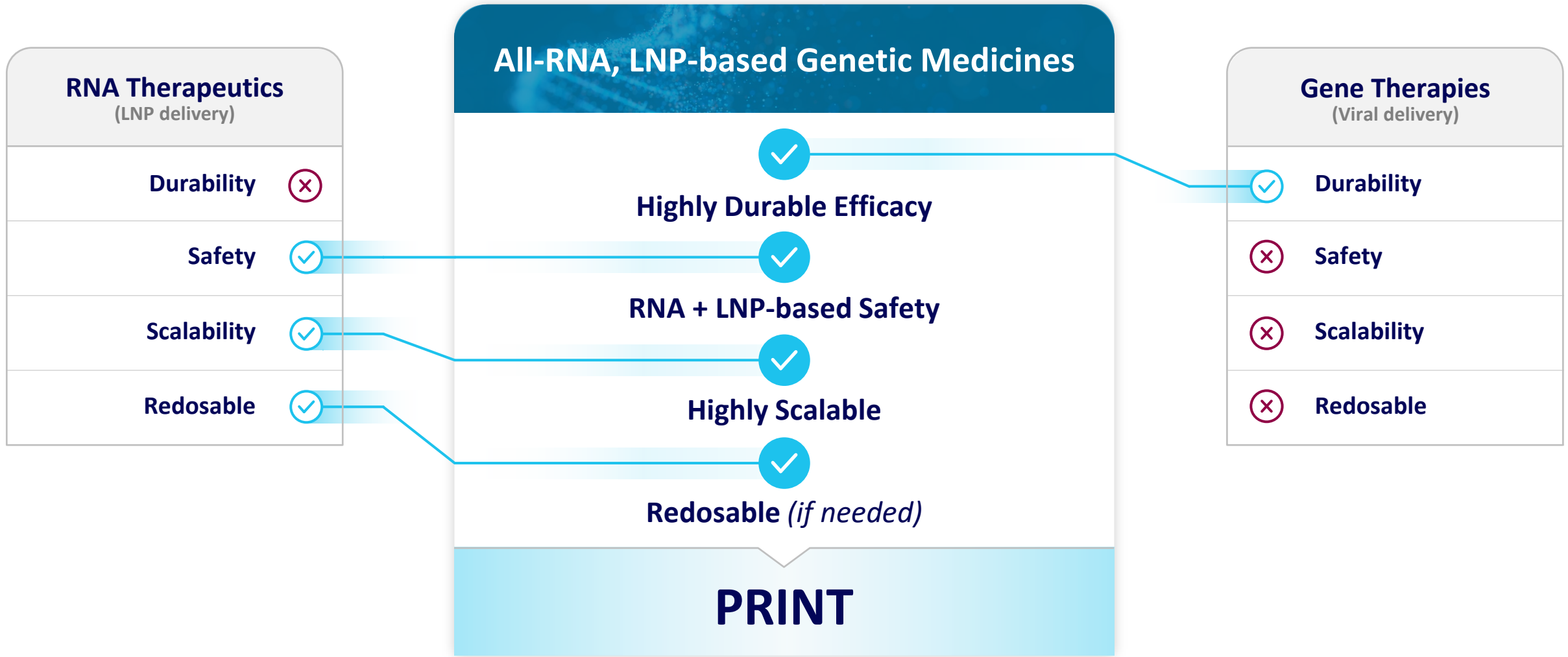


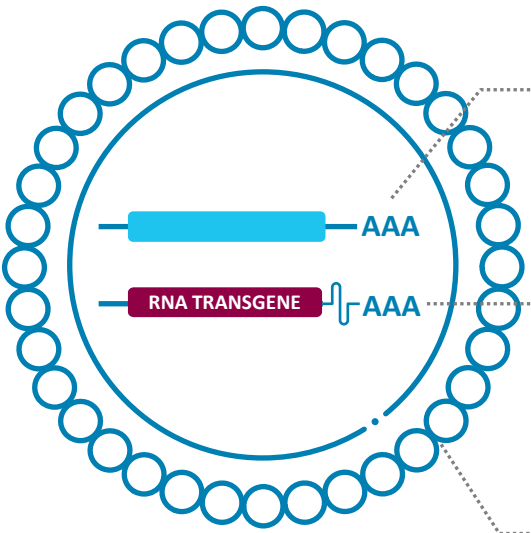
Turning hepatocytes into incretin factories with PRINT™, an RNA-based genetic medicine platform

PRINT: Combining the best of RNA therapeutics and viral gene therapies



Core of our platform is an all-RNA system using an R2 retrotransposase

Precise RNA-mediated Insertion of Transgenes



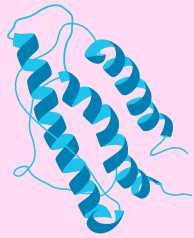
Universal Retrotransposase mRNA
Conventional mRNA

“Plug and Play” Template RNA
Proprietary sequences, Cargo agnostic

Lipid Nanoparticle
Low immunogenicity, repeat dose



Target:
Ribosomal safe harbor
~300-700 copies in human genome



Result:
Protein expression
Endogenous to human genome or Exogenous protein therapeutic

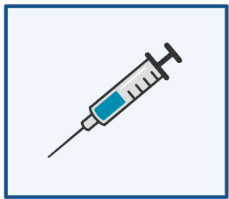
Abstract on PRINT platform (#103) was presented at ASGCT by Greg Cost on Tuesday, May 12th

PRINT is well-suited to address the challenges of severe, medically-driven obesity

Severe, morbid obesity is a lifelong medical condition that cannot be managed through lifestyle changes alone.

Reduces life expectancy by 5-15 years

Current therapies:



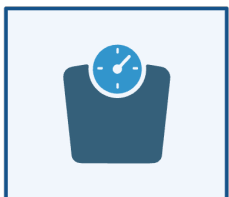
Treatment burden:

Daily or weekly administrations



Side effects burden:

Cardiac, hyper-pigmentation, nausea, ...

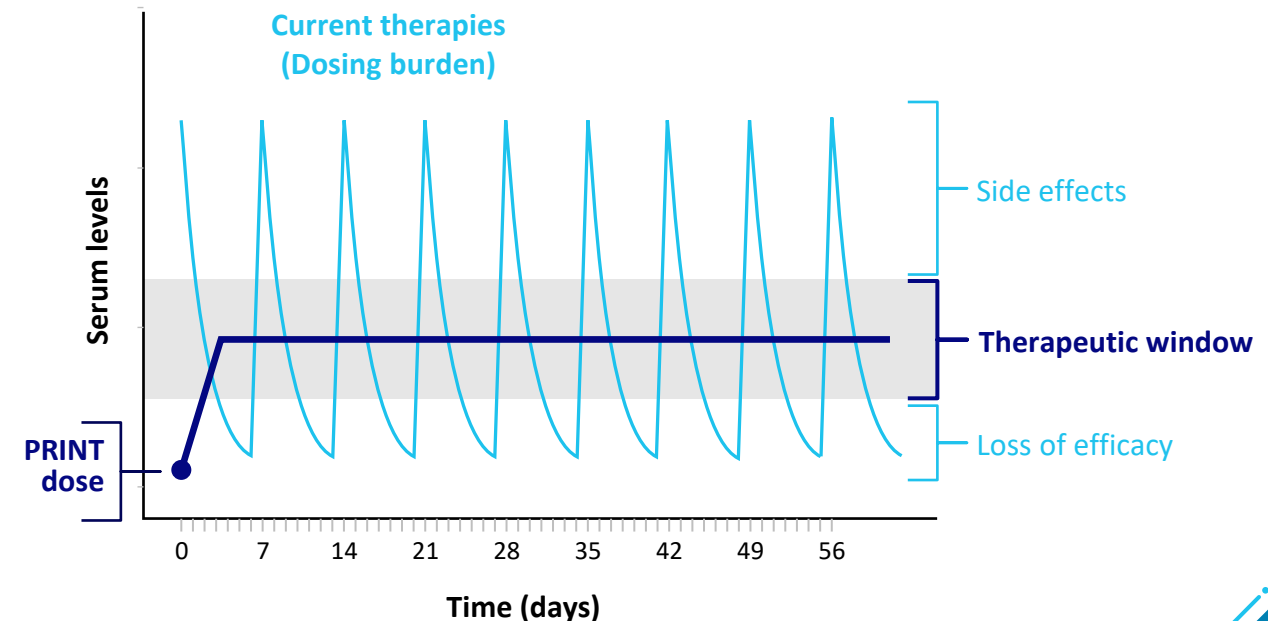


Long term outcome:

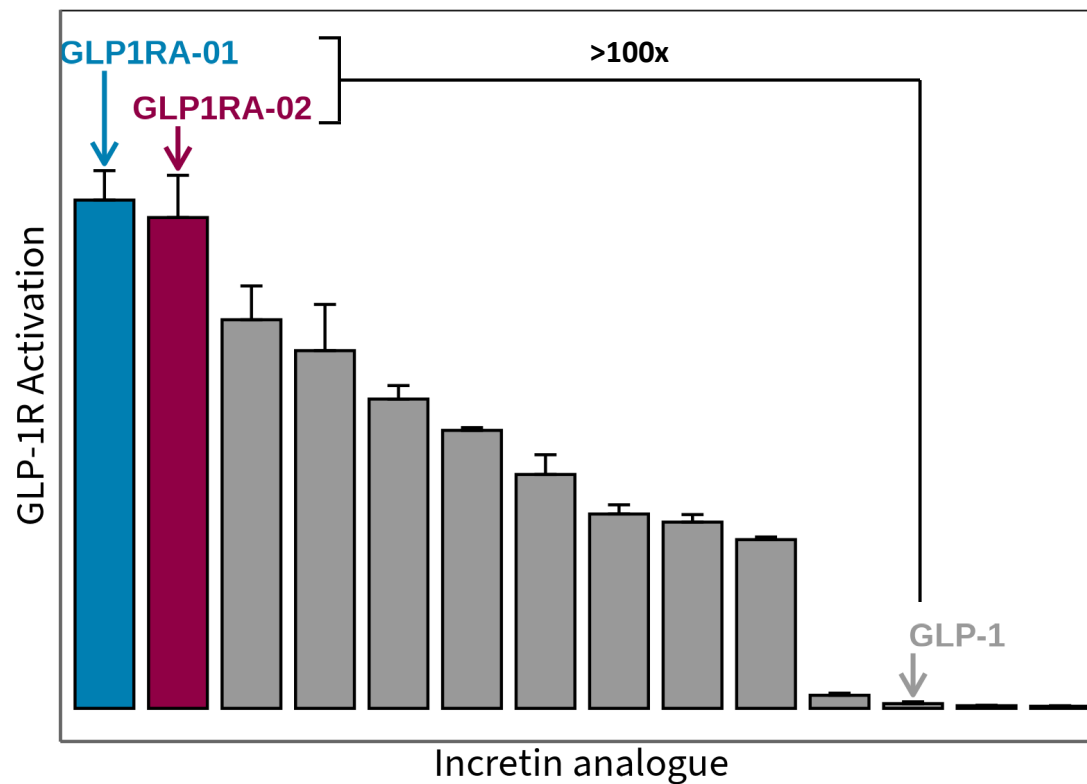
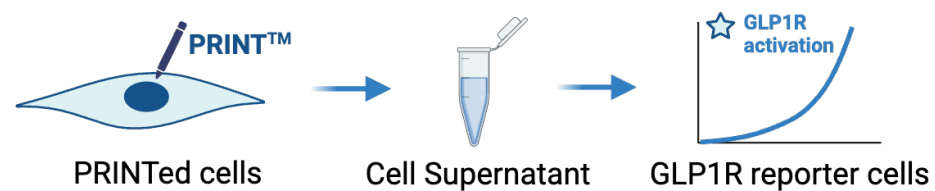
Contingent on compliance

PRINT potential:

- **Multi-year durability** eliminates dosing burden
- **Flat PK** improves efficacy and tolerance
- **Redosability and tunability** control therapeutic exposure



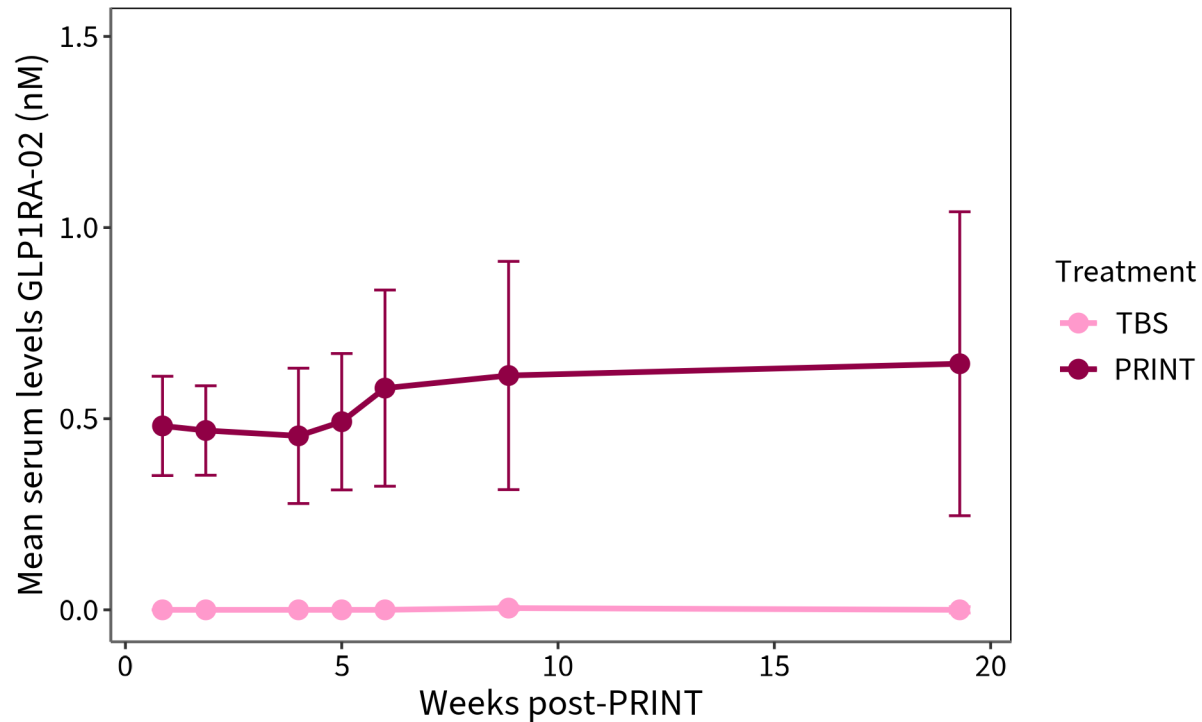
PRINTed incretin mimetics show high potency profile



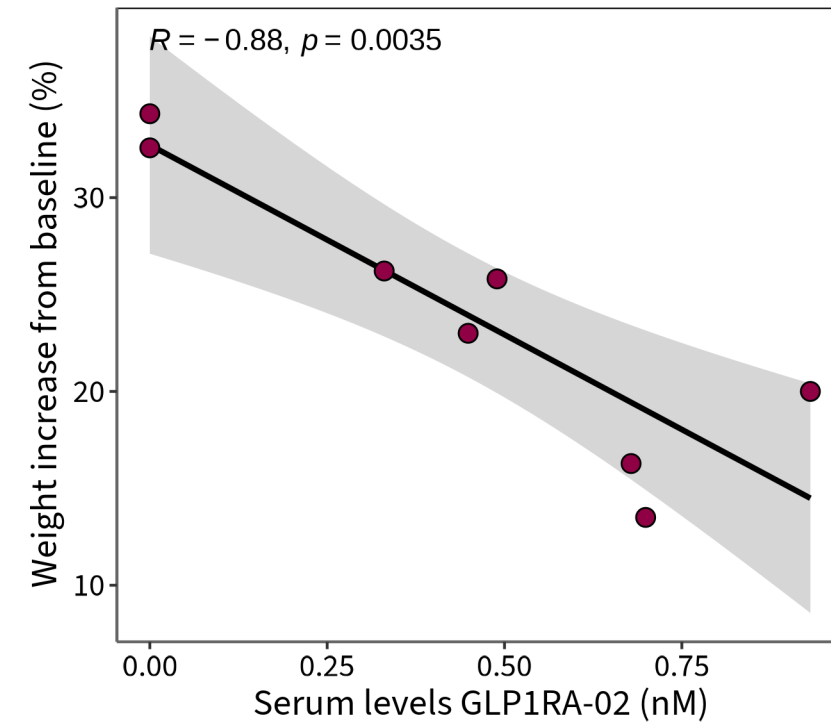
Sustained expression of an incretin mimetic correlates with reduced weight gain in a DIO model

GLP1RA-02 (first generation template)

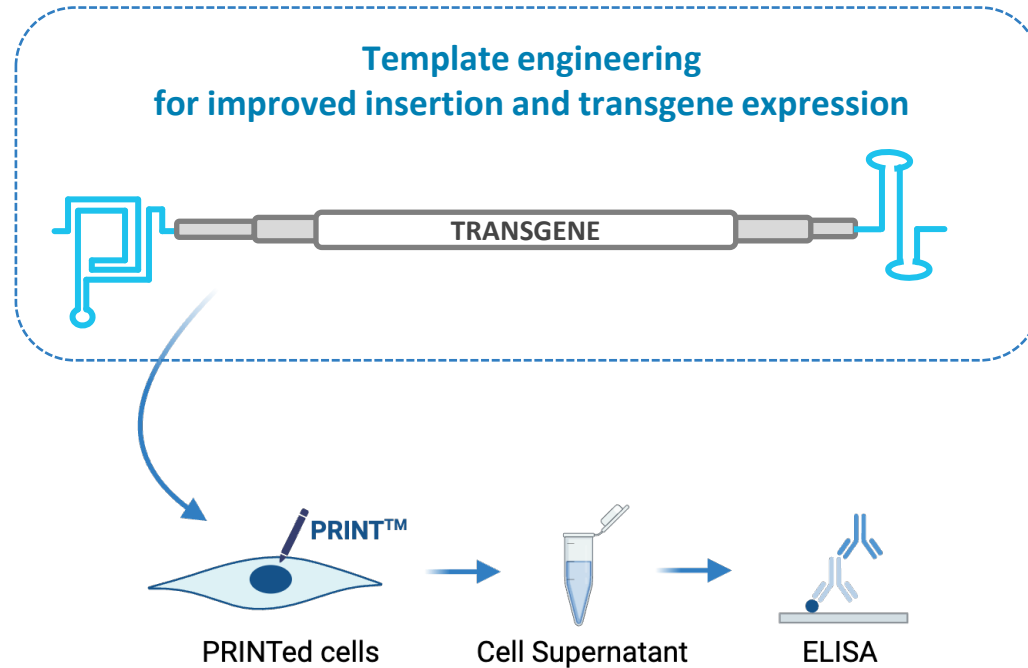
GLP1RA-02 serum levels
(up to 19 weeks post-PRINT)



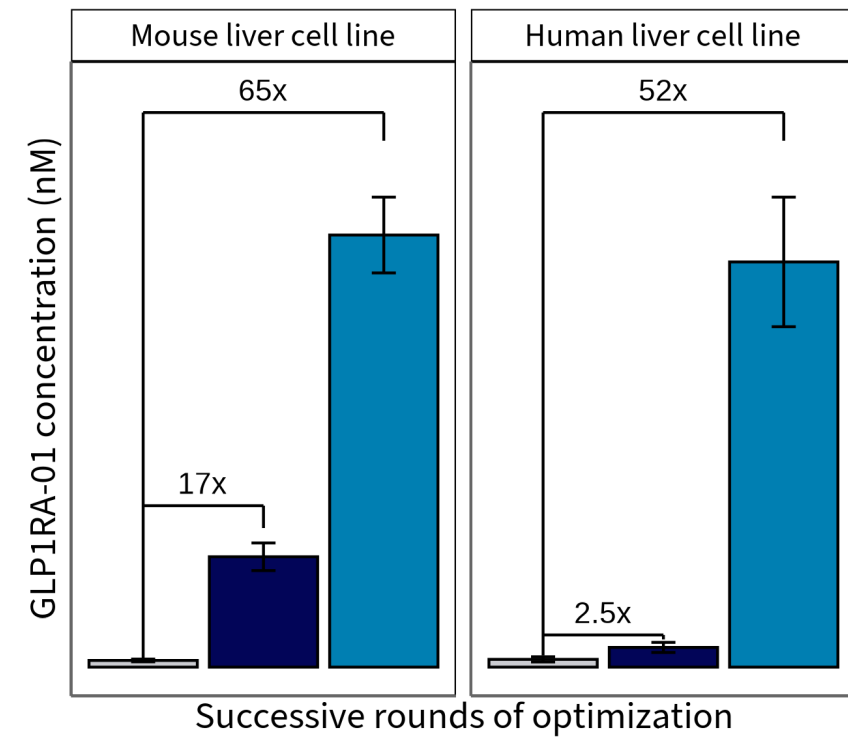
GLP1RA-02 effect on weight gain



Engineering of templates achieves enhanced PRINT of incretin mimetics in cell lines



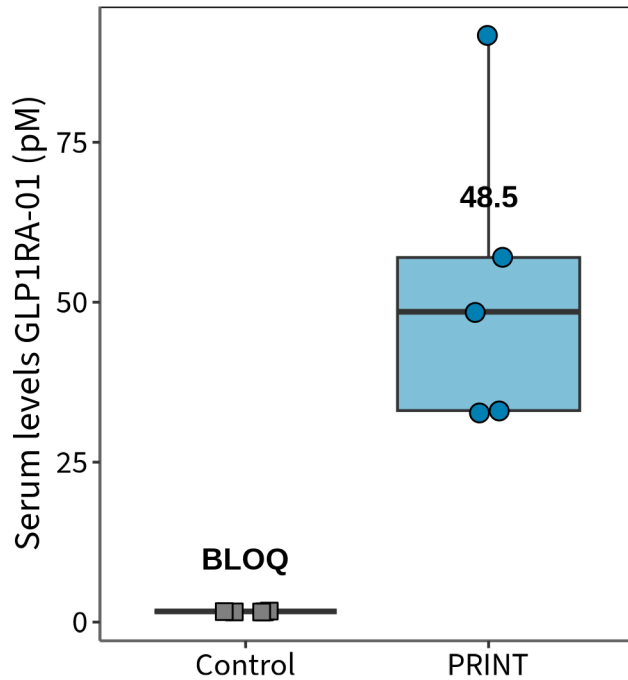
GLP1RA-01 template optimizations



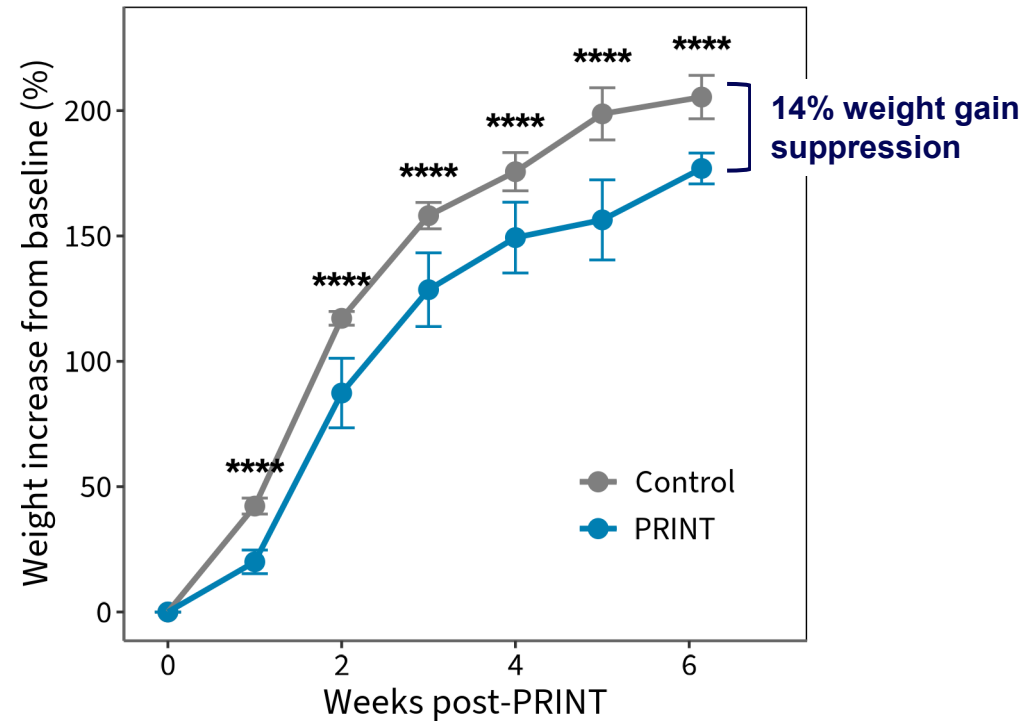
Efficient PRINT of incretin mimetics results in reduced weight gain in mice on standard diet

GLP1RA-01

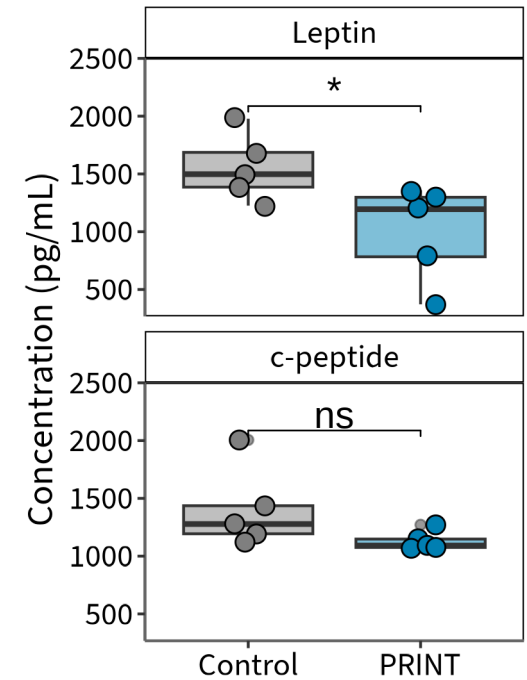
Serum levels (6 weeks post-PRINT)



Weight gain suppression



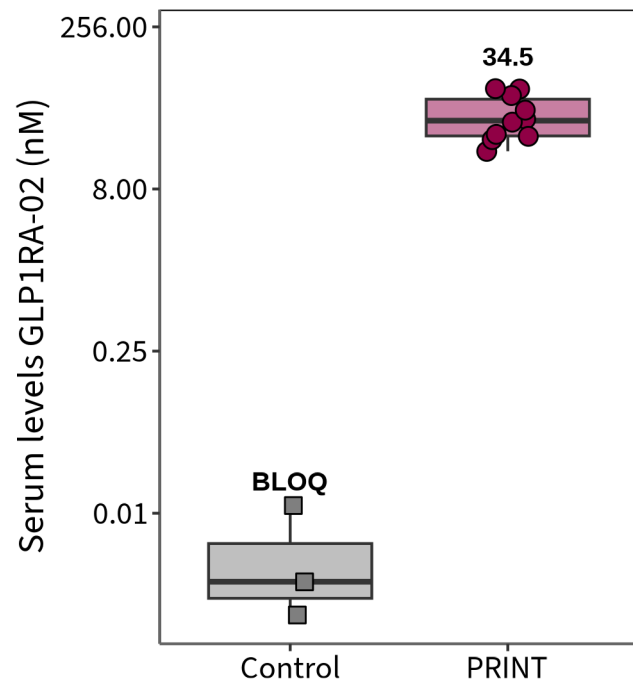
Non-fasting biomarker levels



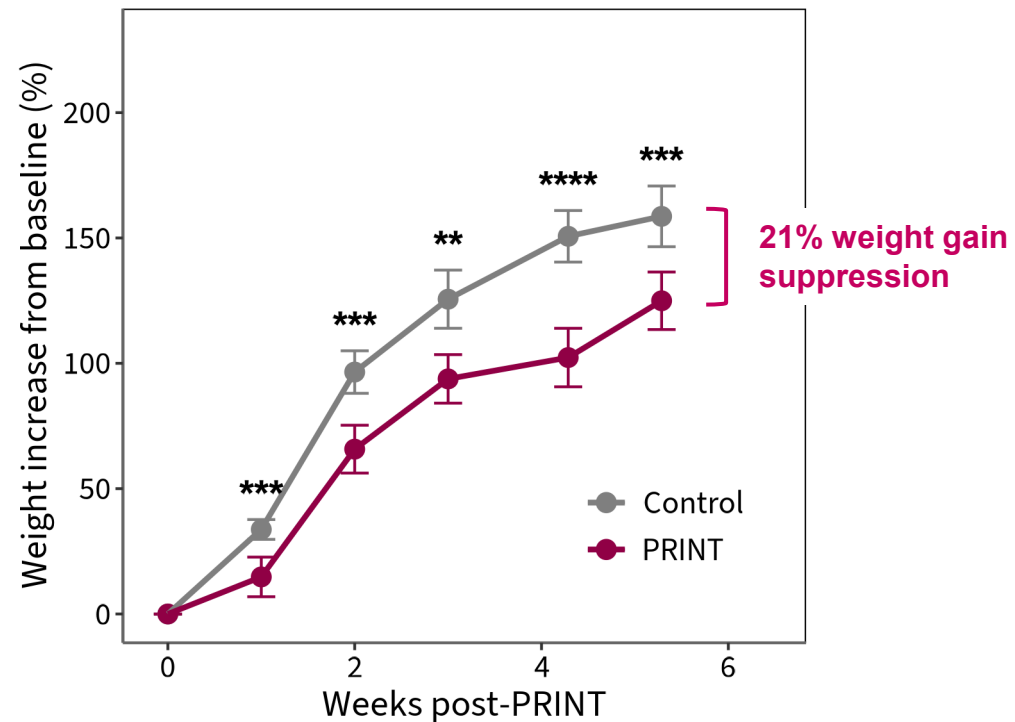
Efficient PRINT of incretin mimetics results in reduced weight gain in mice on standard diet

GLP1RA-02

Serum levels (3 weeks post-PRINT)



Weight gain suppression



Switch allows for dose-dependent, permanent PRINT turn-off with an FDA-approved molecule



Dimmer switch for tunable reversibility

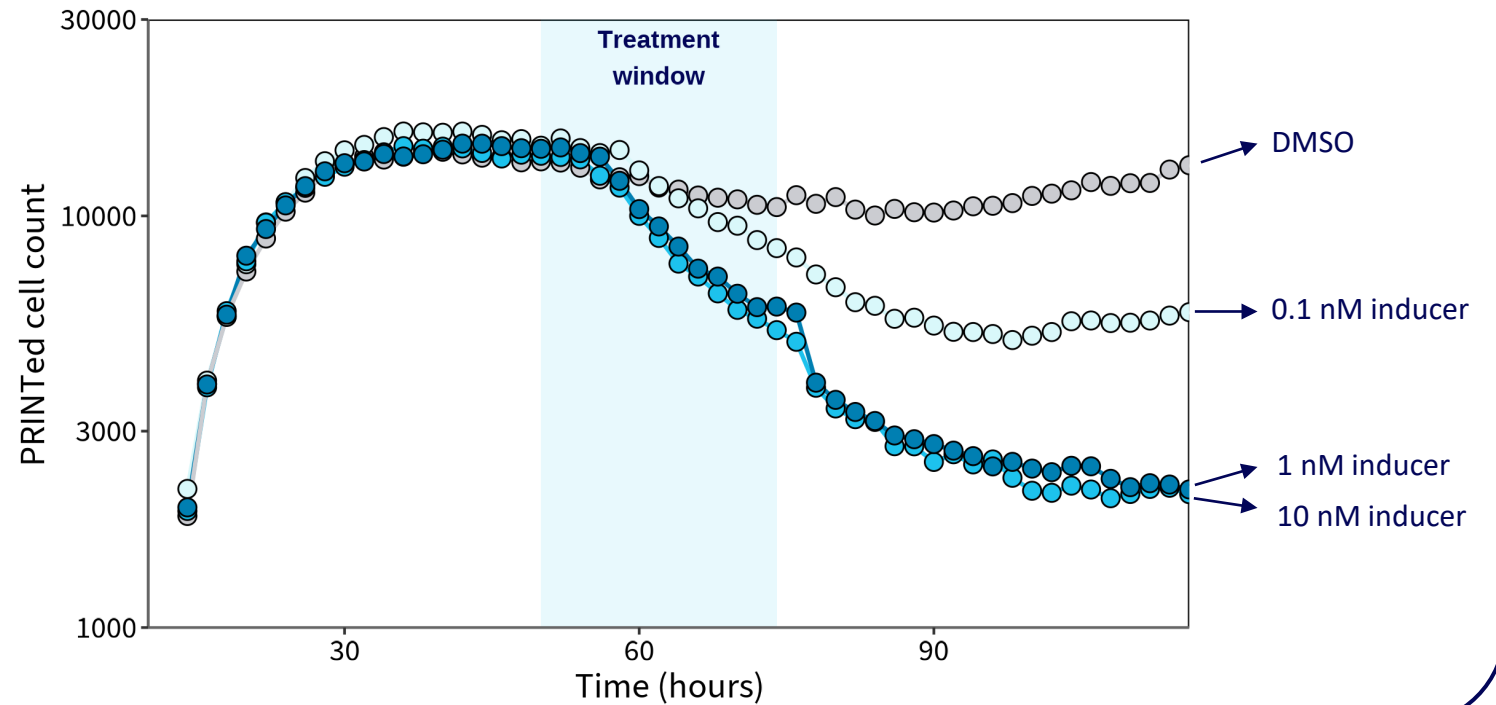


Failsafe switch for complete termination

Inducible apoptosis switch

- Clinical validation and long development history
- Low immunogenicity and genotoxicity
- Short time for activation

In vitro proof-of-concept of PRINT safety switch with fluorescent reporter



PRINT: a path to a durable, tunable therapy for severe, chronic obesity

PRINT

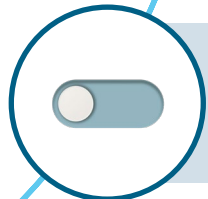
PRINT combines the **tolerability and redosability of RNA therapeutics** with the **durability of gene therapy**, enabling sustained transgene expression from a single, repeatable dose.



Template engineering campaigns successfully **raise expression of incretin mimetics with strong potency profiles**.



PRINTed incretin mimetics result in up to **21% lower weight gain** in mice on standard diet and **exposure correlates with weight gain suppression in a DIO model**.

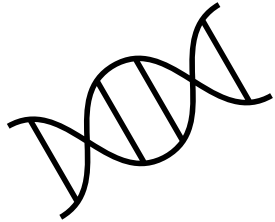


A built-in, clinically validated safety switch enables both **fine-tuning and full reversal** of therapy.

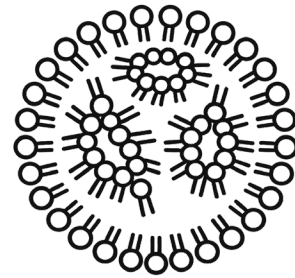
Acknowledgements – it takes a village!



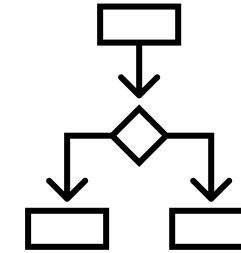
ADDITION THERAPEUTICS



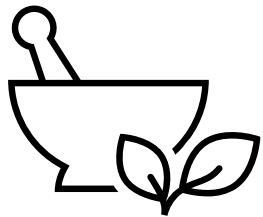
Therapeutic Technologies



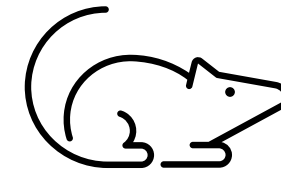
LNP Formulation



Operations



Pharmaceutical Sciences



Pre-clinical Translation

Thank you