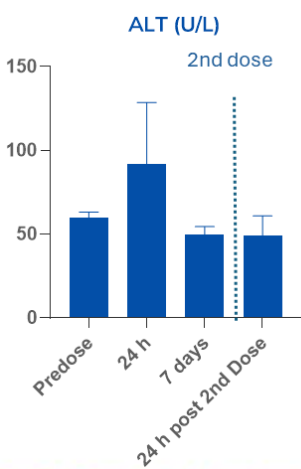
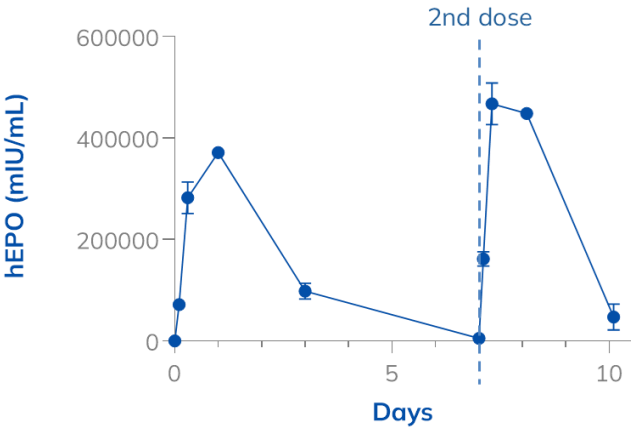


Clinical-Ready ILs

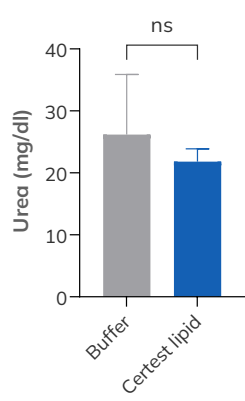
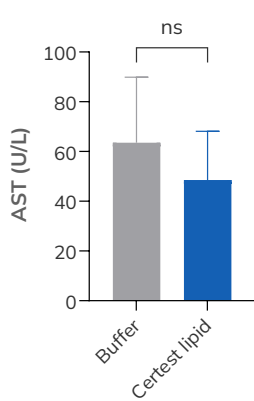
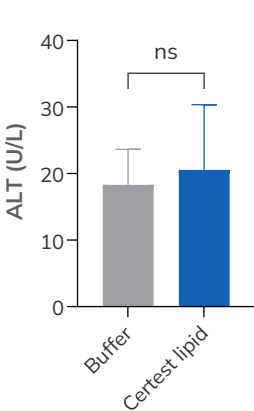


- Large-scale in vivo screening
- Potency and tolerability selected
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- Validated in non-human primates
- **Two Phase 1 clinical programs**

NonHuman Primates studies

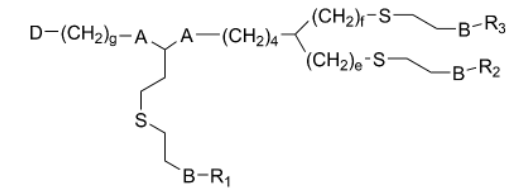


Tolerability studies (2.5 mg/kg dose) in Mice



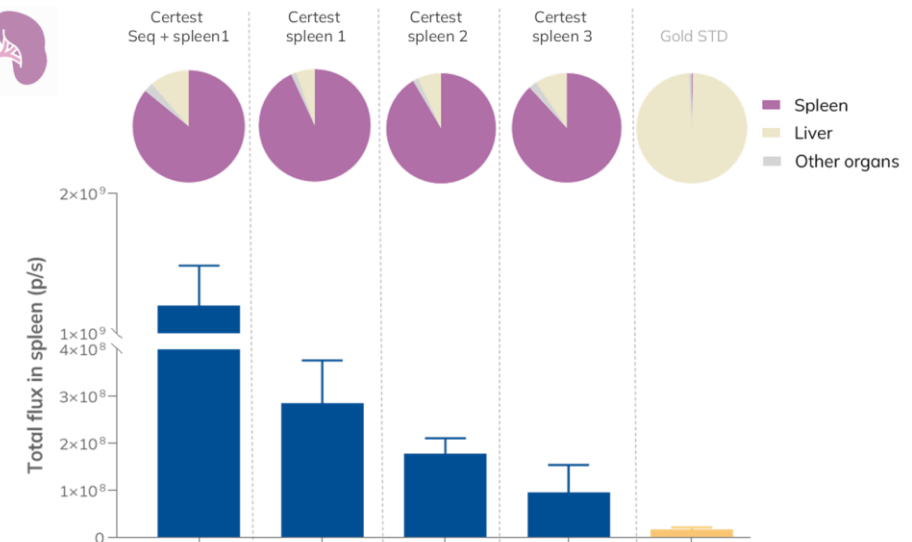
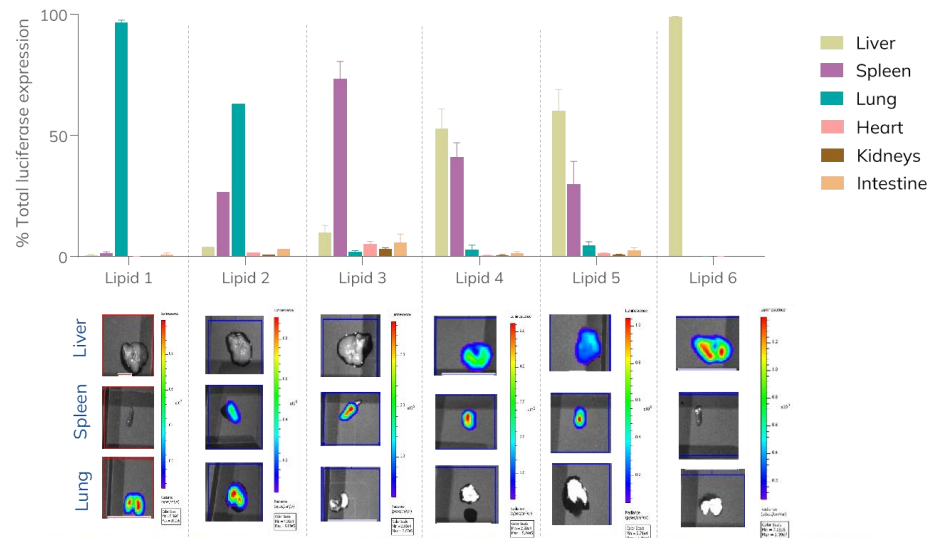
- Selective spleen delivery profiles
- Minimal liver expresión
- Compatible with standard 4-component LNPs
- Extrahepatic biodistribution capabilities
- Optimized for tissue-selective delivery

Extrahepatic ILs

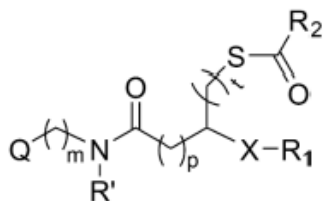


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In vivo TISSUE SELECTIVITY



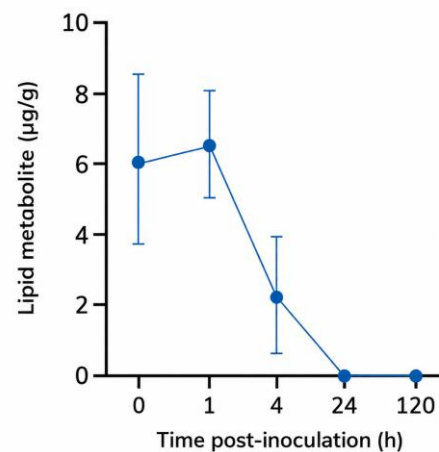
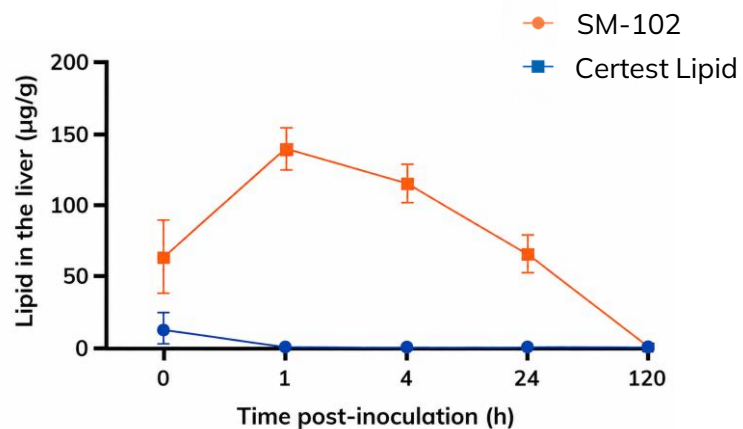
Highly-Biodegradable ILs



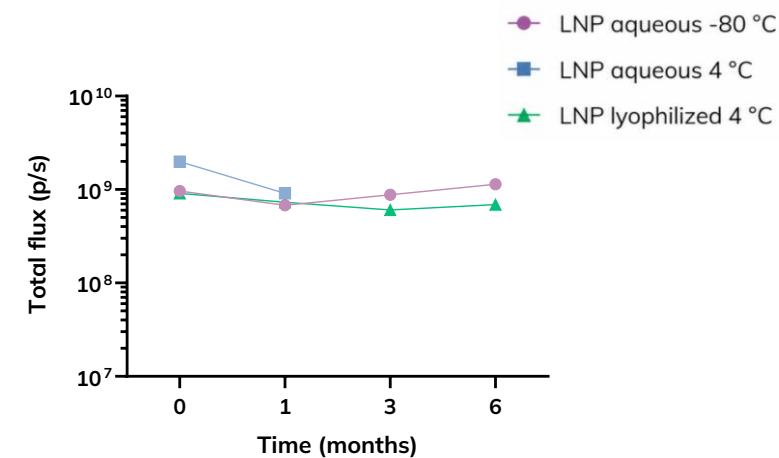
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- Enhanced biodegradability and clearance
- Stable under standard storage conditions
- Improved tolerability at high doses
- Designed for repeat-dose applications
- Optimized for high-exposure RNA therapies

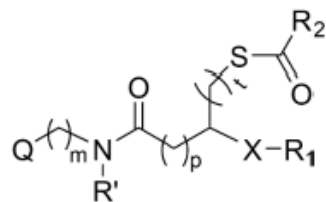
Biodegradability studies



LNP stability



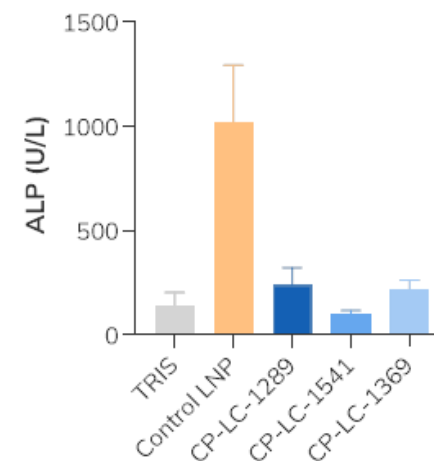
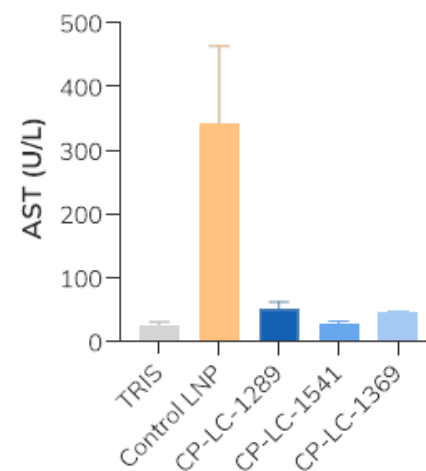
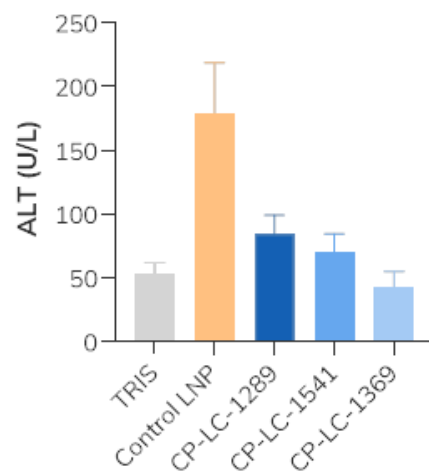
Highly-Biodegradable ILs



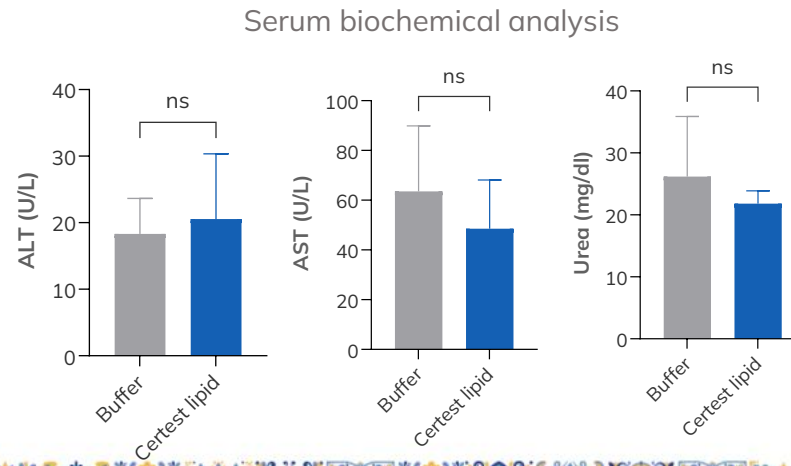
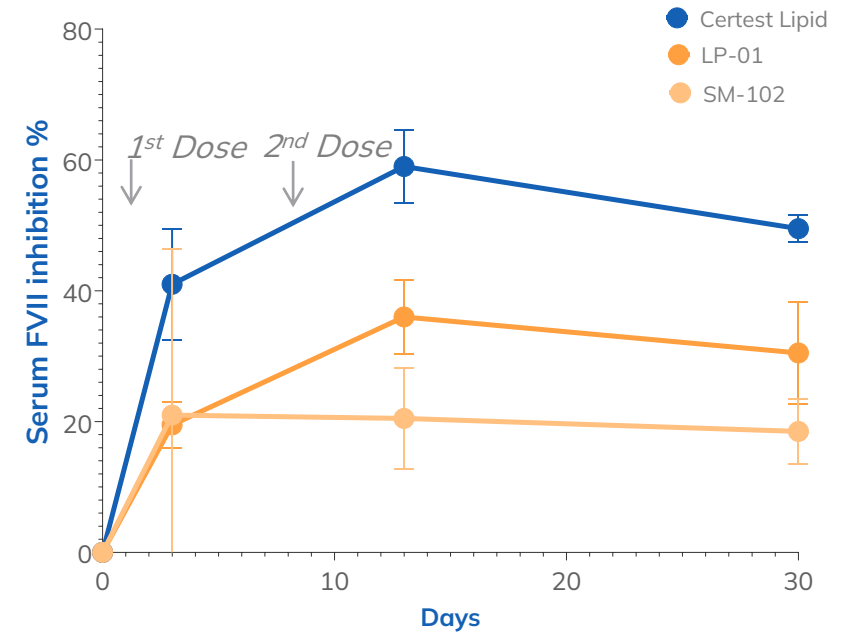
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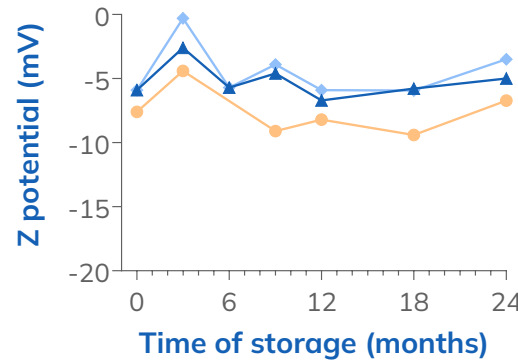
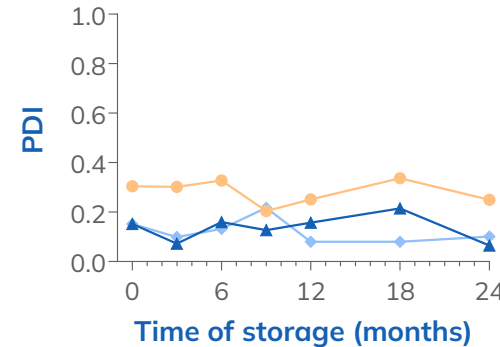
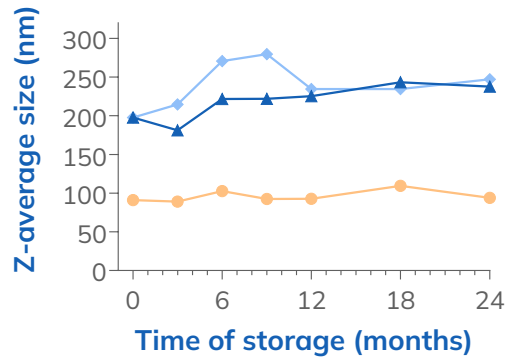
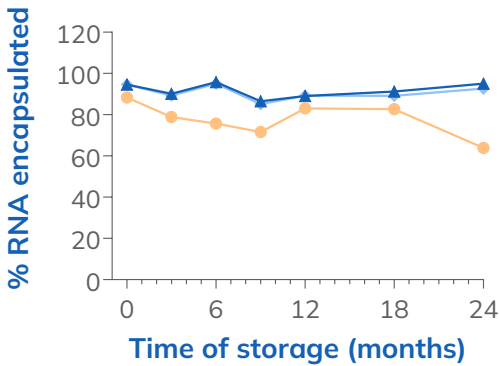
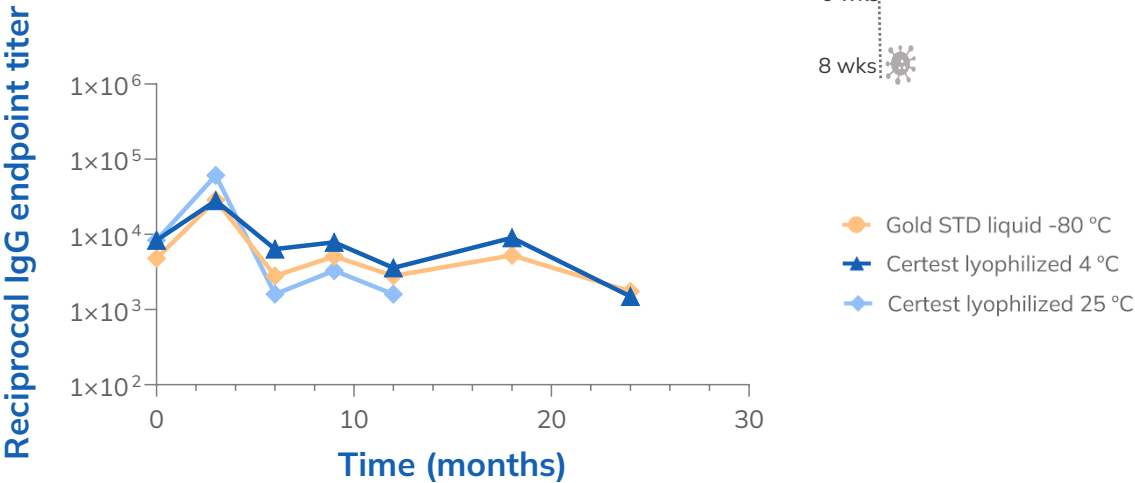
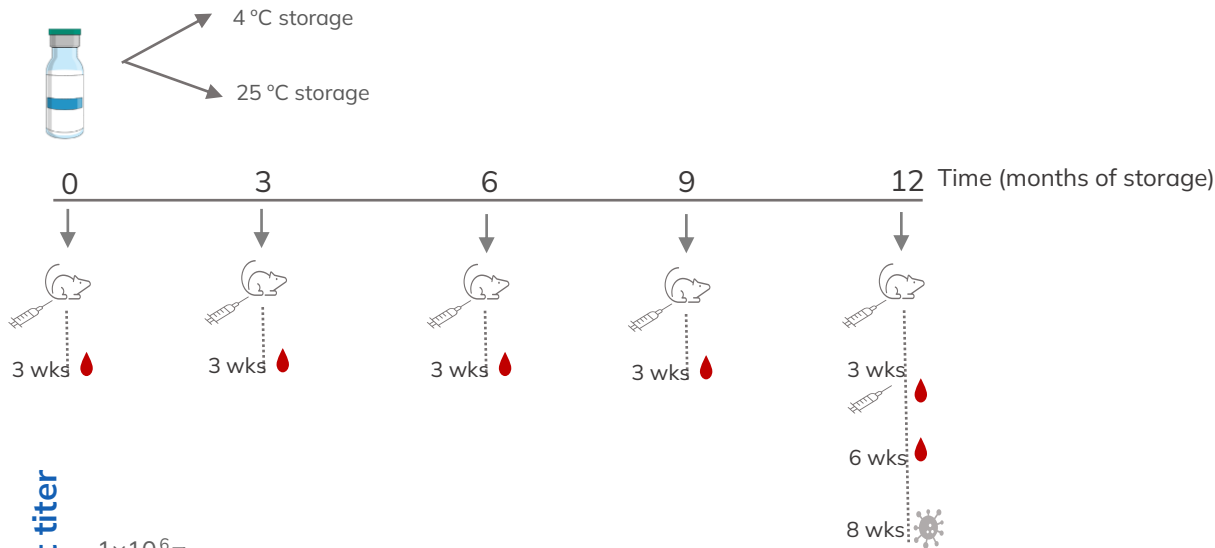
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Tolerability high dose (5 mg/kg)



Enhanced CRISPR/Cas gen editing



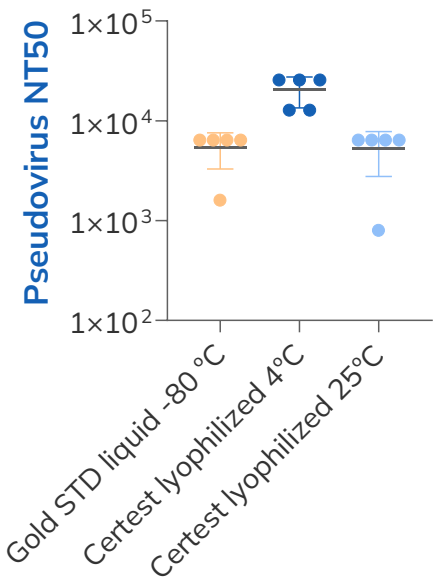


<https://doi.org/10.1038/s41541-025-01201-1>

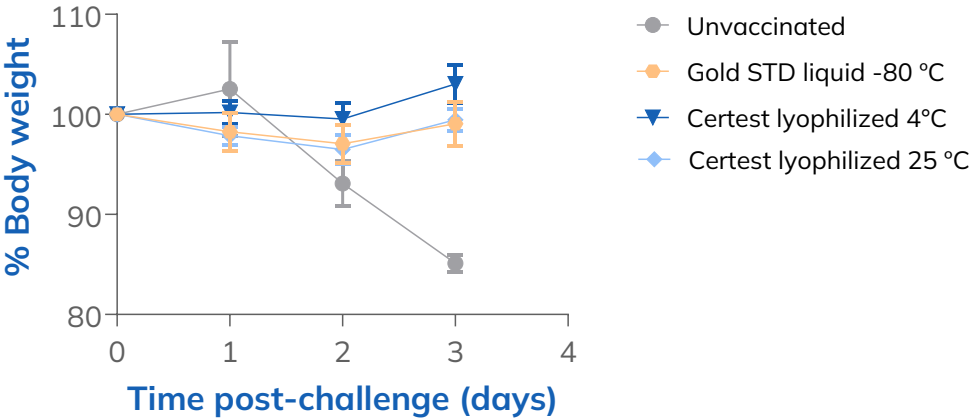
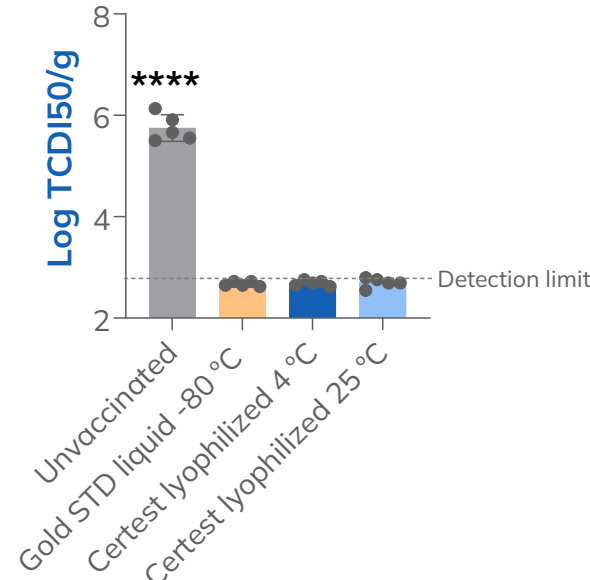


VACCINE EFFICACY AFTER 12 MONTHS OF STORAGE

Serum neutralization titers post-boost



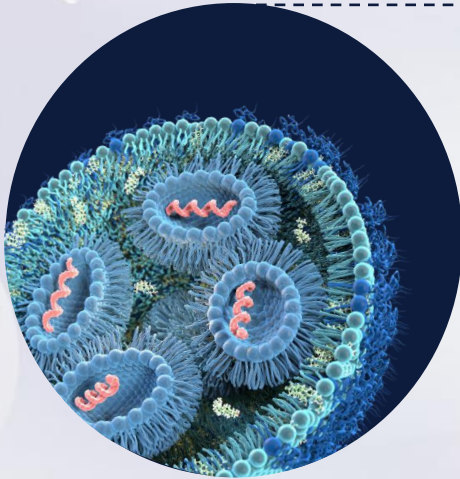
Lung viral load (day 3 post-SARS-CoV-2 challenge)



<https://doi.org/10.1038/s41541-025-01201-1>

Certest Lyophilized vaccines retain their immunogenicity and efficacy after 12 months of storage at temperatures up to 25°C

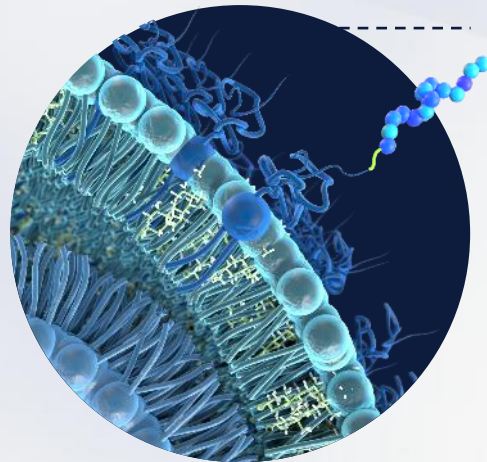
Passive Targeting



Four component LNP

- ★ *Organ-selective delivery through lipid composition*
- ★ *FTO (Freedom to Operate)*
- ★ *Protein corona-driven biodistribution modulation*

Active Targeting



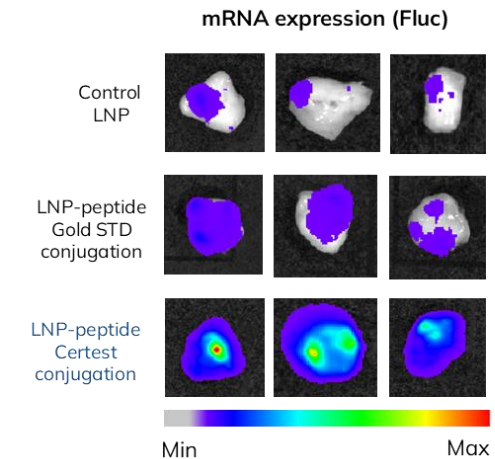
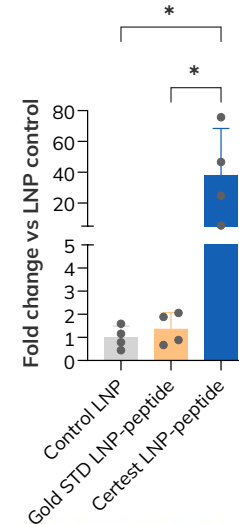
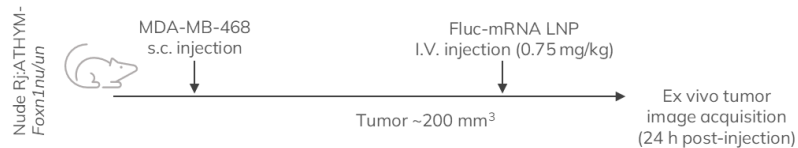
Ligand-conjugated LNPs with CerTarget Lipid Technology

- ★ *Improved conjugation and ligand presentation*
- ★ *Preserved LNP functionality and stability*
- ★ *20-fold in vivo increase in tumor delivery*
- ★ *Compatible with multiple ligands (peptides, AB, Fab)*

CerTarget Lipid



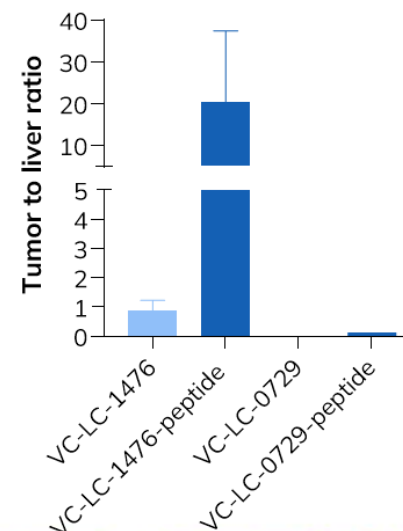
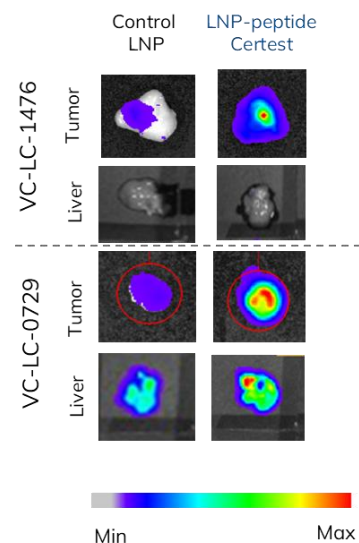
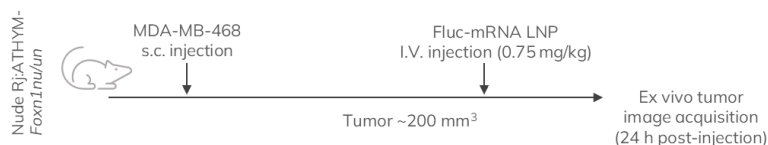
- PEG-alternative surface lipid platform
- Compatible with peptides and antibody conjugation
- Preserved LNP stability and functionality
- Improved receptor-mediated delivery performance
- **20-fold increase in tumor delivery in vivo**
- 3-fold increase in brain delivery in vivo.



CerTarget Lipid



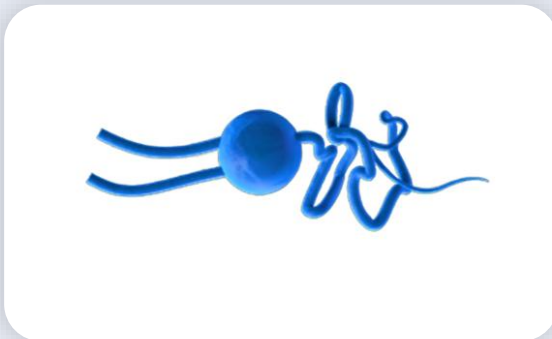
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Compatible with extrahepatic lipids



CerTarget Lipid

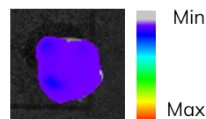


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DSPE-PEG-Peptide



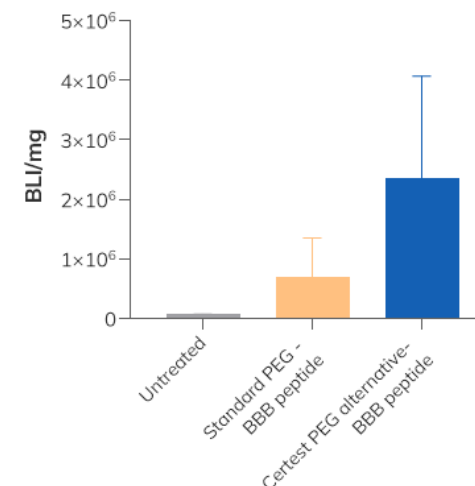
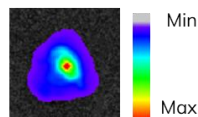
LNP-peptide
PEG
conjugation



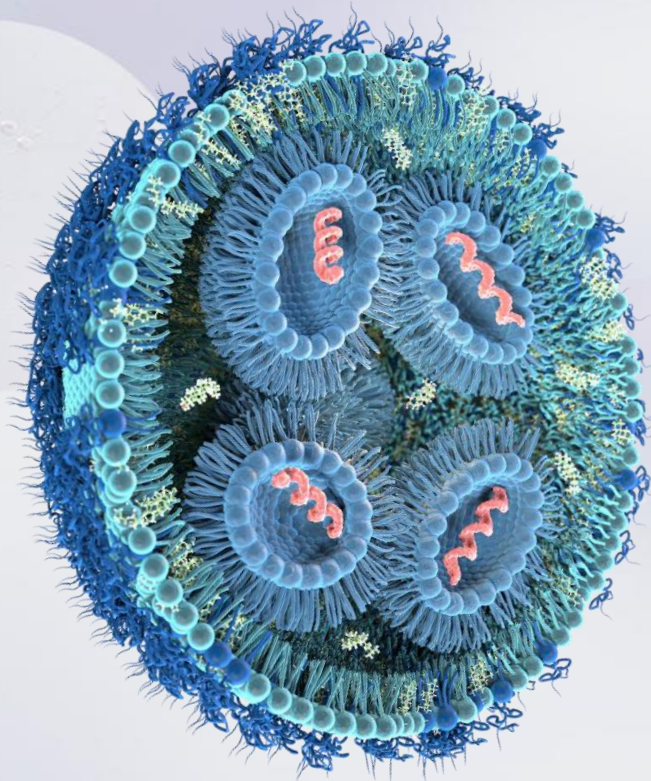
CerTarget lipid-Peptide



LNP-peptide
CerTarget
conjugation



Developing technologies the enhance LNP delivery performance



Optimized Thermostable LNP Formulations

Stability and efficacy of lyophilized LNPs and naked oligonucleotides, as well as liquid formulations

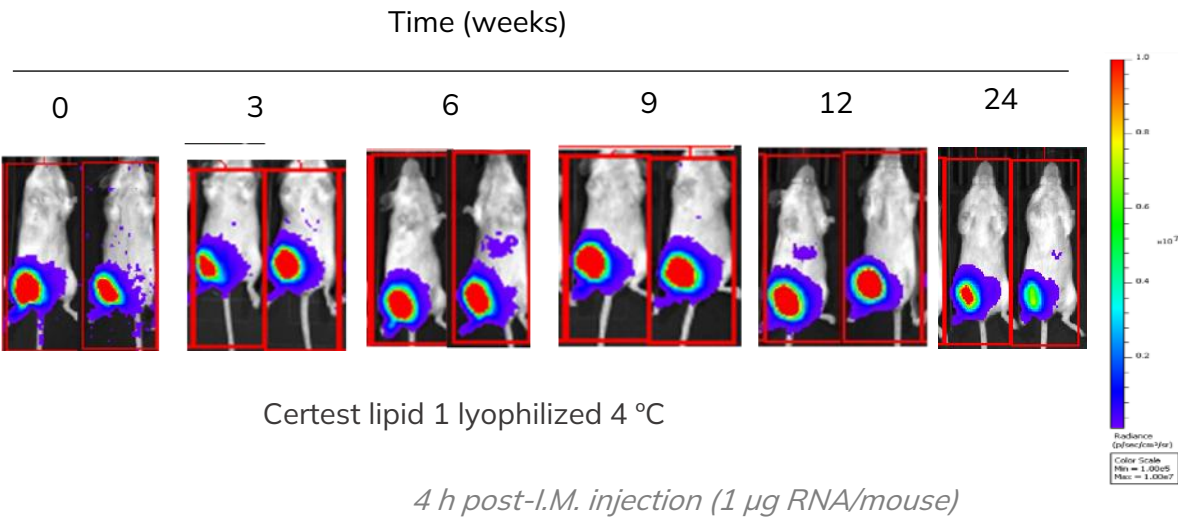
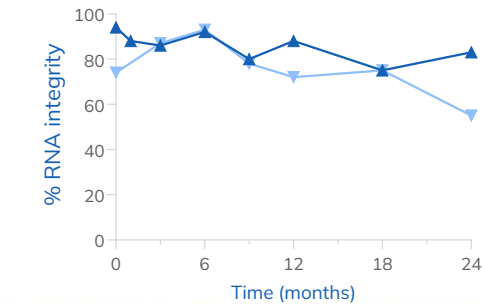
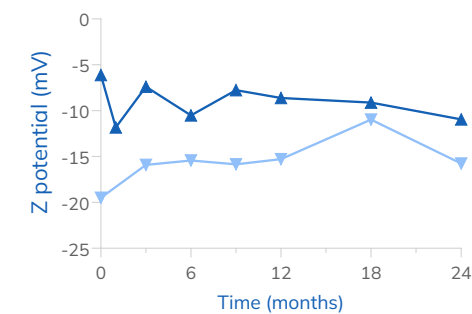
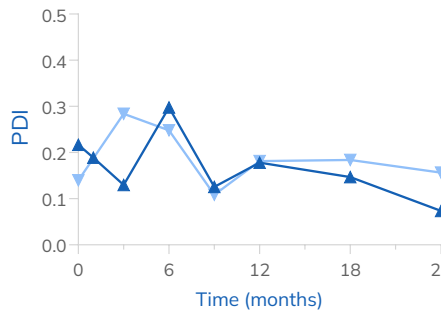
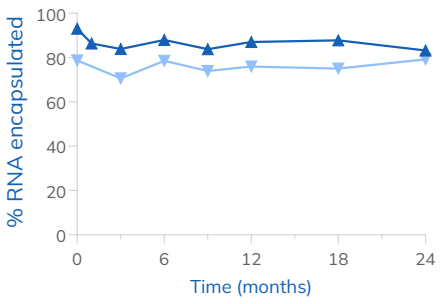
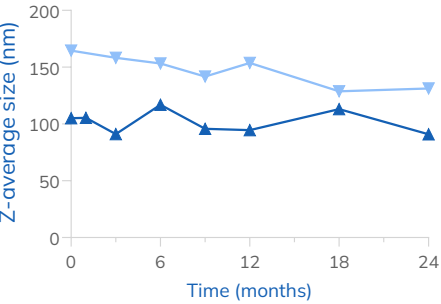


RNA and Oligonucleotide modifications

Tailored modifications to improve stability, specificity, and functionality

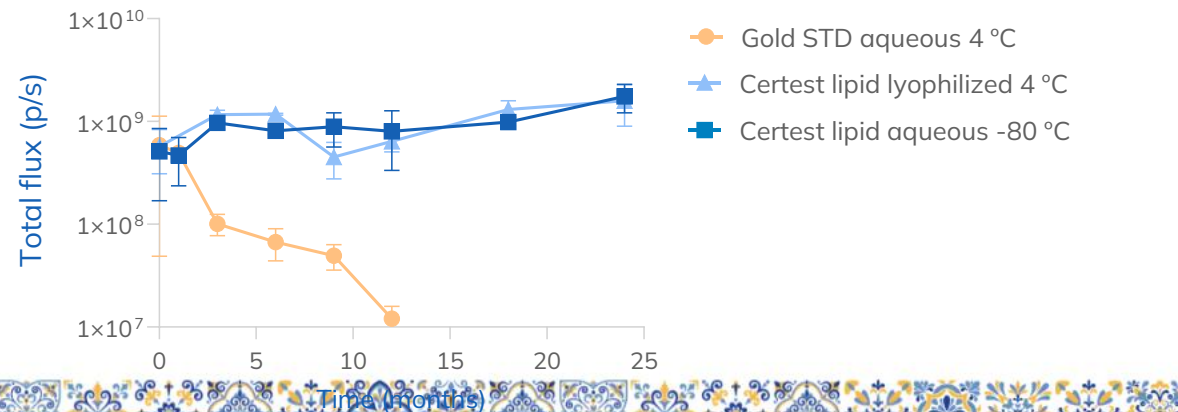


STABILITY AFTER 2 YEARS

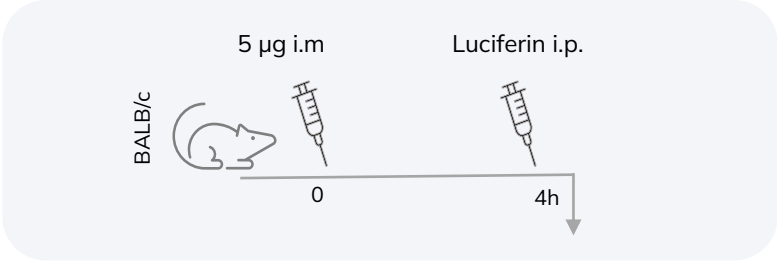
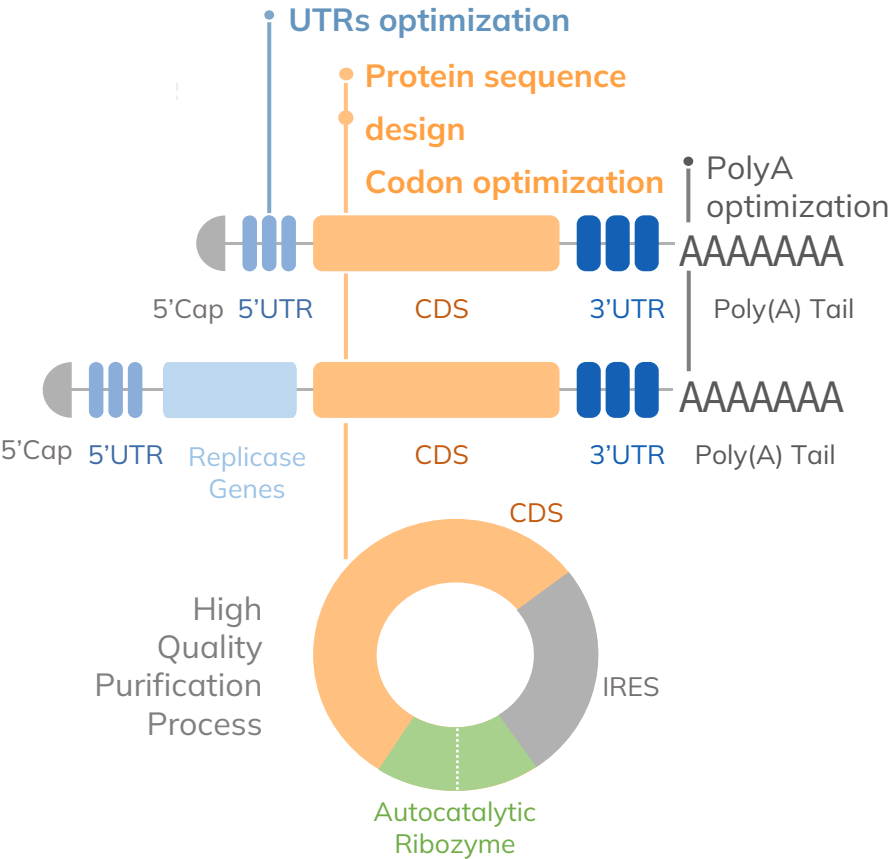


Certest lipid 1 lyophilized 4 °C

4 h post-I.M. injection (1 µg RNA/mouse)

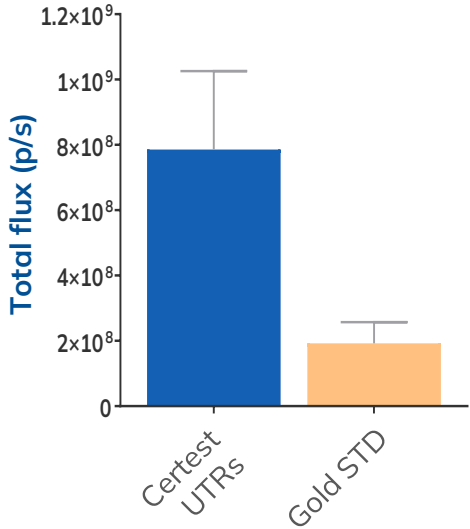


>> mRNA, saRNA and circRNA available



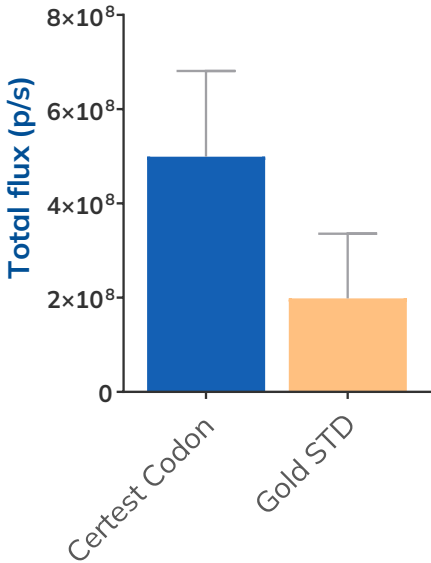
UTRs OPTIMIZATION

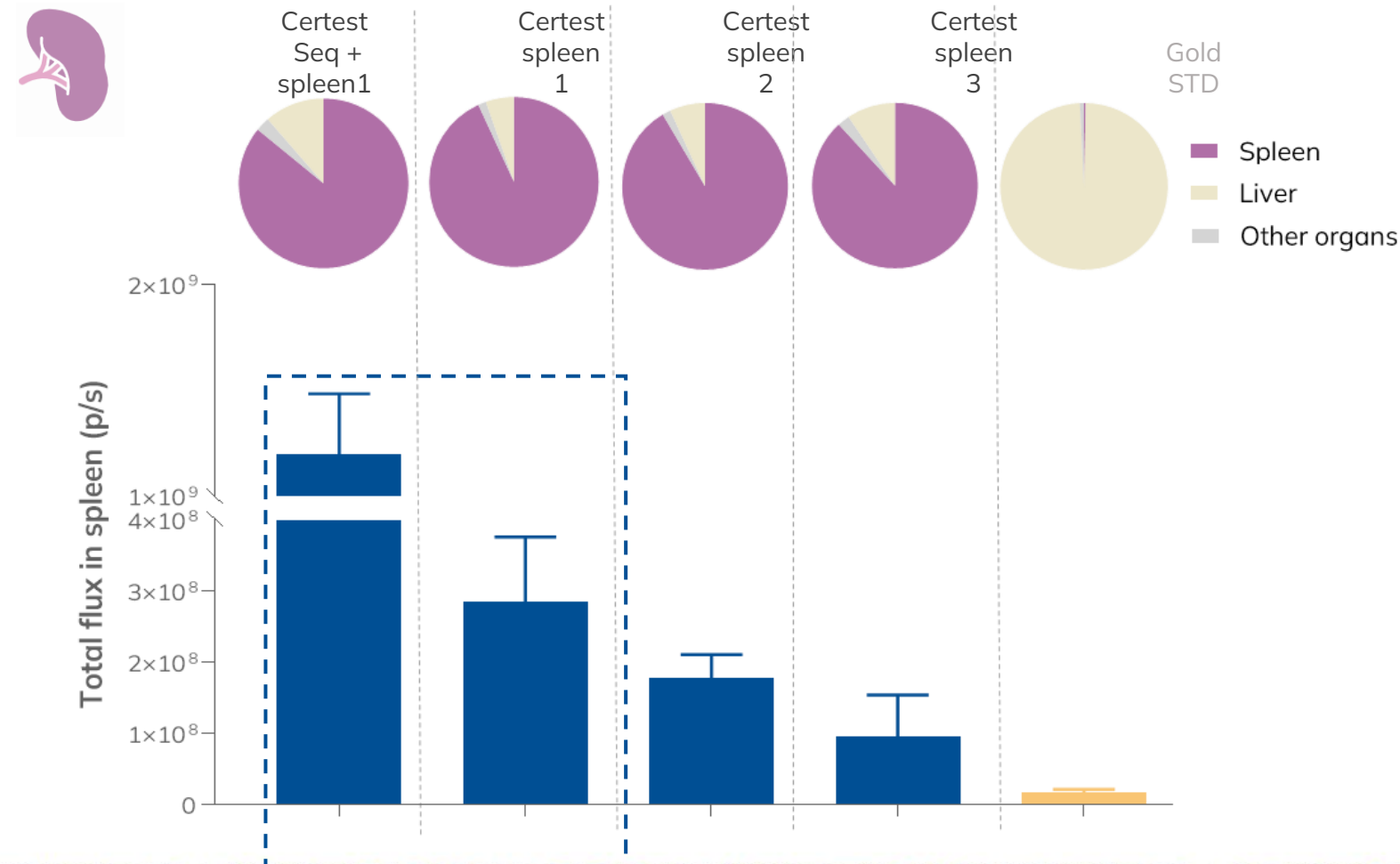
4x protein production



CODON OPTIMIZATION

3x protein production





Spleen specific UTRs increase protein expression for targeted





3,000
sqm of dedicated
labs



75+
people



Thank you – Questions?



<https://pharma.certest.es>



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pharma@certest.es