

Bringing new hope with mRNA-targeted therapies

# Introduction to Veritas In Silico

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## Forward-looking statements

This presentation may contain forward-looking statements as defined in the United States Private Securities Litigation Reform Act of 1995. The forward-looking statements do not mean that the Company's management guarantees its future performance. The Company may use expressions such as "aim", "predict", "believe", "continue", "attempt", "estimate", "expect", "plan", "intend", "possible", "plan", "potential", "probability", "project", "risk", "seek", "should", "make effort", "propose", "will", as well as other similar expressions to explain the forward-looking statements. Forward-looking statements may also be identified by discussions on strategies, plans or intentions. Forward-looking statements contained herein are based on the current assumption and judgement of the Company which are made based on currently available information. As such, these forward-looking statements include known and unknown risks, uncertainties and other factors. The Company's actual results or financial conditions could differ materially from those expressed in or suggested by the forward-looking statements due to such risks, uncertainties and other factors.

## Veritas In Silico (VIS) at a glance

Japan-based biotech company focusing on mRNA-targeted therapies



#### Founded 2016 IPO 2024 (TSE Growth)

- Pioneer in mRNA targeting
- Technology based on decades of mRNA research



Our mission: Deliver life-changing therapies

- Informatics and biology to create effective and safe therapies
- Potential for application to most diseases



Strategic partnerships with pharma/biotech

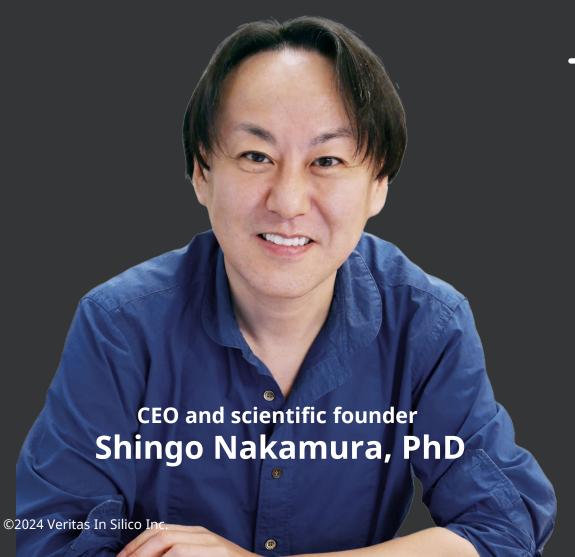
- Multiple partnerships with global and Japanese companies
- EU: Oncodesign Services, Liverpool ChiroChem



Pipeline for diseases of unmet medical need

- Transition from platform to hybrid business
- Aiming for first- or bestin class opportunities

## A VISionary's journey in mRNA targeting



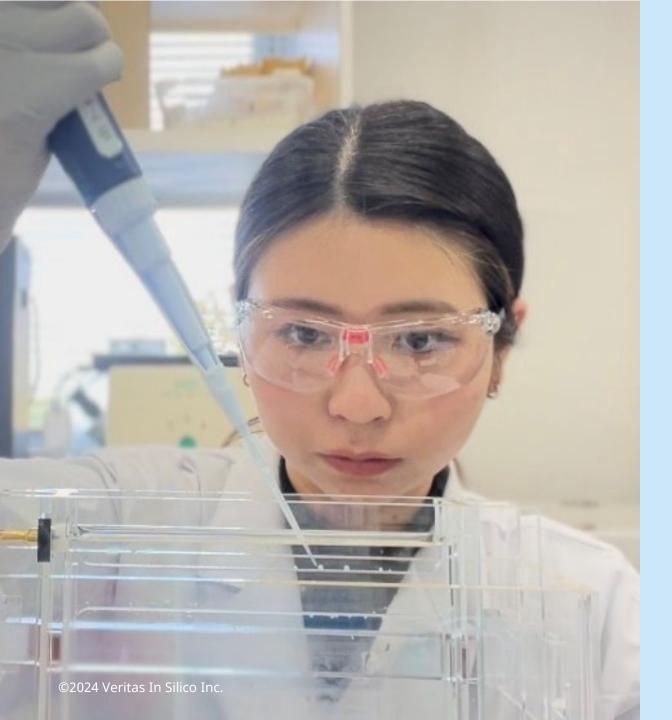


Statistical mechanics

- + Thermodynamics+ Artificial intelligence (AI)
  - = mRNA structural analysis



World's first business model patent for mRNA-targeted small molecule drug discovery in 2004



### Our team

## World-class scientists with expertise in:

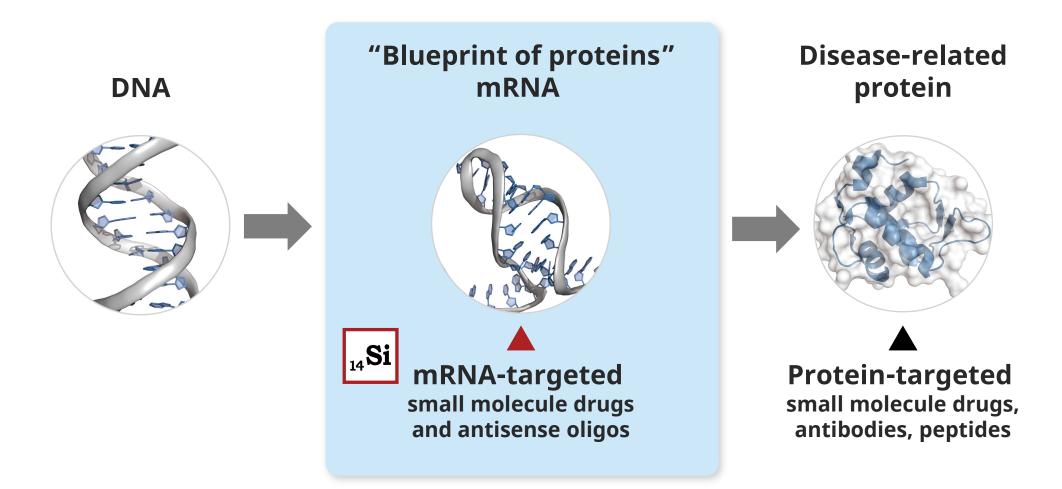
Drug discovery mRNA biology Informatics

## Leadership team with extensive experience in:

Pharmaceutical and biotech industries

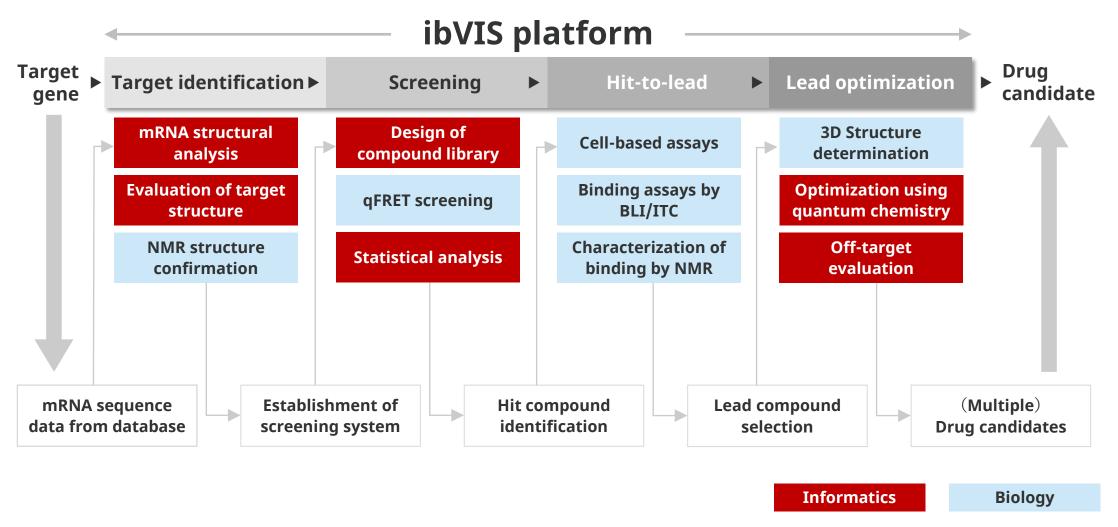
## Why target mRNA?

Targeting mRNA allows us to reach the "undruggable" space and create new therapies



## ibVIS: mRNA-targeted drug discovery platform

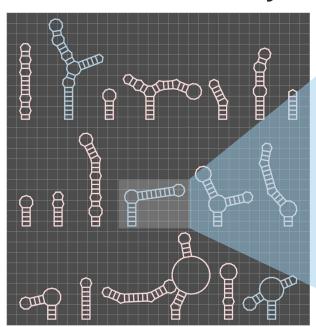
A one-stop solution that integrates informatics and biological tools for mRNA targeting



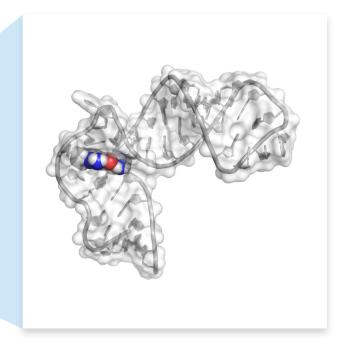
## mRNA structural analysis technology

VIS' core strength that enables the identification of druggable structures on mRNAs

## In silico mRNA structural analysis



## Small molecule target structure on the mRNA



#### 01 Fast

Target structures on mRNA can be identified in significantly less time than using experimental methods

#### 02 Accurate

The target structures identified using our in silico technologies have an accuracy of >90%, while other computational methods have an accuracy of only 60–70% based on our evaluation

#### **03** Well-validated targets

The target structures that we select are rigorously tested to ensure they are optimal targets for small molecule drug discovery

## Scoring system for evaluating mRNA structures

Based on results of over 50 screening programs, constantly updated

## **Structural conservation Thermodynamic** Existence stability probability ΔG Location **Druggability Total score** •—AUG **Current version 5.5**

## Rigorous validation of candidate mRNA targets

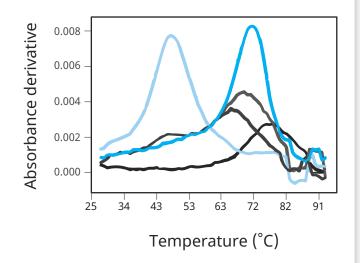
Conduct tests to check structure and suitability for screening

# Verify target structure

To verify that the mRNA sequences adopt the therapeutically relevant structure

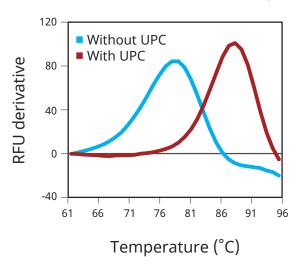
<sup>1</sup>H Chemical shift (ppm)

#### **Determine assayability**



To determine whether the melting temperature (Tm) of the mRNA sequence falls within the detection range

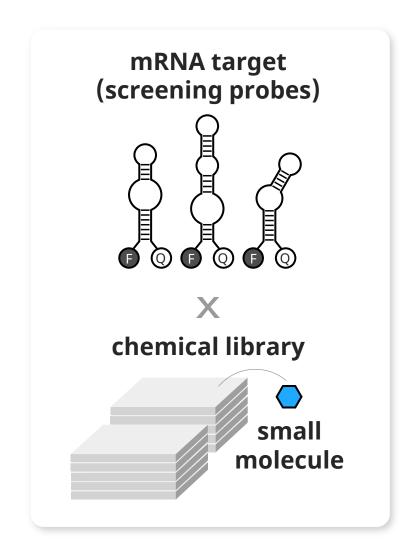
#### **Confirm functionality**

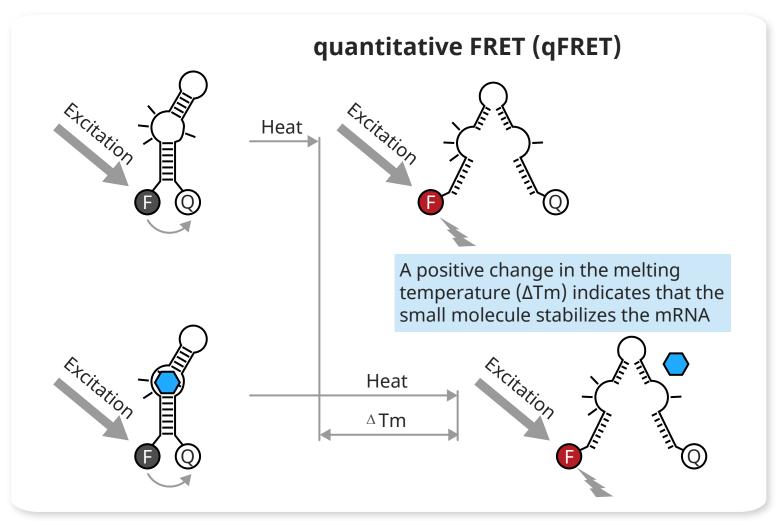


Tm of the screening probes in the presence and absence of a universal positive control (UPC) are measured to check functionality

## High-throughput screening using qFRET

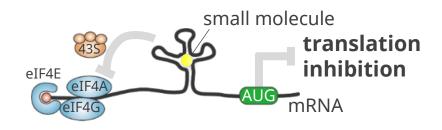
Identifies small molecules that bind to and stabilize mRNA structures

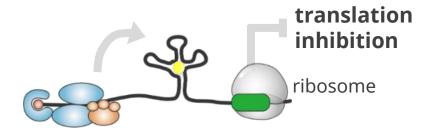




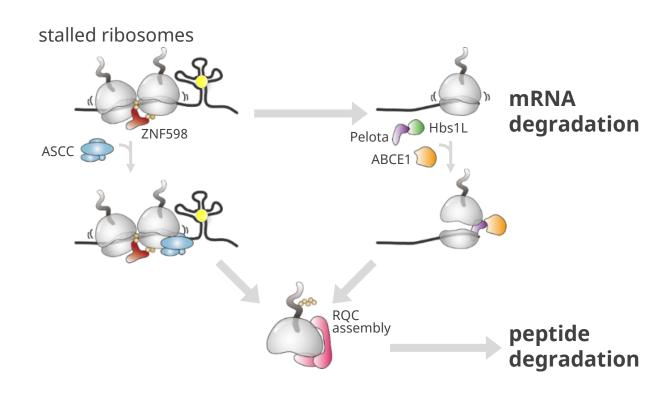
## Stable mRNA structures inhibit protein production

## Prevent the formation of the translation pre-initiation complex





## Promote ribosomal collision that signals peptide or mRNA degradation

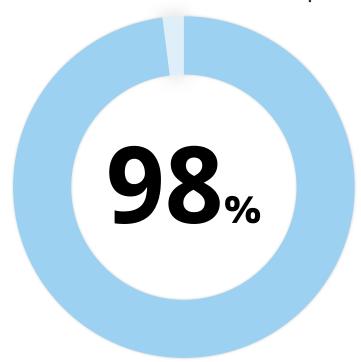


Adapted from *Nat. Rev. Mol. Cell Biol.* **19**,158-174 (2018).

Adapted from *Mol. Cell* **79**, 603-614 (2020).

## ibVIS screening technology with high success rate

Obtained small molecule hit compounds for



of the 50+ mRNA targets we screened



**Target Identification** 

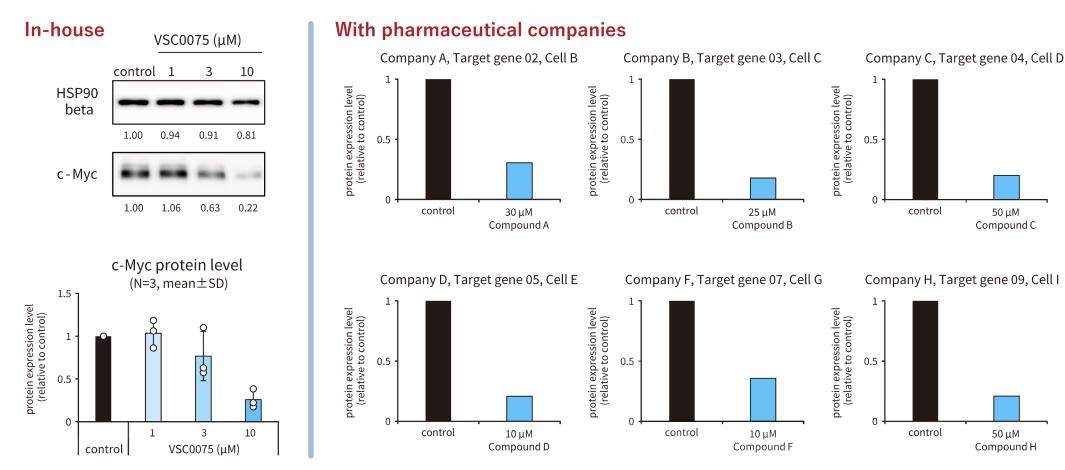
50 μΜ

Compound C

50 μM

Compound H

## Hit compounds show cellular activity

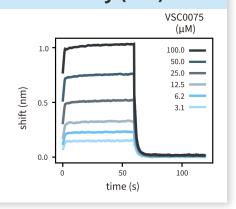


Left: Addition of the compound (VSC0075)—obtained in the screening against a target structure found on the mRNA of a disease-related protein (c-Myc)—to the cells resulted in a predominant decrease of the c-Myc protein. HSP90 beta, a representative of common proteins, was hardly affected. Right: Cell-based assay results of the compounds (Compound A-H) obtained in the screening for six partners are shown, with one example for each.

## Techniques to understand binding and identify leads

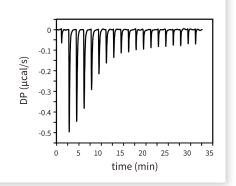
#### **Biolayer interferometry (BLI)**

Examine the binding rate between the RNA and the compounds



#### **Isothermal titration calorimetry (ITC)**

Precisely determine the binding strength of the compounds to the RNA



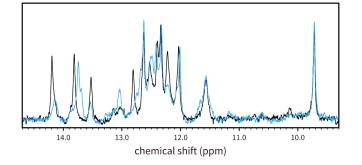
#### Synthesis of derivatives

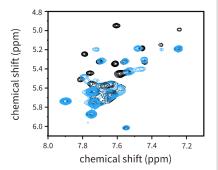
**SAR studies** 

#### Nuclear magnetic resonance (NMR) measurement

Investigate the binding sites of the compounds on the RNA

- --- With small molecule
- --- Without small molecule

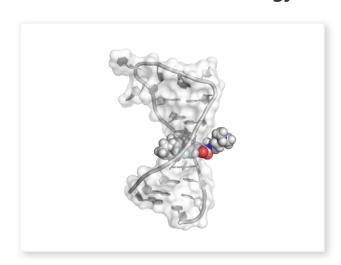






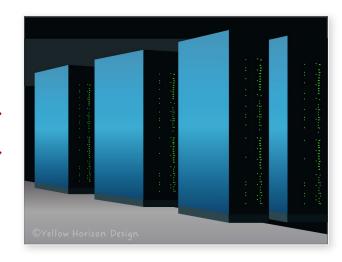
## RNA structure determination and quantum mechanics for lead optimization

## RNA-tailored structure determination technology



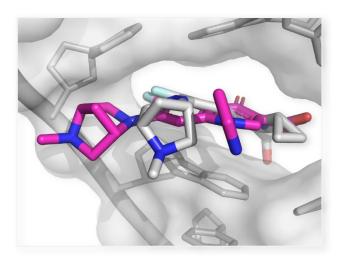
We are uniquely equipped to perform RNA 3D structure determination, with our scientists having rare expertise in RNA X-ray crystallography and NMR spectroscopy

## RNA -small molecule interaction analysis technology



We use Japan-made quantum mechanics methods and the Fugaku supercomputer to thoroughly analyze the interactions between mRNA targets and small molecule binders

## Rational optimization of mRNA-targeted small molecules



Our technologies enable us to rationally design small molecules with higher potency and selectivity for mRNA. As a result, we can optimize small molecules faster and more efficiently than traditional methods

## Our pipeline of mRNA-targeted small molecules

We are developing an in-house pipeline to address diseases of unmet medical need

	Target identification ►	Screening	•	Hit-to-lead	<b>•</b>	Lead optimization
Cancer						
Lymphoma (gene: c-Myc)					I I	
Prostate cancer (gene: AR)						
Various cancers (gene: STAT3)					į	
Various cancers						
Various cancers					 	
Various cancers					1	
CNS diseases						
Pain disease					 	
Rare diseases						
Cardiovascular disease						
Muscular disease					 	

## Milestone achievements in joint drug discovery

Research with Takeda, Shionogi, RaQualia, and Toray all progressing smoothly

PRESS RELEASE in the past year

**Takeda** 

2024.06.19 Veritas In Silico announces milestone achievement in collaborative research with Takeda

**SHIONOGI** 

2024.09.24 Notification regarding achievement of milestone in joint drug discovery research with Shionogi & Co., Ltd.

**RaQualia** 

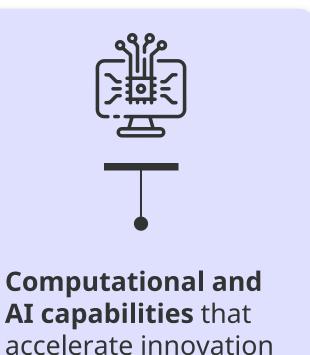
2023.12.21 RaQualia Pharma and Veritas In Silico announce that both companies have achieved the milestone in a joint research collaboration for mRNA-targeted small molecule drug discovery

## Why partner with Veritas In Silico?

When you collaborate with us, you gain access to:



ibVIS: a cutting-edge mRNA-targeted drug discovery platform





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