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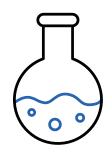
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This presentation concerns drug candidates, some of which are undergoing nonclinical studies and others of which are under clinical investigation, and all of which have not yet been approved for marketing by the U.S. Food and Drug Administration. These drug candidates are currently limited by federal law to investigational use, and no representation is made as to their safety or effectiveness for the purposes for which they are being investigated.

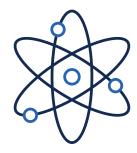


# Aligos Therapeutics Highlights



**Our Mission** 

Develop novel therapeutics to address unmet medical needs in viral and liver diseases



**Our Platforms** 

Proprietary oligonucleotide and small molecule platforms

Leverage complementary modalities to broaden the range of therapeutic targets we can address



Our Strategy

Develop pharmacologically optimized drug candidates against clinically validated targets

Pursue combination regimens designed to achieve improved treatment outcomes



**Our Team** 

Decades of drug discovery and development experience

Proven track record of success in the areas of viral infections and liver diseases

Our goal is to become a world leader in the development of targeted, antiviral therapies for CHB and COVID-19 and to leverage our expertise in liver diseases to create targeted therapeutics for NASH



# Our Platforms, Capabilities and Collaborations



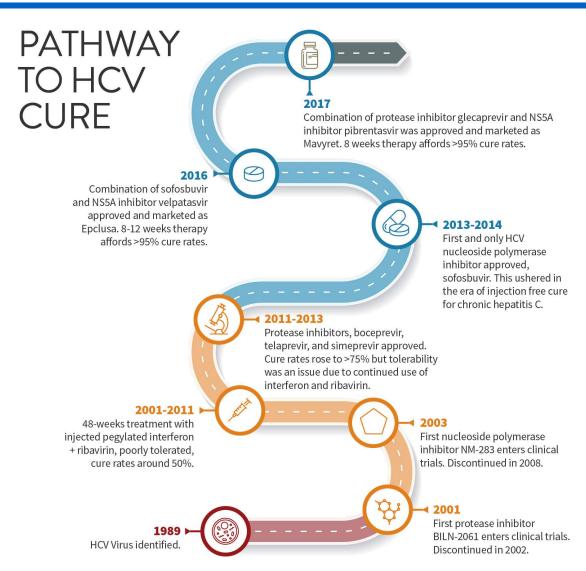
Enabling efficient drug discovery in pursuit of optimized combination regimens



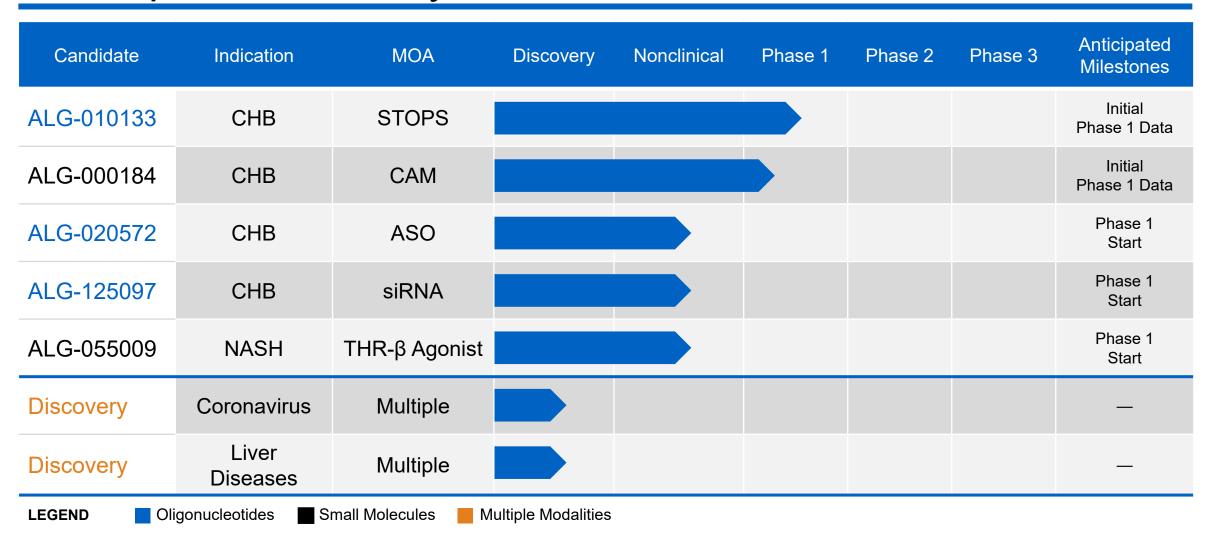
# A Lesson from Chronic Hepatitis C From First Movers to Best-in-Class Regimens

- Efficacy → Safety → Convenience
- Standard of care treatment until 2011
  - Pegylated interferon + ribavirin
  - 48-weeks, ~50% cure rate, poorly tolerated
- Protease inhibitors
  - First clinical entrant BILN-2061 (2001)
  - First approvals were boceprevir and telaprevir (2011)
  - Simeprevir was third (2013)
- Nucleos(t)ide analogs
  - First clinical entrant NM283 (2003)
    - Followed by balapiravir (2004) and IDX-184 (2010)
    - Only approved compound, sofosbuvir (2013)
- Current standard of care includes NS5A inhibitors
  - Cure in 8-12 weeks. >95% cure rate
  - Epclusa: sofosbuvir + velpatasvir (Gilead, 2016)
  - Mavyret: glecaprevir + pibrentasvir (AbbVie, 2017)





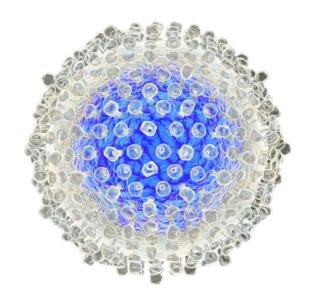
# Our Pipeline of Wholly Owned Assets







# Chronic Hepatitis B



# CHB Epidemiology & Our Portfolio

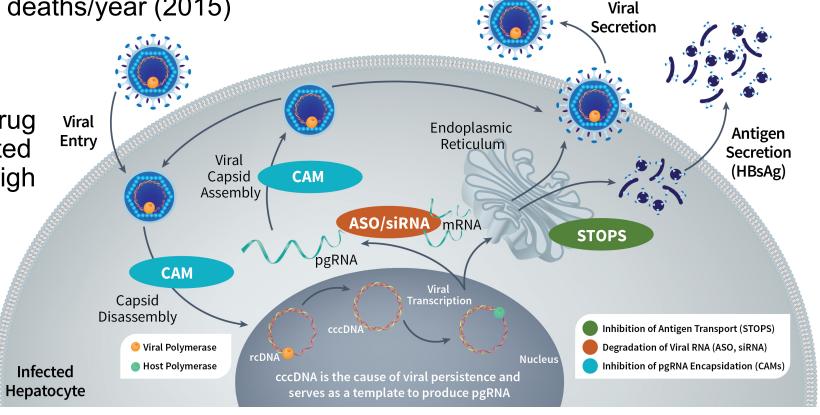
- CHB Epidemiology\* Most common chronic viral infection in world
  - >290 million carriers worldwide (2020)
  - ~8 million in US, EU, Japan (2015)

- Responsible for ~900,000 deaths/year (2015)

#### Our Portfolio

 Potentially best-in-class drug candidates against validated targets with potential for high functional cure rates

- > STOPS<sup>TM</sup>
- > CAM
- > Oligonucleotides
  - ASO
  - siRNA

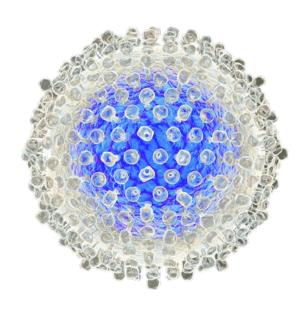






# **STOPS**

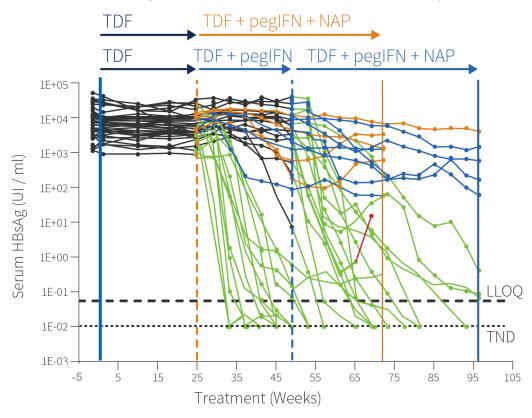
ALG-010133, a novel, chemically optimized poly-AC oligonucleotide analog



# Poly AC Oligonucleotides Have Clinical Validation

- Replicor 102 Study\*
  - REP 2139-Ca Monotherapy
  - 500 mg Q-weekly, 2-hr IV administration
  - Significant reduction in HBsAg observed
- Replicor 401 Study (Figure\*\*)
  - REP 2139-Mg or REP 2165-Mg
  - 250 mg Q-weekly, 2-hr IV administration
  - Triple combination
    - With tenofovir and pegylated-IFN
  - Reported a ~39% functional cure rate\*\*\*

#### **HBsAg Reduction in REP 401 Study**

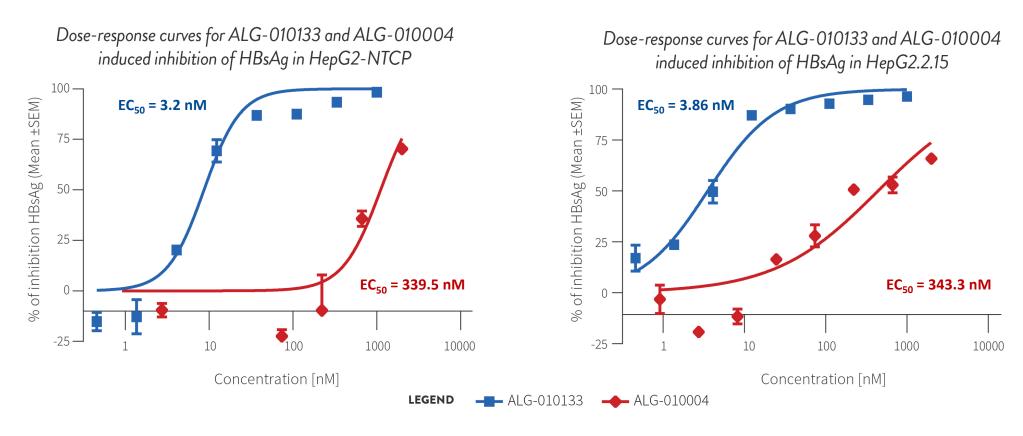


Treatment results in multiple log<sub>10</sub> reductions of HBsAg and higher rates of functional cure



# **ALG-010133**Our Lead STOPS Molecule

#### Significantly greater potency vs. reference Poly-AC oligonucleotide\*



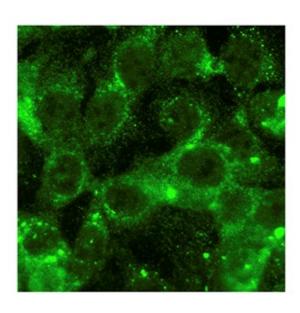
Similar potency improvement also seen in primary human hepatic cells





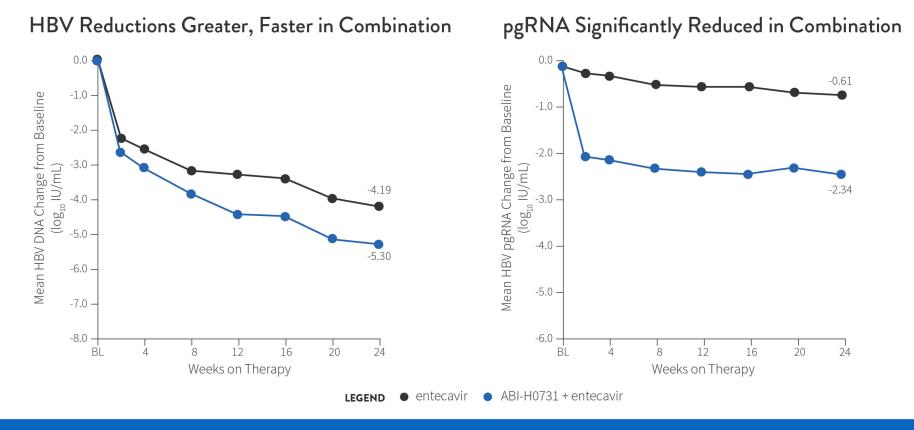
# **CAM**

ALG-000184, a prodrug of ALG-001075



# CAMs Have Demonstrated Clinical Anti-HBV Activity

Assembly ABI-H0731 at 300 mg QD in HBeAg (+) treatment naïve CHB patients in combination with the nucleoside analog entecavir



Unlike nucleoside analogues when used in monotherapy, reductions in HBV pgRNA were also observed



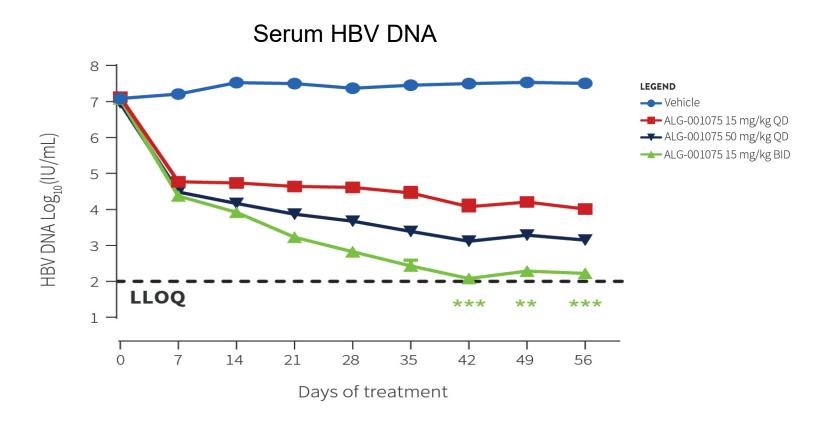
# Our CAM Demonstrates Superior In Vitro Potency vs. Other Known CAMs in Clinical Development

Compound	Current Status	HBV DNA reduction (EC <sub>50</sub> nM)	Cell Type
Assembly ABI-H0731	Phase 2	172	AD38
Assembly ABI-H2158	Phase 2	22	AD38
Assembly ABI-H3733	Phase 1	5	AD38
Janssen JNJ-6379	Phase 2	54	HepG2.117
Janssen JNJ-0440	Completed Phase 1	12	HepG2.117
Enanta EDP-514	Phase 1	17	HepG2.2.15
Aligos ALG-000184	Phase 1	0.63	HepG2.117
		0.53	HepG2.2.15

With the exception of ALG-000184, data was sourced from publicly available literature, posters and presentations. ALG-000184 data was generated by Aligos on the parent compound ALG-001075.



## Potent In Vivo Activity in the AAV-HBV Mouse Model



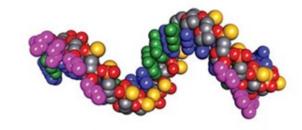
Dose dependent reductions in HBV DNA observed across treatment groups >5 log<sub>10</sub> IU/mL reduction in HBV DNA relative to placebo in the 15 mg/kg BID treatment group



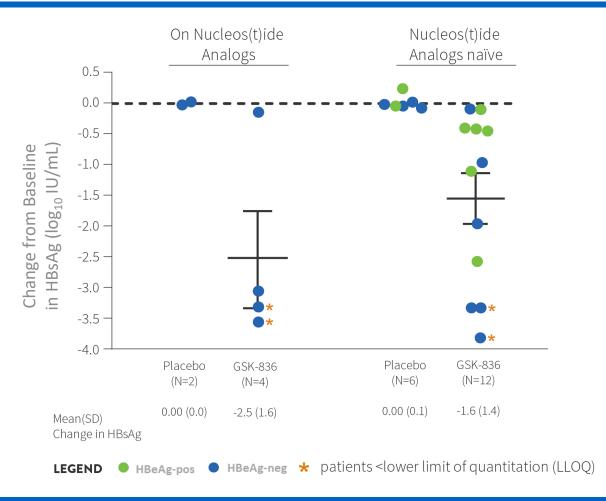


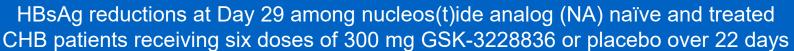
# Oligonucleotide Approaches

ASO (ALG-020572) siRNA (ALG-125097)



# ASOs Have Demonstrated Clinical Anti-HBV Activity





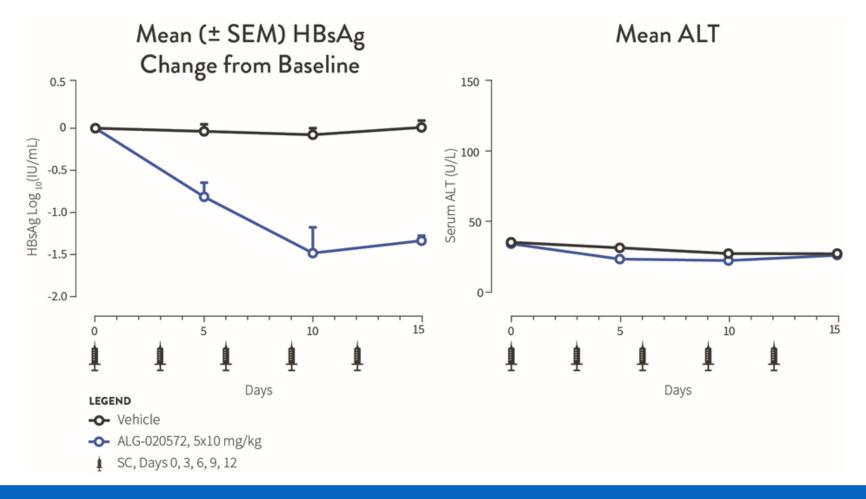


# Discovery of our ASO Drug Candidate

- We have utilized our bioinformatics approach in the selection and optimization of novel ASO constructs
- We optimized our sequences using our proprietary Luxna Biotech chemistries to potentially enhance in vitro potency, stability and safety
  - Luxna xeno nucleic acids (XNA) are 3<sup>rd</sup> generation bridged nucleic acids (BNA) that may improve nuclease resistance and reduce hepatotoxicity
- Potent ASOs were conjugated to our proprietary AM Chemicals GalNAc ligands for targeted liver delivery
- This approach has led to the selection of our ASO drug candidate ALG-020572



### Our ASO Drug Candidate in the Mouse AAV-HBV Model

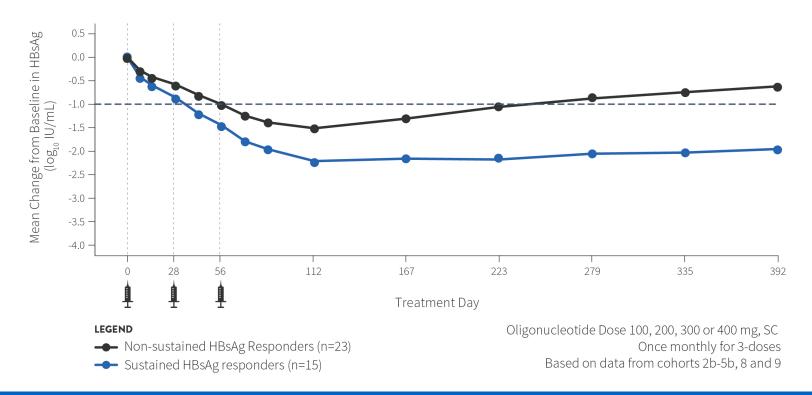


Potent in vivo activity of ALG-020572 in the AAV-HBV mouse model without ALT elevation



# siRNAs Have Demonstrated Clinical Anti-HBV Activity

JNJ-3989, a GalNAc conjugated, dual (X + S) trigger siRNA



JNJ-3989 in combination with a nucleos(t)ide analog demonstrated durable HBsAg reductions following three monthly doses

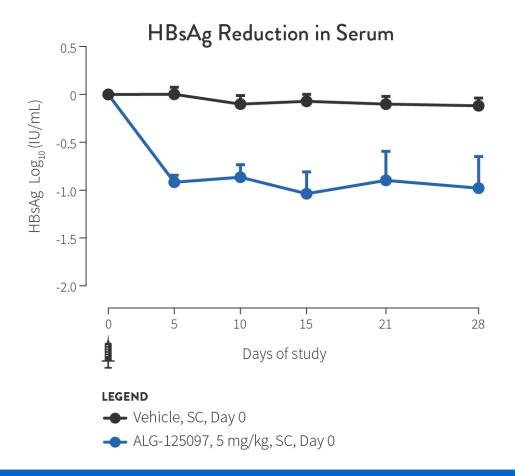


## Discovery of our siRNA Drug Candidate

- We have utilized our bioinformatics approach in the selection and optimization of novel siRNA target sequences
- We optimized siRNA sequences using our proprietary technology including chemical modifications and liver targeting conjugation to maximize in vivo potency
- Our approach to developing siRNAs may have safety, stability and potency advantages over ASOs and other siRNAs
- ALG-125097 has optimized chemistry enabling potency and durability in nonclinical animal models of CHB



## siRNA Drug Candidate in the Mouse AAV-HBV Model



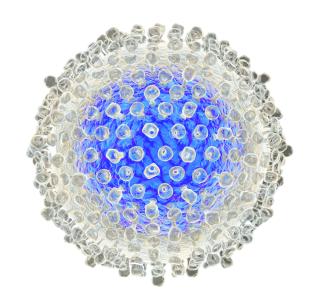
Potent and sustained antiviral activity following a single dose through 28-days of monitoring



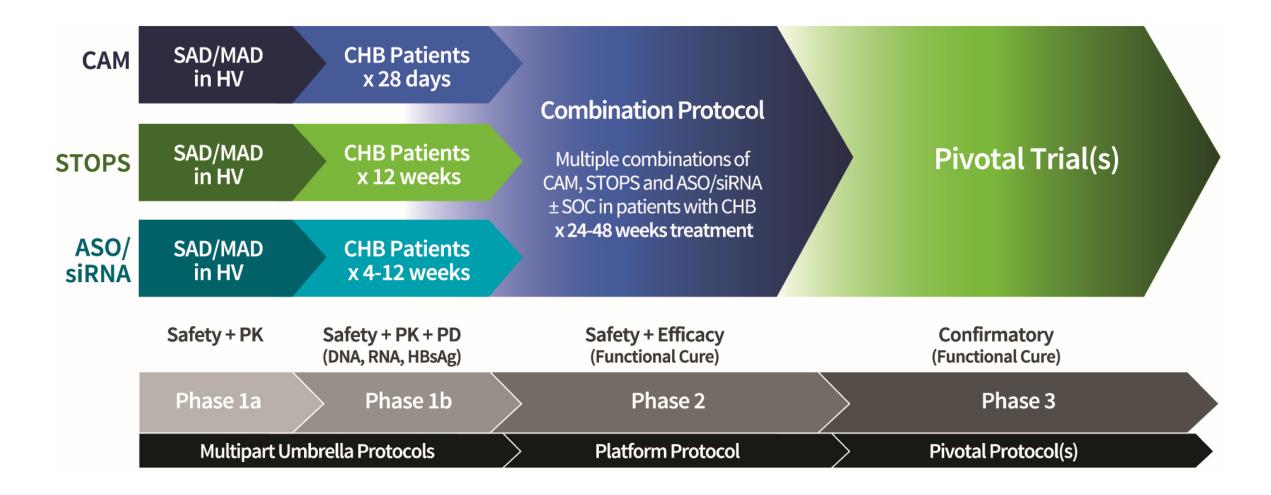


# Chronic Hepatitis B

Clinical Development Combination Strategy



# Our CHB Clinical Development Strategy





# Aligos CHB Clinical Development Timelines Phase 1 Target Timelines\*

ALG-010133 SAD/MAD in HV Multiple dosing CHB (STOPS) ALG-000184 SAD/MAD in HV Multiple dosing CHB (CAM) Multiple ALG-025072 SAD/MAD in HV (ASO) **Dosing CHB** SAD/ ALG-125097 MAD (siRNA) in HV Mar 2021 2022 **AASLD APASL APASL EASL AASLD** 





# **NASH**

ALG-055009



# THR-β Agonists are Clinically Validated

THR-β Drug^	Serum LDL Lowering*	Liver Fat Reduction at 12 Weeks* (MRI-PDFF)	NASH Resolution Without Fibrosis Worsening* at 36 weeks (Liver Biopsy)
Resmetirom**	17.3% (30 weeks)	22.5%	18.2%
VK-2809***	21.8% (12 weeks)	49.2%	_

THR-β agonists have demonstrated beneficial effects on serum lipid levels and liver histology



<sup>^</sup>All dose levels pooled.

<sup>\*</sup>Placebo corrected values.

<sup>\*\*</sup> In patients with NASH, Harrison, Lancet, 2019. Resmetirom = MGL-3196.

<sup>\*\*\*</sup> In patient with NAFLD, hyperlipidemia, Loomba, AASLD, 2018.

LDL: Low Density Lipoprotein, MRI-PDFF: magnetic resonance imaging proton density fat fractionation.

#### More Potent and Selective In Vitro than MGL-3196 and VK-2809

#### Relative THR-α and THR-β Activity in Cell-Based Assays

	EC <sub>50</sub> α (nM)	EC <sub>50</sub> β (nM)	Relative THR- $\beta$ Selectivity ( $\alpha/\beta$ )
Т3	14.2	11.6	1.2
MGL-3196	5927	2366	2.5
VK-2809 parent	366	269	1.4
ALG-055009	191	50	3.8

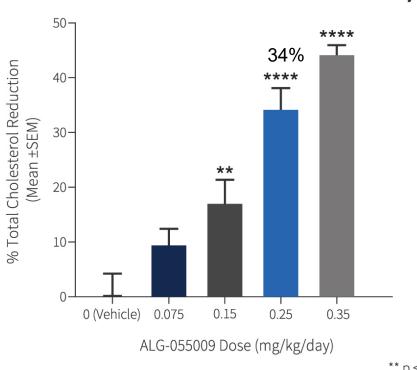
In vitro, ALG-055009 is 5-47x more potent compared to resmetirom (MGL-3196) and 2-3x more selective for THR-β than VK-2809\*

High β selectivity and potency may improve risk-benefit profile

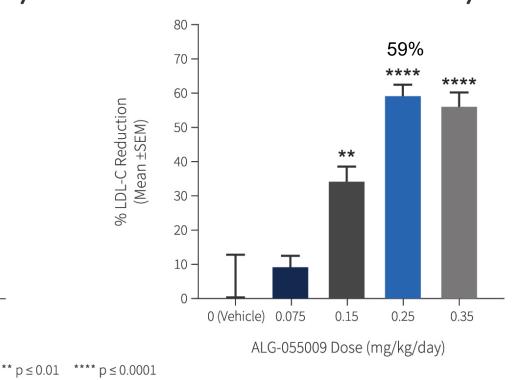


## Dose-Related Effect on Serum Lipids in the DIO Mouse Model





#### LDL-C Reduction on Day 28



0.25 mg/kg ALG-055009 resulted in greater serum lipid reductions in the DIO model compared to what has been previously reported for VK-2809 and resmetirom at exposures equivalent to Phase 2b/3 clinical doses without a thyroid hormone safety signal



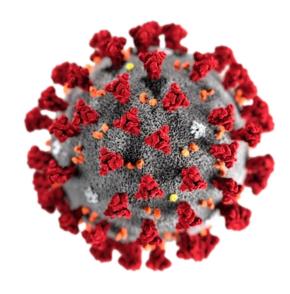
# Our NASH Strategy

- Advance our lead THR-β drug, ALG-055009
  - Phase 1 proof of concept study planned for 2021-2022
    - > Provide clinical confirmation of the safety, PK and efficacy advantages identified in nonclinical studies
    - Data readouts expected to begin in 2022
- Madrigal Phase 3 and VK-2809 Phase 2b readouts are expected in late 2021 through early 2022
  - We believe that enthusiasm for the THR-β MOA is likely to be high following these data
- If enthusiasm for the MOA is high, we plan to partner ALG-055009 with a NASH company with a complementary MOA





# Coronavirus



# Our Approach to Developing a Broad Coronavirus Treatment

- We are leveraging our virology and chemistry expertise to develop purpose-built drugs with distinct MOAs
  - Coronavirus protease inhibitors in collaboration with CD3/Rega at KU Leuven
  - Oligonucleotides
- We are aiming to develop drug candidates that
  - Are broadly active against diverse coronaviruses
  - Have a high barrier to resistance
  - Have a therapeutic window compatible with prophylaxis and treatment
  - Can be combined to prevent emergence of resistance and provide broader strain coverage
- We are prioritizing regimens with MOAs that may offer therapeutic benefit for potential zoonotic coronavirus transmissions in the future



# Coronavirus Protease Inhibitor Collaboration CD3/Rega at KU Leuven

Centre for Drug Design and Discovery ("CD3")



CD3



Rega Institute
Professor Johan Neyts
KU Leuven

Identified multiple nM potent coronavirus protease inhibitors

ALIGOS



# Aligos Therapeutics Summary

- We have assembled a world-class team with a proven track record of success
- Strong management team supported by top tier investors
  - \$100M\* Series A in September 2018, \$125M\*\* Series B in December 2019 / October 2020
  - \$167.2M gross proceeds from our IPO in October 2020 and the underwriters exercise of its overallotment option
- We are advancing multiple drug candidates with clinically-validated MOAs with the goal of rapidly developing optimized combination regimens
  - CHB
    - > STOPS molecule Phase 1 clinical trial initiated in August 2020 (ALG-010133)
    - → CAM Phase 1 clinical trial initiated in October 2020 (ALG-000184)
    - > ASO and siRNA oligonucleotide compounds are planned to advance into the clinic
  - NASH
    - > We are planning to advance ALG-055009 into clinical development and partner after Phase 1
  - Coronavirus
    - We have identified multiple nM potent coronavirus protease inhibitors



# ALIGOS THERAPEUTICS