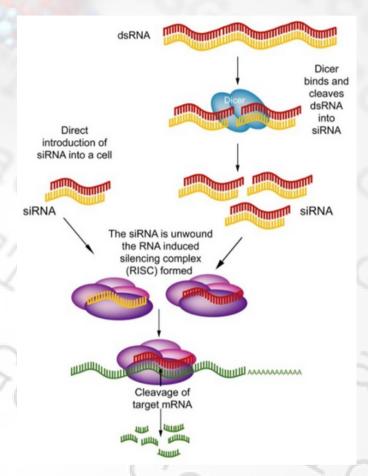


Provider of Preclinical Research Services (GLP/non-GLP) for Drug Discovery Efficacy and Pharm/Tox IND contract research studies (clients worldwide) 100+ Xenograft Models (validated in-house) and IND-enabling Toxicology studies 100% IP belongs to client, experienced IACUC-regulated barrier facility

# Targeted siRNA qPCR Assays

## RNA Interference and qPCR Assays

- 1. Introduction of dsRNA into the cell triggers cleavage by Dicer to yield siRNAs
- 2. RISC binds to, processes, incorporates siRNA
- 3. RISC uses siRNA to seek out complementary sequences and degraded them silencing gene expression



http://zimdarsgen564s14.weebly.com/rna-interference.html



## RNA Interference and Drug Discovery

- Specific, targeted gene silencing
- Expression is significantly reduced, but not eliminated (gene knockout)
- Represents an exploding field of Drug Discovery
  - 1. Identifying drug targets
  - 2. Elucidating cellular signaling pathways
- Applications with novel drug discovery but also RNA-based drugs



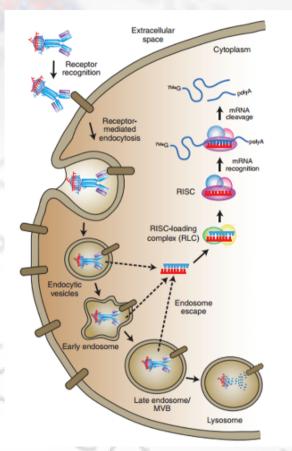
## Challenges of RNA Interference

#### Delivery:

- Exogenous dsRNA gaining entry to the cell
- Escaping from the endosome

#### Stability of dsRNA:

 Avoiding degradation to induce RNAi silencing



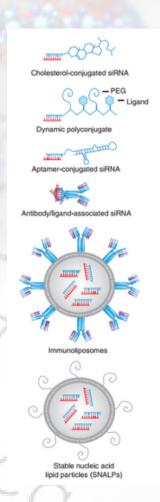
Celluar delivery of siRNA

Monika Dominska, Derek M. Dykxhoorn J Cell Sci 2010 123: 1183-1189



## Efficient RNA Delivery

- Altogen Labs is experienced in efficient delivery of RNA to produce targeted gene silencing via the RNA interference pathway (liposomal encapsulation and chemical modification RNA)
- Altogen Labs has extensive knowledge and expertise of encapsulation and modification for efficient delivery of RNA



Different RNA modification/encap sulation methods for enhancing delivery

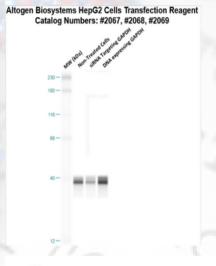
Monika D., Dykxhoorn, D.M. 3 Cell Sci 2010 123: 1183-1189



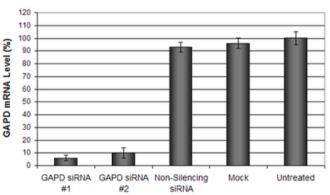


#### Representative In Vitro RNAi Knockdown

- siRNA is delivered to target cell line via cationic transfection reagent
- Knockdown efficiency is evaluated by RT-PCR (mRNA) and Western Blot (mRNA)



HepG2 cells transfected with siRNA targeting Glyceraldehyde-3-Phosphate Dehydrogenase



Post-transfection data displaying targeted knockdown of GAPD mRNA (RT-PCR, top) and protein expression (right)

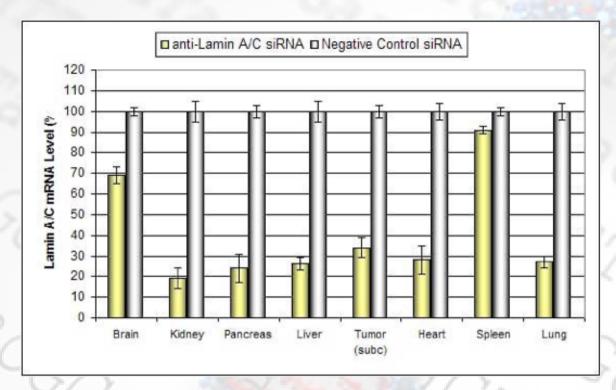


## Representative In Vivo RNAi Knockdown

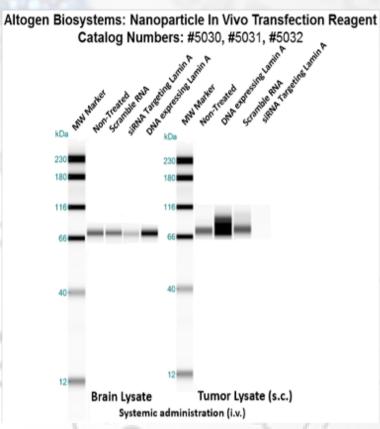
- siRNA is introduced via nanoparticlebased reagent intravenously (IV) or intratumorally (IT)
- Knockdown efficiency is evaluated by RT-PCR (mRNA) and Western Blot (mRNA) on tissue and tumor lysates



#### Systematic administration of anti-Lamin A/C siRNA



Post-intravenous injection data displaying targeted knockdown of Lamin A/C (RT-PCR, top) and protein expression (right)





#### Contact Us

#### Altogen Labs RNAi Services:

- Targeted gene silencing via siRNA
- Analytical methods to characterize reduction in mRNA concentration (RT-PCR) and concurrent knockdown in protein expression (Western Blot)
- Cell-based assays
- In vitro and in vivo RNA stability studies
- RNA chemical modification and encapsulation



Contact us to discuss details, timeline estimates, and price!

