VIDEO DROP

Assessing the particle concentration of mRNA-LNP using VIDEODROP: A new metric for LNP manufacturing, optimization & standardization

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Videodrop is an innovative nanoscale imaging technology. Based on the principle of interferometry, **Videodrop** makes it possible to measure the size and concentration of nanoparticles:

- In real-time **(40 s)**
- In a single drop (7 μL)
- Between **80 nm to 500 nm**
- Within a concentration range of 1E8 to 1E10
 part/mL
- Without labelling

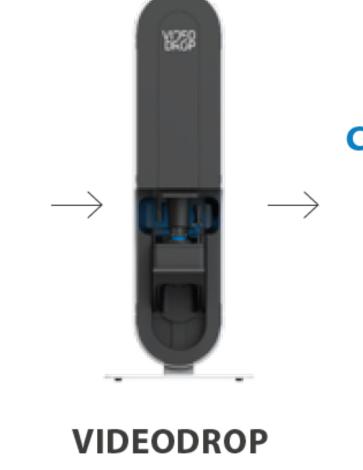
Here, we propose to use Videodrop for the analysis on mRNA-LNP and highlight the interest of measuring the particle concentration of mRNA-LNP.

The current analytical strategies employed for mRNA-LNP characterization lie primarily on size, polydispersity, and zeta-potential.

However, particle concentration, evaluated through a **single particle measurement technique**, is an important parameter to monitor for several reasons:

- Quality control tests
- Stability assessment
- Standardization for comparative studies





Particle concentration

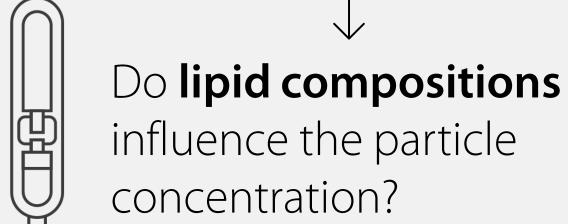
Particle & Size distribution

Quality control tests
Stability assessment
Standardization for
comparative studies

Why is LNP particle concentration important?



SOLUTION PREPARATION RNA/LIPID



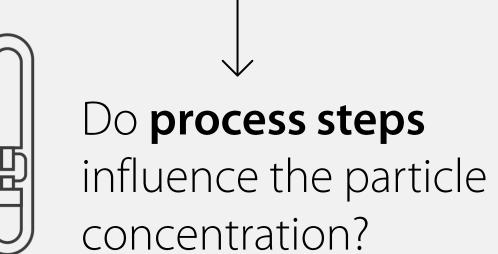
2

LNP FORMULATION

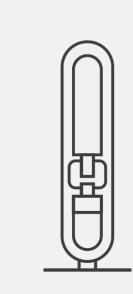


Do microfluidic process conditions influence the particle concentration?

BUFFER EXCHANGE & CONCENTRATION

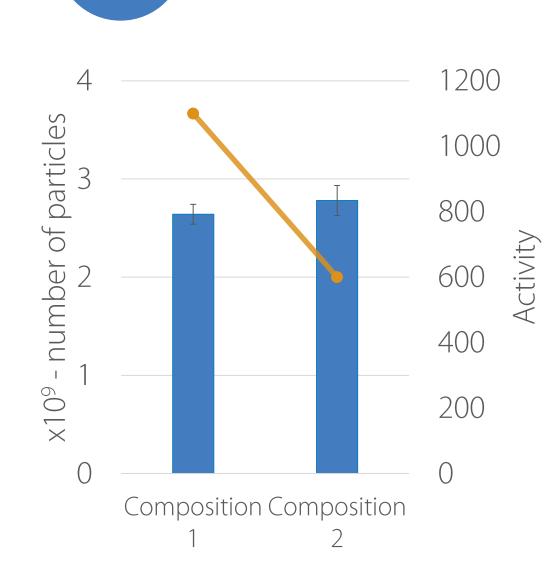






Do **storage conditions** influence the particle concentration?

SOLUTION PREPARATION RNA/LIPID



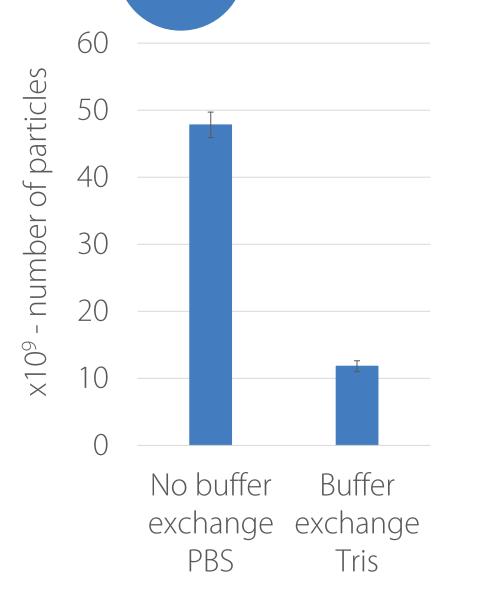
Lipid composition optimization

In this experiment, two LNPs with different lipid compositions were produced using the same process.

Particle concentration was not affected by the lipid composition; however, lipid composition 1 exhibits higher activity compared to lipid composition 2.

Despite having approximately, the same number of particles, lipid composition 1 is more efficient compared to lipid composition 2.





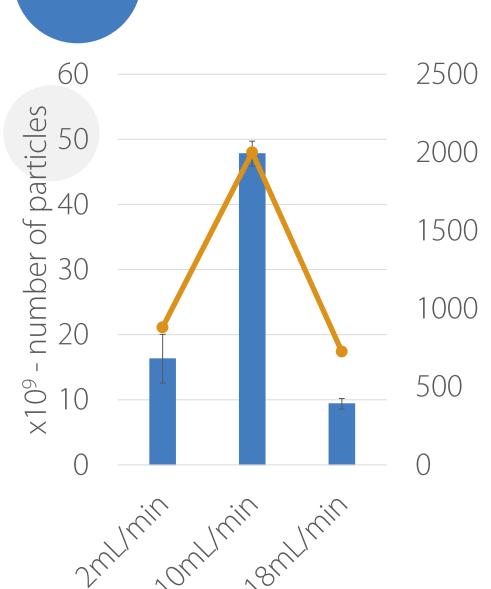
Buffer exchange and concentration steps are crucial and require optimization to improve the yield.

Buffer exchange optimization

This experiment examines the influence of incorporating a buffer exchange step.

In this case, the buffer exchange step results in a 75% reduction in the number of particles.

2 LNP FORMULATION

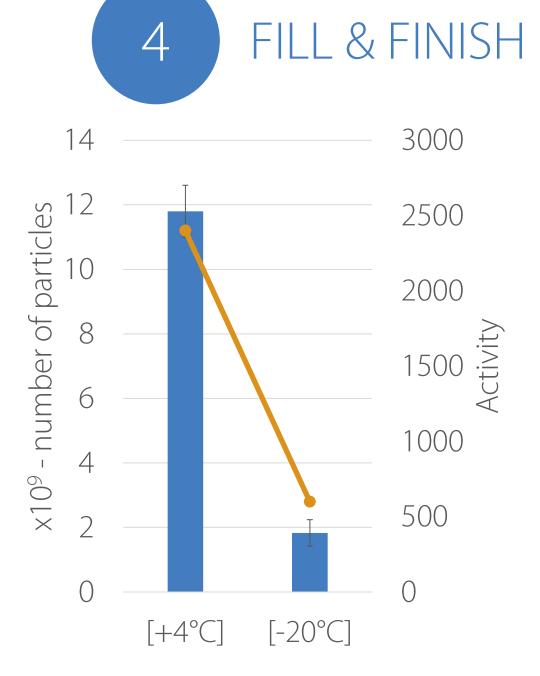


Mixing speed optimization

Here, we compare LNPs having the same composition but prepared with different mixing speeds on the microfluidic device.

The mixing speed step has a significant influence on the concentration of the generated particles.

In this case, the activity is directly related to the number of particles.



Storage optimization

To ensure product quality, it is essential to optimize storage conditions.

In this experiment, we tested the same product stored at +4°C or -20°C.

The results indicate that the storage at -20°C leads to a decrease in the number of particles as well as a loss in activity.

Videodrop is a ready-to-use tool for measuring the particle concentration and size distribution of LNPs in a single drop.

- Videodrop enables better control of the production process by quickly and easily measuring the concentration.
- Moreover, in comparative studies, standardizing cell-based assays (transfection activity) using the number of nanoparticles improves the evaluation of LNPs produced under different conditions.

VIDEO DROP

VIDEODROP IS AN IDEAL TOOL TO BE INCORPORATED INTO AN ANALYTICAL STRATEGY TO ASSESS LNP QUALITY.