

Accelerate your mRNA-based proof-of-concepts studies with Tebubio

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# Webinar Agenda

1. Tebubio at a glance

2. In-house µ-scale mRNA production service

3. Q&A session



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## Tebubio at a glance

We facilitate Life Sciences Research everyday, contributing to advance Science for a brighter future.



Founded in 1953, privately-owned



Offices operating across Europe & Contract Research Services (CRS) Laboratory based in Europe



Unique combination of innovative solutions and Contract Research Services

With Tebubio, concentrate your efforts and accelerate what's most important to you: **Your Research.** 



# Tebubio at a glance

Our CRS activity



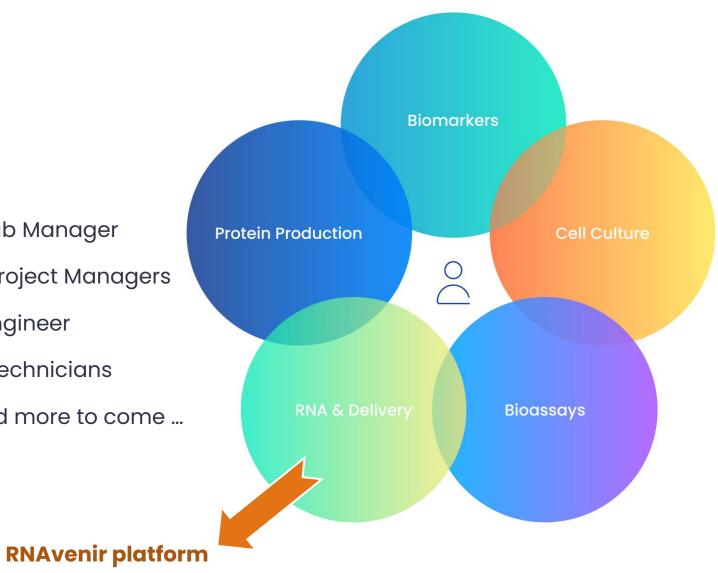
1 Lab Manager

5 Project Managers

1 Engineer

5 Technicians

And more to come ...





# Supporting you from A to Z in your projects

Preclinical Research Phase

Clinical Phase

MRNA μ-scale production

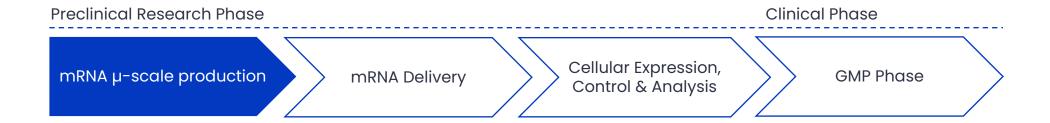
mRNA Delivery

Cellular Expression,
Control & Analysis

GMP Phase



# Supporting you from A to Z in your projects





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1. Tebubio at a glance

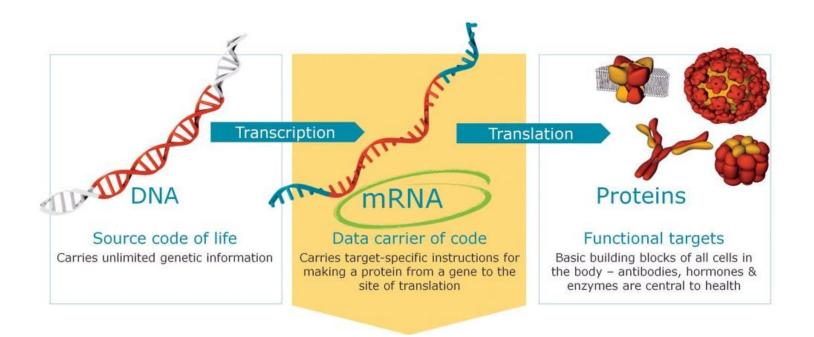
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## **RNA Therapeutics**

Since the Covid-19 outbreak, mRNA-based therapeutics represent an increasingly used alternative to classical vaccines and solutions



- Faster
- Safer
- Cost Effective

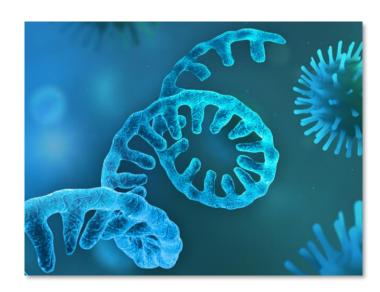


## **RNA Therapeutics**

Not only a vaccine alternative for infectious diseases

### Applications for mRNA

- mRNA vaccines
- Cancer immunotherapy
- Cell therapy (CAR-T)
- Replacement therapy
- Anti-virals
- Rare diseases

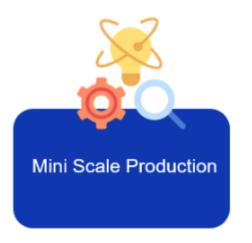


### **Applications for RNA**

- Analytical methods
- Diagnostic methods...



# A service tuned to your preclinical needs









5 days

### μ-scale mRNA production service

Quickly obtainable, small amounts of custom mRNA for screening purposes and proof of concept



### mRNA structure

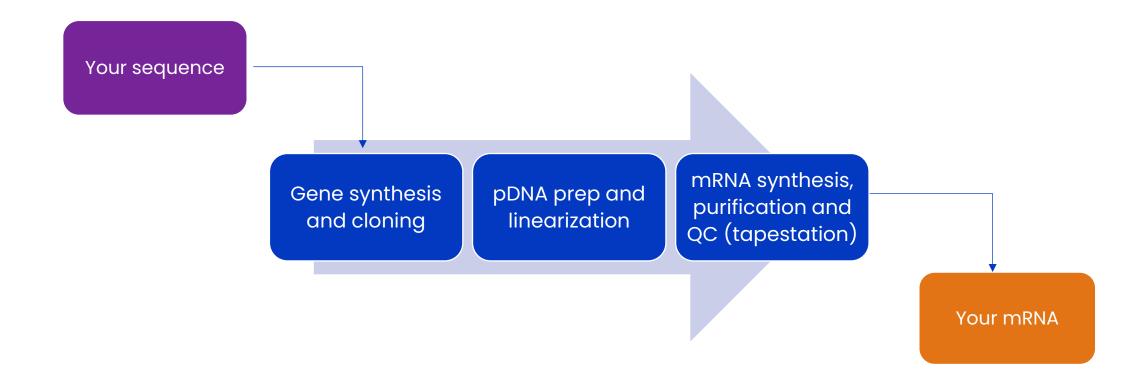
### Key mRNA components

- Coding sequence
- Untranslated regions (UTRs)
- Capping (Uncapped, ARCA, Cap1)
- Poly-A tail





### mRNA Production workflow





### Optimal sequence for better transcription and translation

#### Benefit from...

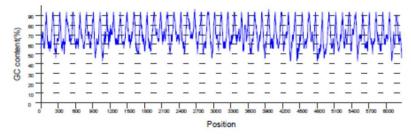
- Sequence optimization (Hairpin, GC...) to improve transcription
- Codon optimization for on optimal expression in your model (species)



Design and Optimize a functional mRNA sequence

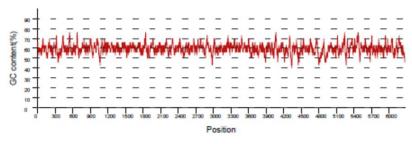
#### **GC Content Analysis**

Hairpin Analysis	Wild-type	Optimized
Small Hairpins	27	5
Medium Hairpins	14	1
Large Hairpins	3	0



#### Average GC content = 70%

Plot of GC content of wild-type gene. Y and X axis represent local GC content (windows qize = 30 bp) and base position, respectively



#### Average GC content = 59%

Plot of GC content of optimized gene. Y and X axis represent local GC content (windows qize = 30 bp) and base position, respectively



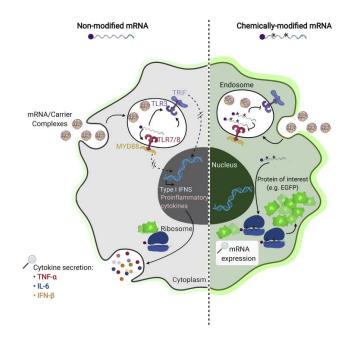
## Optimal sequence for lower immunogenicity and higher expression

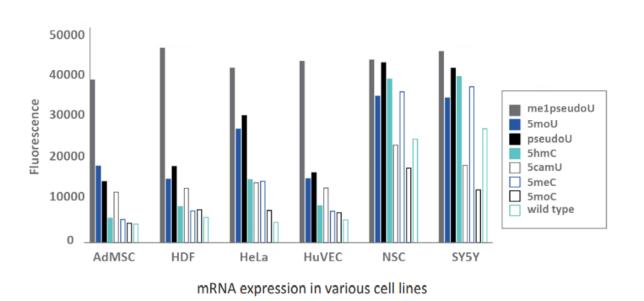
#### Benefit from...

- A U-depletion
- Introduction of modified UTPs (melpseudoU pseudoU 5moU...)









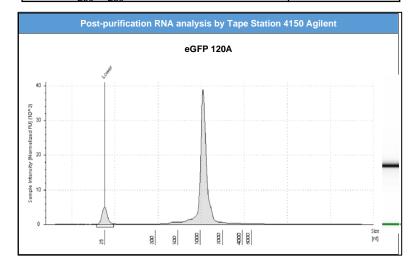
## Optimized vector

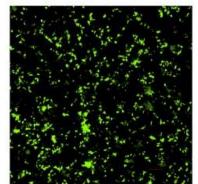
#### Start from...

- Your own DNA template
- A customized template designed for you
- Cloning in our proprietary pDNA template with optimized UTRs & poly-A tail (GMP compatible)

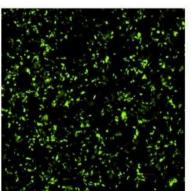


	eGFP 120A
RNA concentration (ng/µL)	1220
Ratio A <sub>260</sub> /A <sub>280</sub>	2.082
Ratio A <sub>260</sub> /A <sub>230</sub>	2.225





Tebubio eGFP mRNA



Trilink eGFP mRNA

Representative Image of HEK293 cells transfected with eGFP mRNA from Tebubio or Trilink



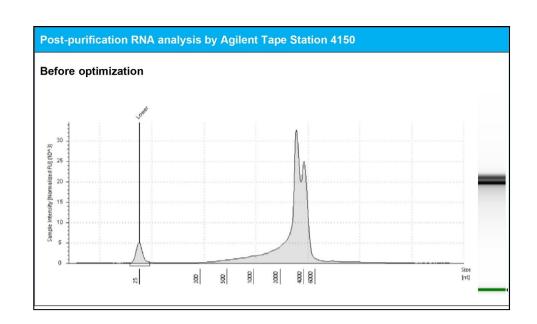
# Optimal transcription

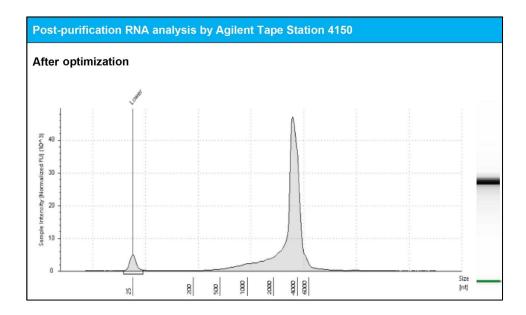
#### Benefit from...

Optimization of production and purification conditions



Design and Optimize a functional mRNA sequence



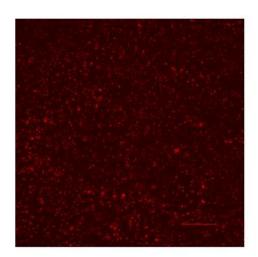


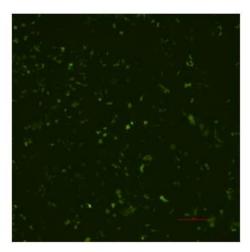


## Monitor mRNA delivery

#### Benefit from...

- Introduction of fluorescent UTPs, or others
- Green, Red, Far Red, Orange...

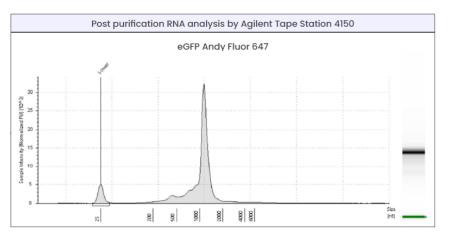




Example of eGFP mRNA labelled with a red fluorophore transfected into HEK293 cells. Visualisation of transfected mRNA (red) and eGFP protein produced.

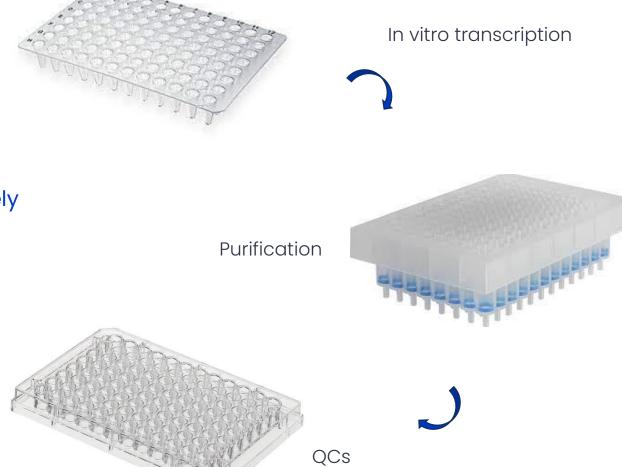


	eGFP mRNA with Andy FLuo 647
RNA concentration (ng/µL)	2992
Ratio A <sub>260</sub> /A <sub>280</sub>	2.059
Ratio A <sub>260</sub> /A <sub>230</sub>	2.120





# Speed up your sequence development and screening

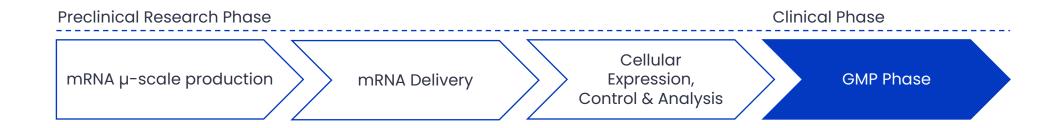


Find the best sequence quickly and cost-effectively

Up to 192 RNA in parallel per batch (100µg)



## A DNA template compatible for GMP production



Our proprietary plasmid DNA template is **GMP-compatible**, for a **seamless path from proof of concept to large-scale manufacturing**.



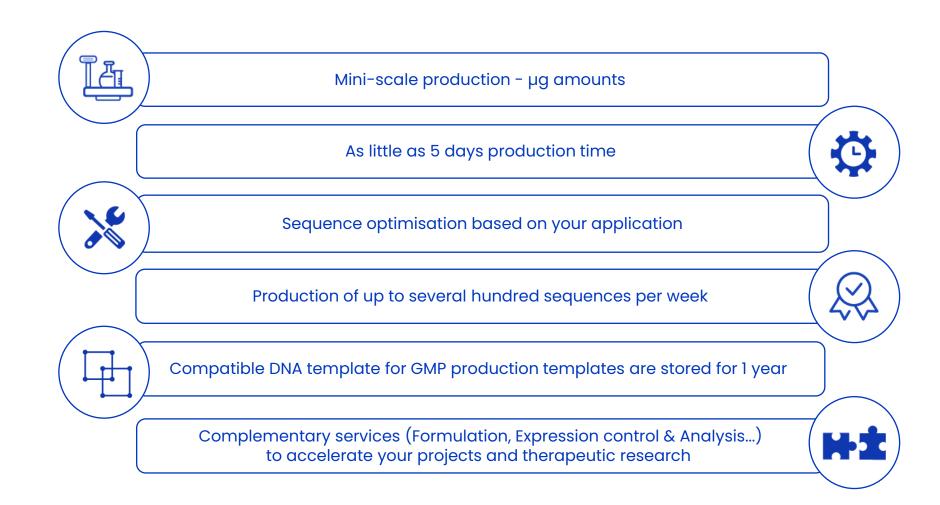
# A complete offer to speed up all your projects



- Optimize your mRNA delivery Mechanism & Control your transfection (Formulation platform)
- Get fast and effortless access to Cellular Models (Cell Culture Platform)
- Take care of your Quality control & Read-Out Analysis (Biomarkers and Biostatistics platform)



## Take Home Message





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