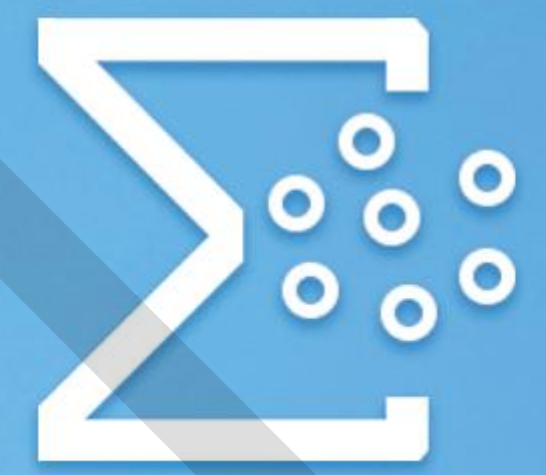


# NanoAnalyzer

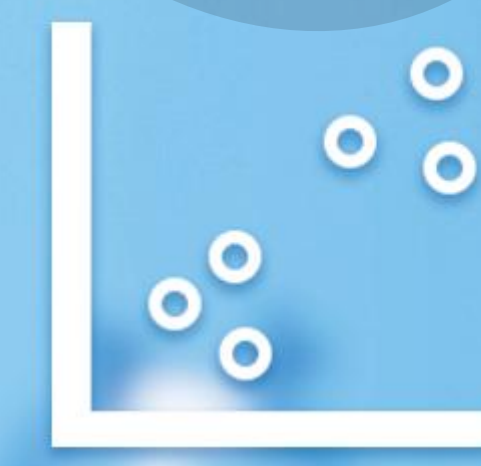
- Unmatched Performance
- Superior Sensitivity
- Innovation for NanoWorld



Size Distribution



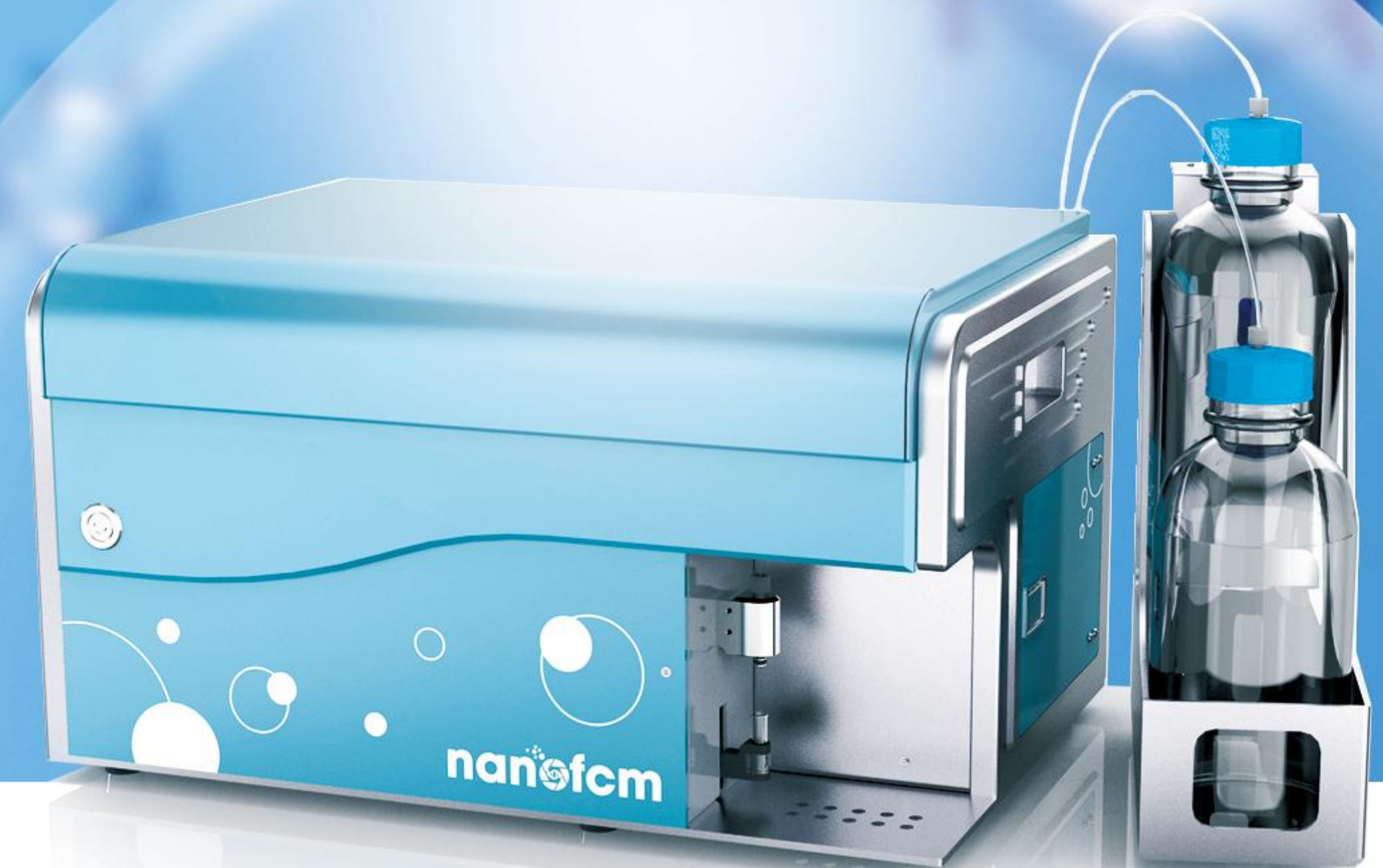
Particle Concentration



Phenotyping



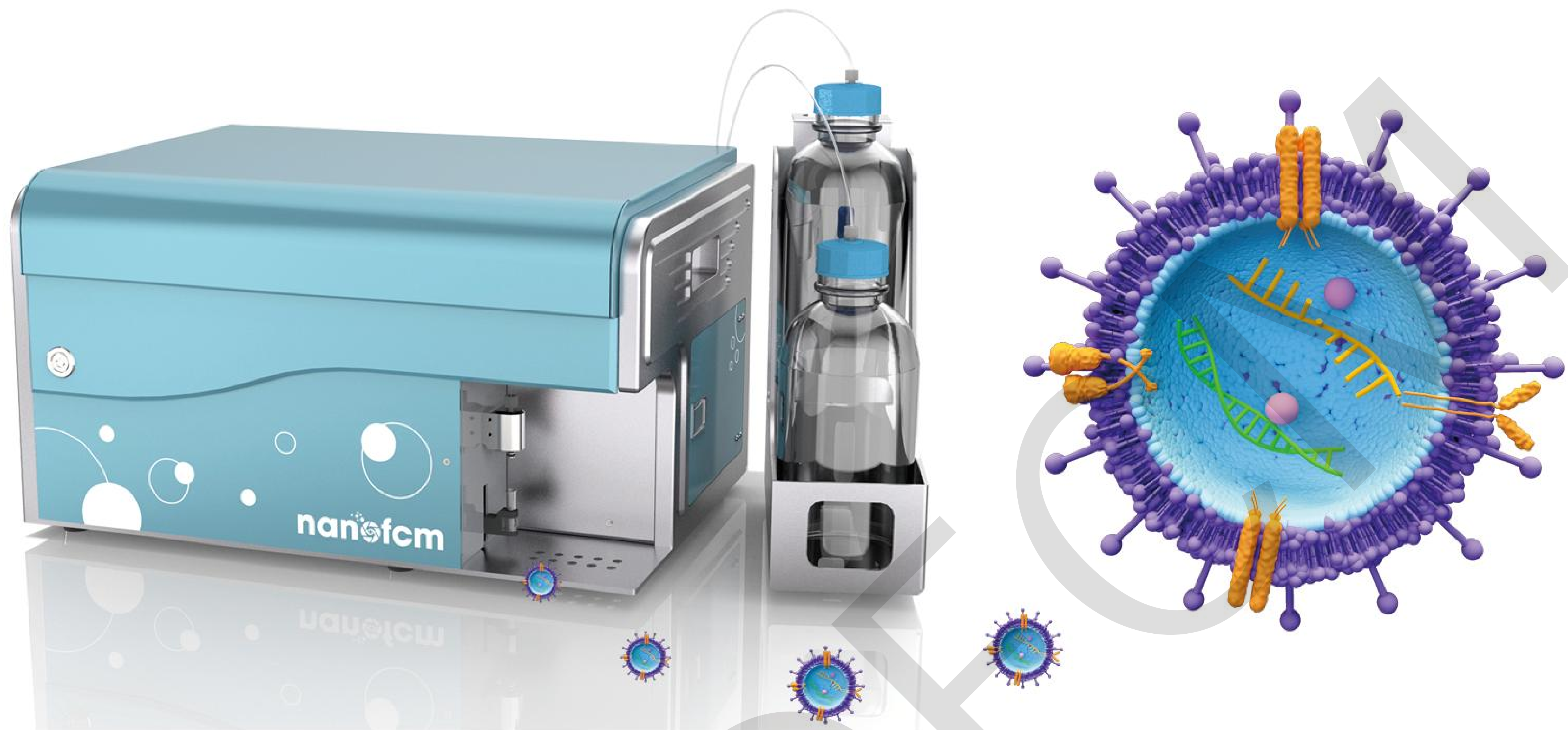
Multiparameter



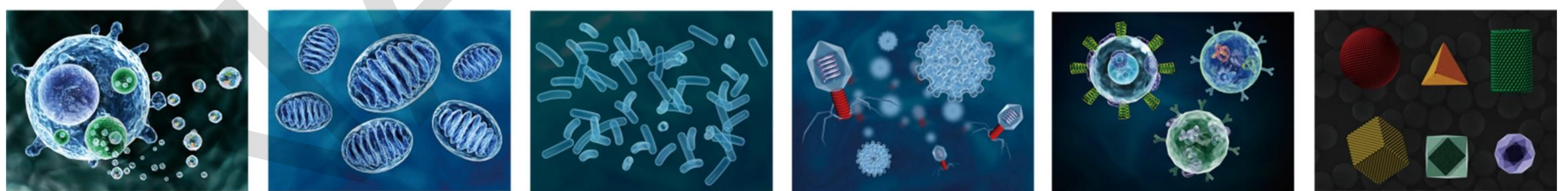
# Flow NanoAnalyzer

## High Sensitivity Flow Cytometry for Nanoparticle Analysis

Deciphering Bio-Nanoparticles (7-1000 nm)



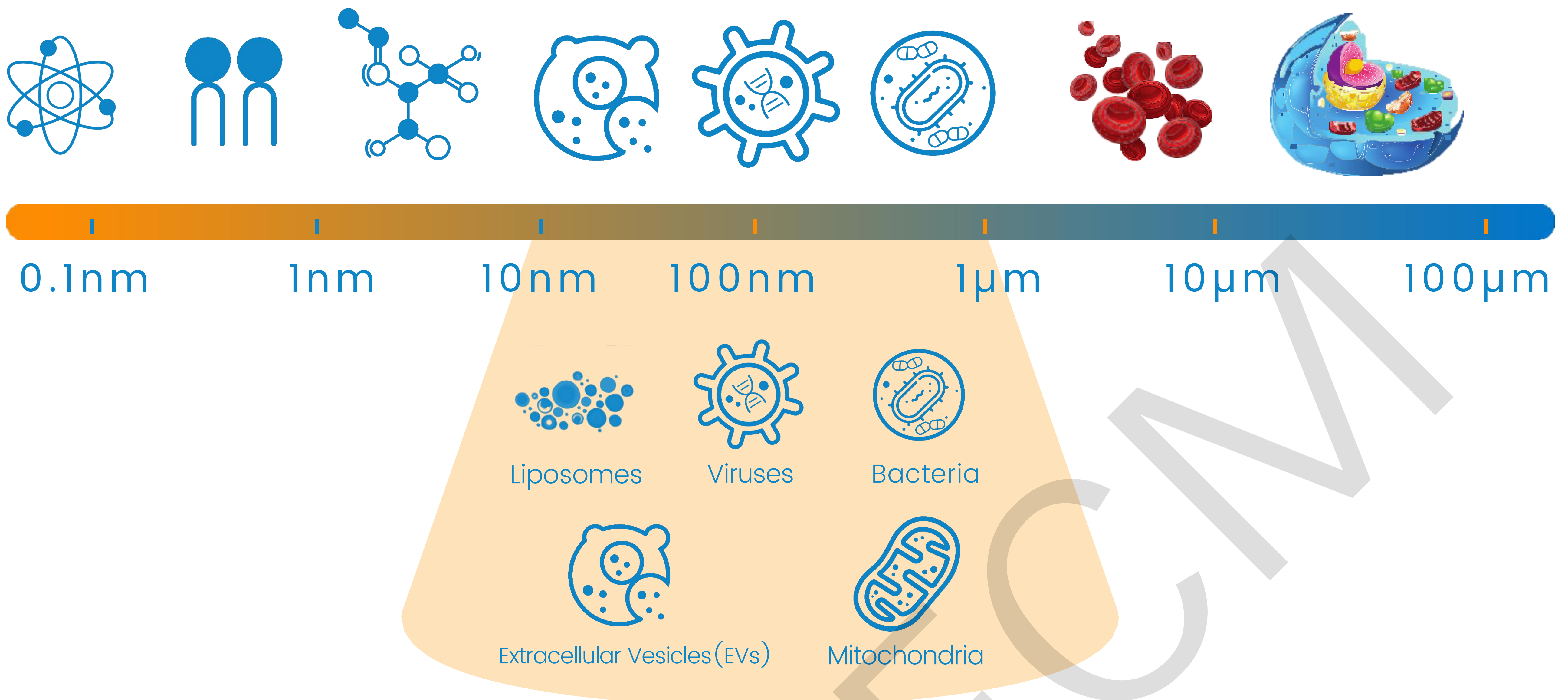
**NanoAnalyzer** is expected to become a powerful tool for life science, nanoscience and nanotechnology studies. NanoAnalyzer can be used for the multiparameter characterization of natural and synthetic nanoparticles (7-1000 nm) at the single-particle level, such as extracellular vesicles, mitochondria, bacteria, viruses, nanomedicines and nanomaterials. Combining light scattering and fluorescence detection, high-resolution distributions of particle size and biochemical properties can be acquired simultaneously in 1-2 minutes.



- Unprecedented Sensitivity for Scatter (24 nm) and Fluorescence (single PE).
- Multiparameter Analysis of Nanoparticles.
- Complementary to Conventional FCM in Size Below 200 nm.
- Covers the Entire Size Range of EVs.

# Application Fields

Measurement range from 7 to 1000 nm



## Application Areas:

- Diagnostics
- Therapeutics
  - Naive EVs &
  - Engineered EVs
- Cancer Treatment
- Fundamental Research

## Data Required:

- Particle Size
- Concentration
  - Total Populations &
  - Sub Populations
- Surface Functionalities
- Cargo/Payload

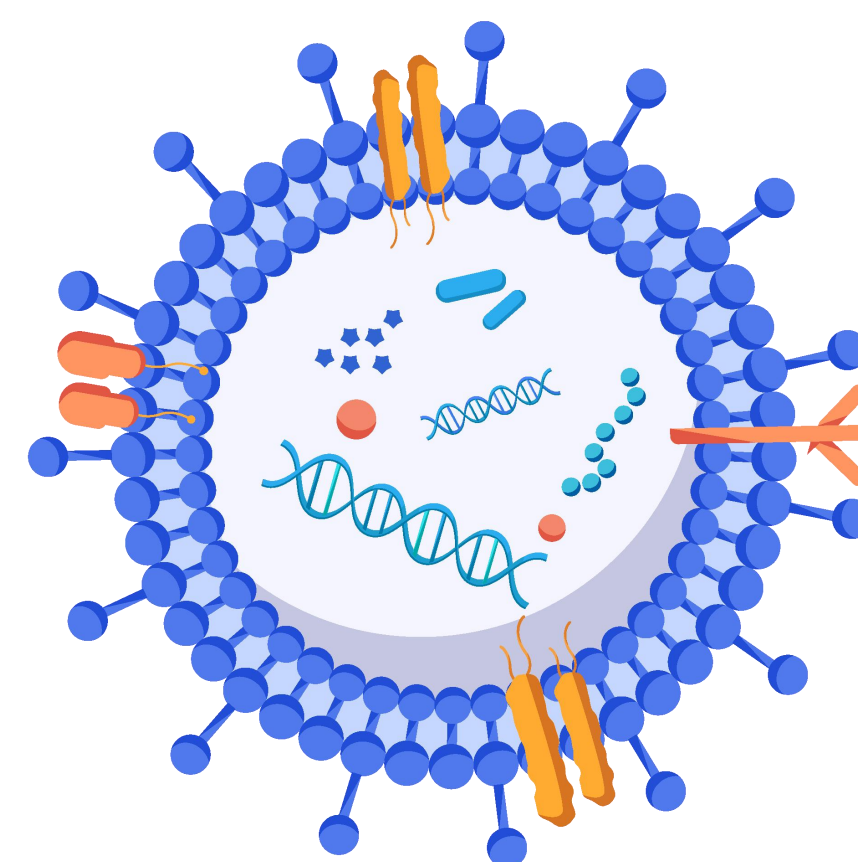
# Products & Services

## NanoAnalyzer



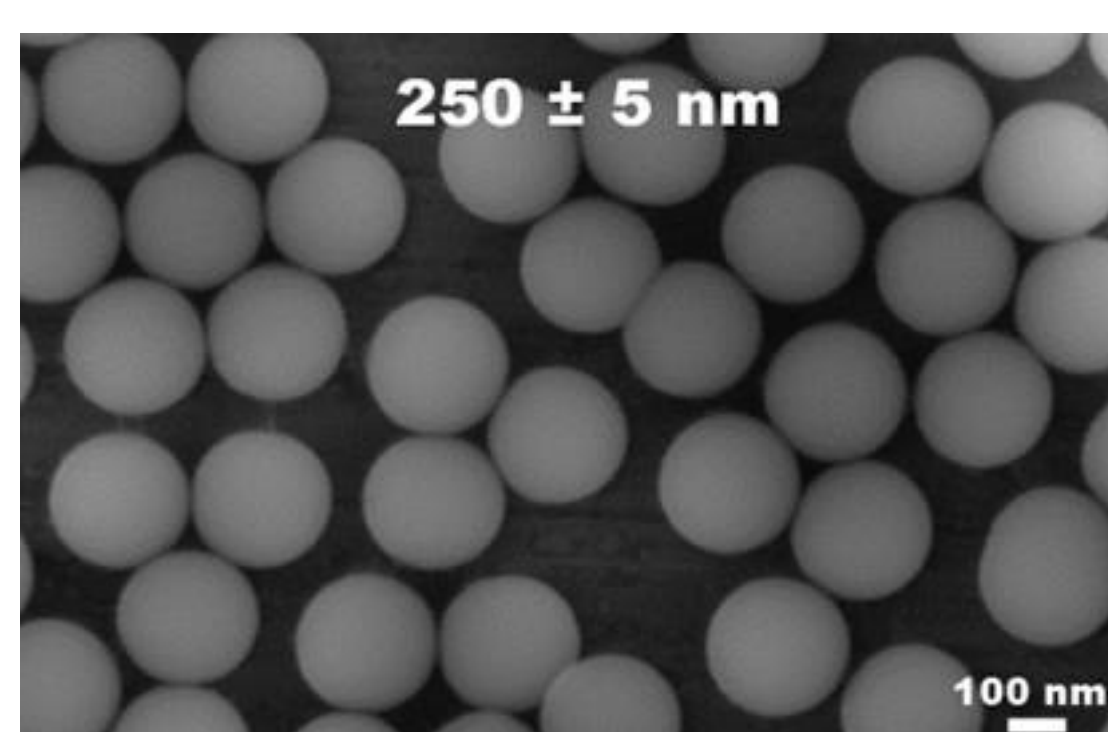
- Particle
- Size
- Concentration
- Phenotyping

## Sample Analysis



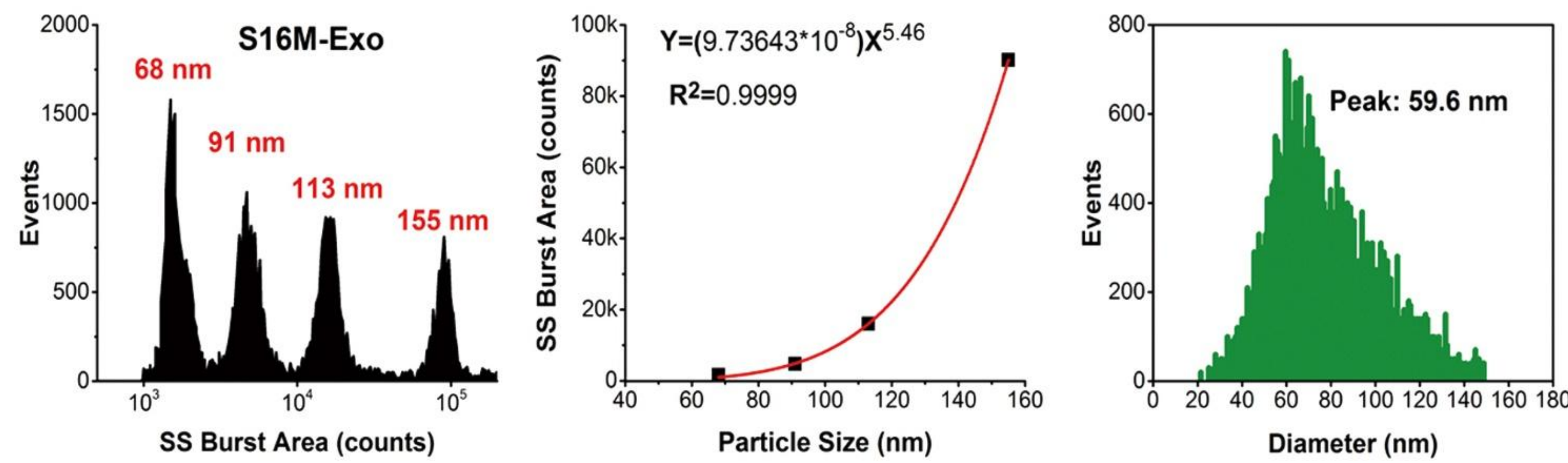
- Chemotherapeutics
- Nucleic Acid
- Imaging Agent (FC, PET, MRI)
- Targeting Ligand

## Silica Nanoparticles & Quality Control Beads



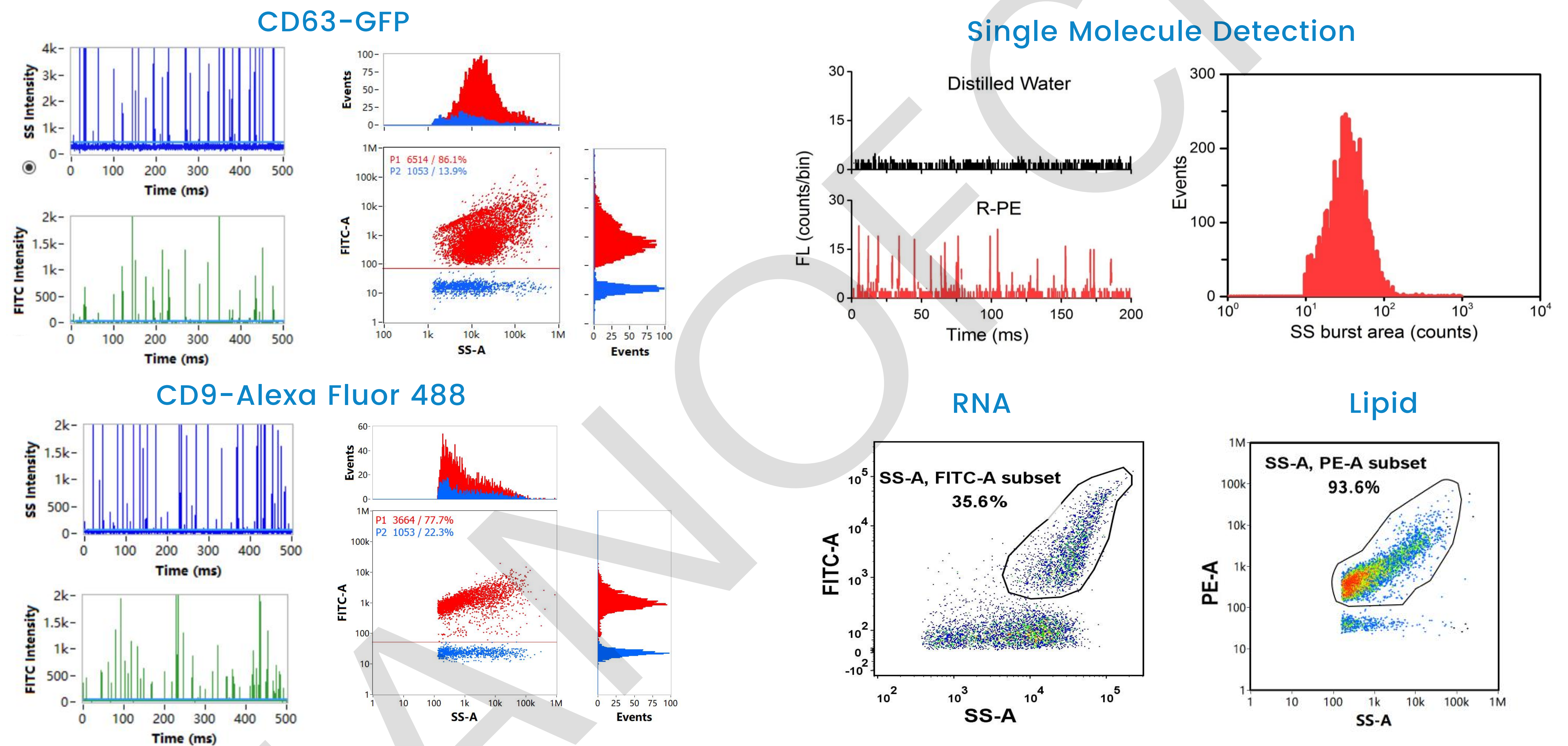
NanoFCM Quality Control Nanospheres Series with accurate size and concentration are designed both for adjusting the alignment of NanoAnalyzers and for use as an internal or external standard for sizing and concentration measurement.

### High-Resolution Size Distribution Analysis

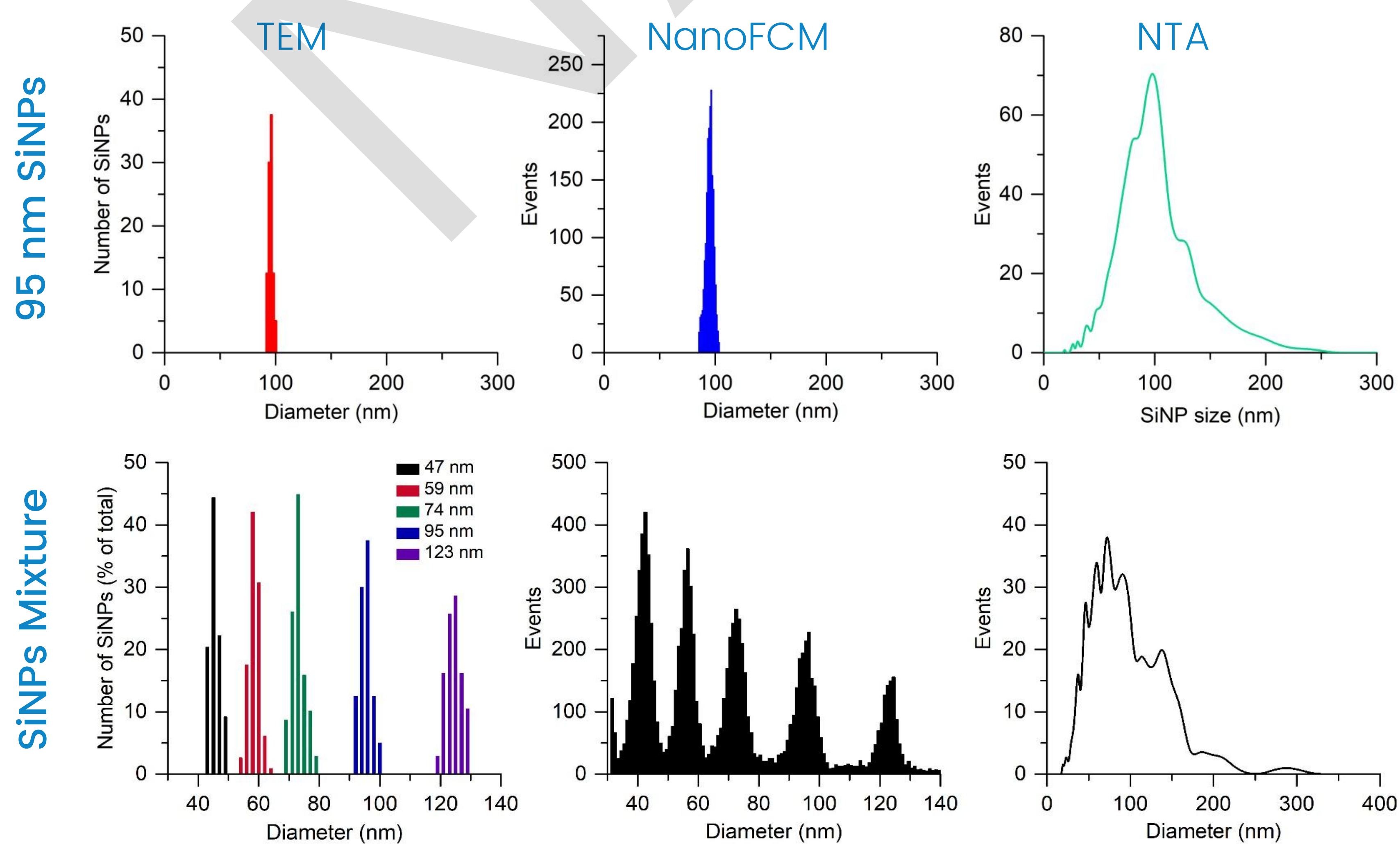


Employing S16M-Exo (NanoFCM) as size standards, a calibration curve will be constructed between the particle size and side scatter intensity, the SS intensity of each EV particle could be converted to size. The size distribution of EV matches well with that acquired from Cryo-TEM.

### EVs Phenotyping at Single Particle Level



### Comparison with First-Generation Techniques



For silica NPs of 95 nm, the size distribution measured by NanoFCM is consistent with TEM, that is, sharp histogram centered at 95 nm is achieved. In contrast, a broad size distribution (20-250 nm) containing massive false signals is observed by NTA, originating from the lack of both sensitivity and resolution.

The ultrahigh sensitivity and resolution of NanoFCM is perfectly illustrated by the measurement of silica NPs with mixed sizes (47, 59, 74, 95, and 123 nm). NanoFCM offers comparable results as TEM, far beyond the best result obtained by NTA. NanoFCM enables users to accomplish complicate analysis for heterogeneous samples without losing details.

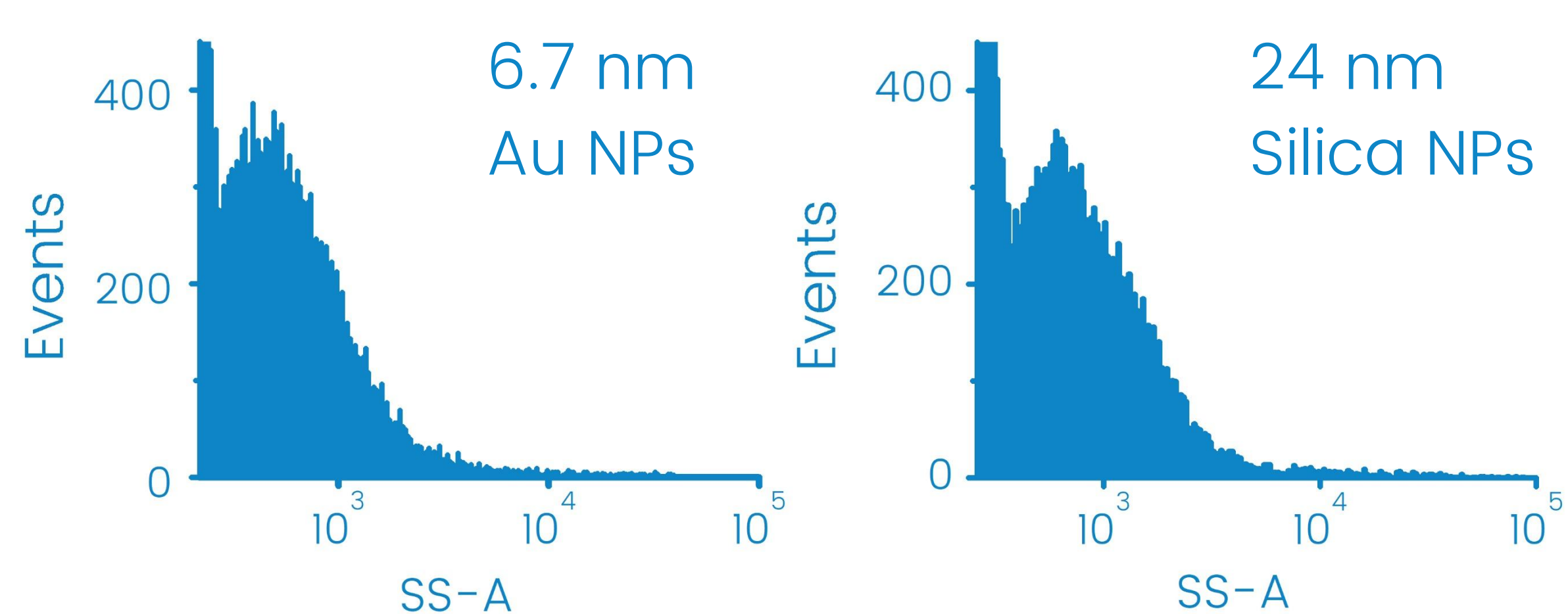


# Performance



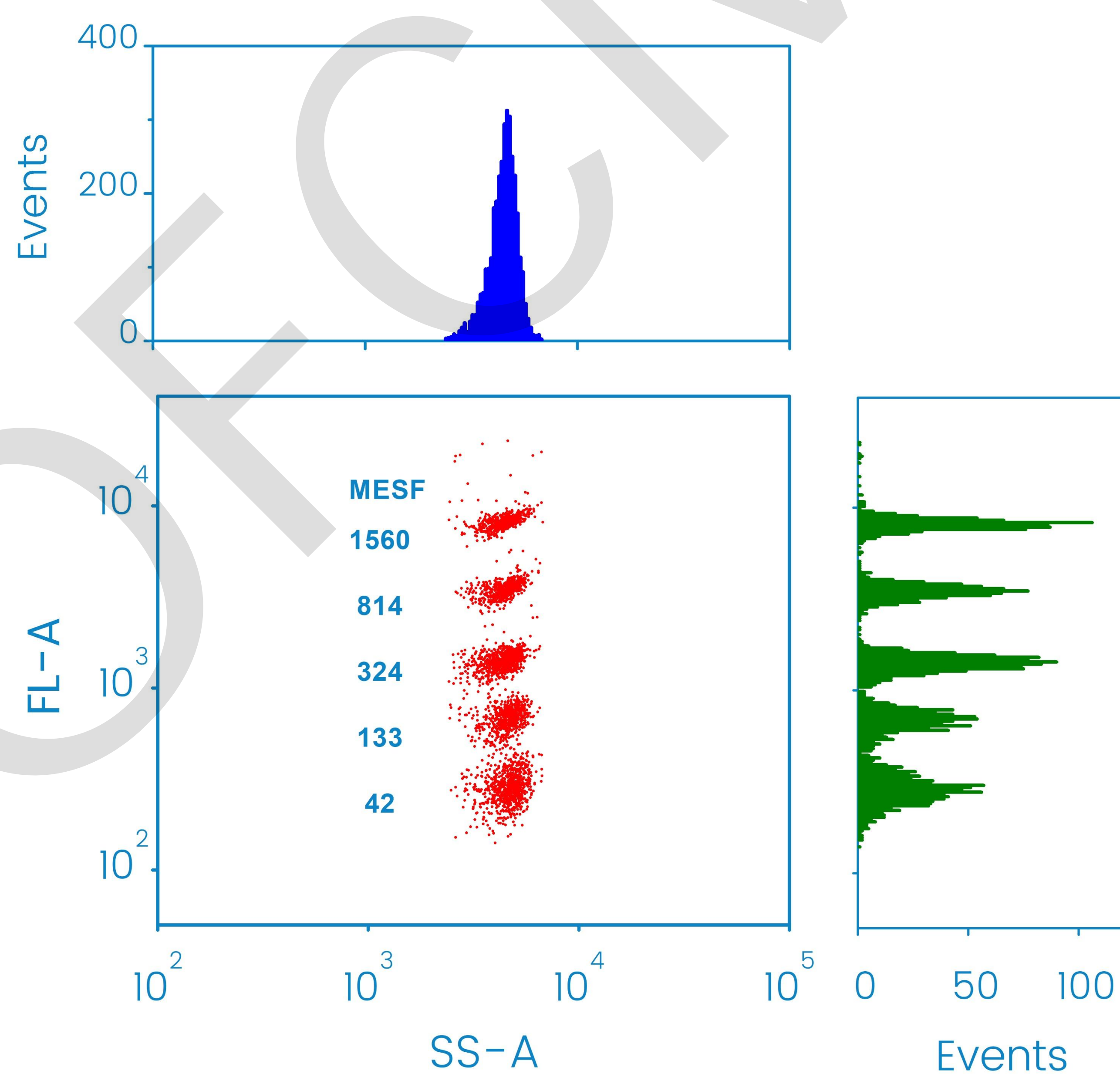
- SSC Sensitivity < 30 nm NPs
- SSC Resolution 40/50 nm NPs
- Fluorescence Sensitivity AF488 < 10
- Fluorescence Resolution 42/133 ERF
- Particle Size 7-1000 nm
- Sample Acquisition Rate 10,000 events/min

## Sensitivity for Scattering Detection (7-1000 nm)

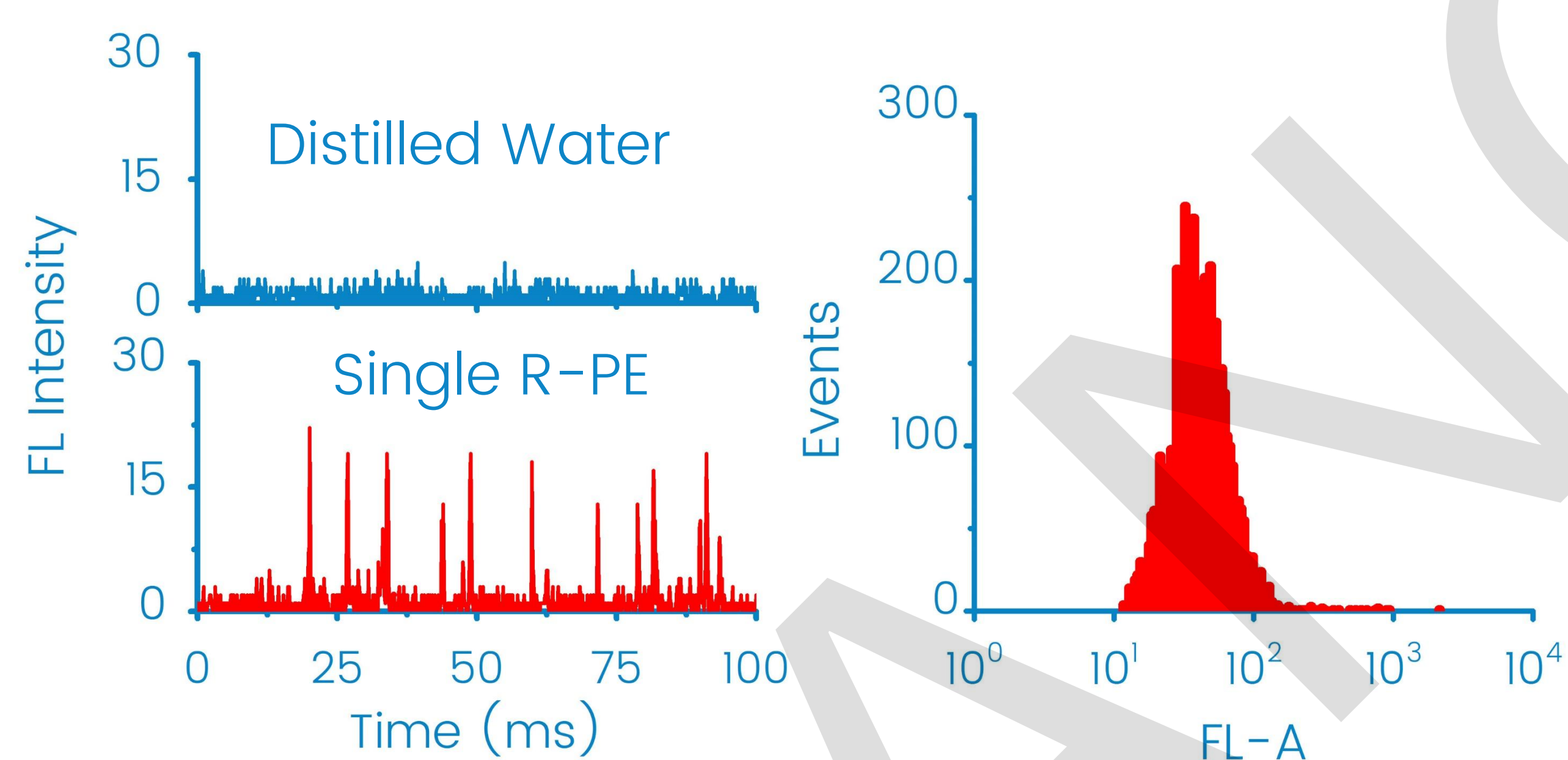


## Resolution for Fluorescence Detection

Mixture of 212 nm Fluorescent Silica NPs

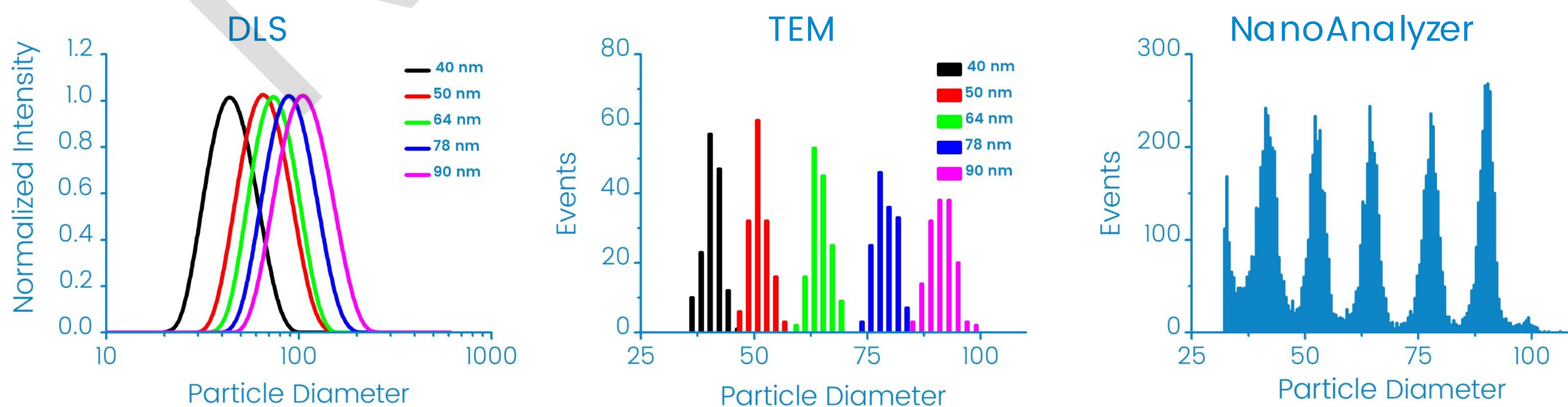


## Sensitivity for Fluorescence Detection



## Resolution Comparable to Electron Microscopy

Mixture of 40, 50, 64, 78 and 90 nm Silica Nanoparticles



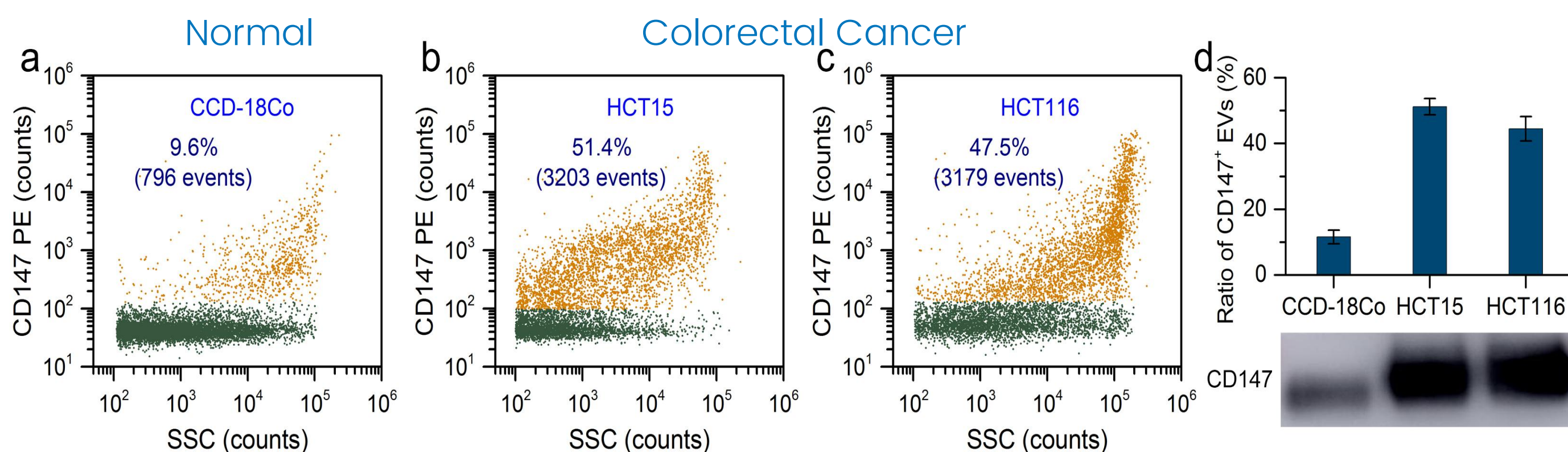
Unmatched Performance • Superior Sensitivity • Innovation for NanoWorld



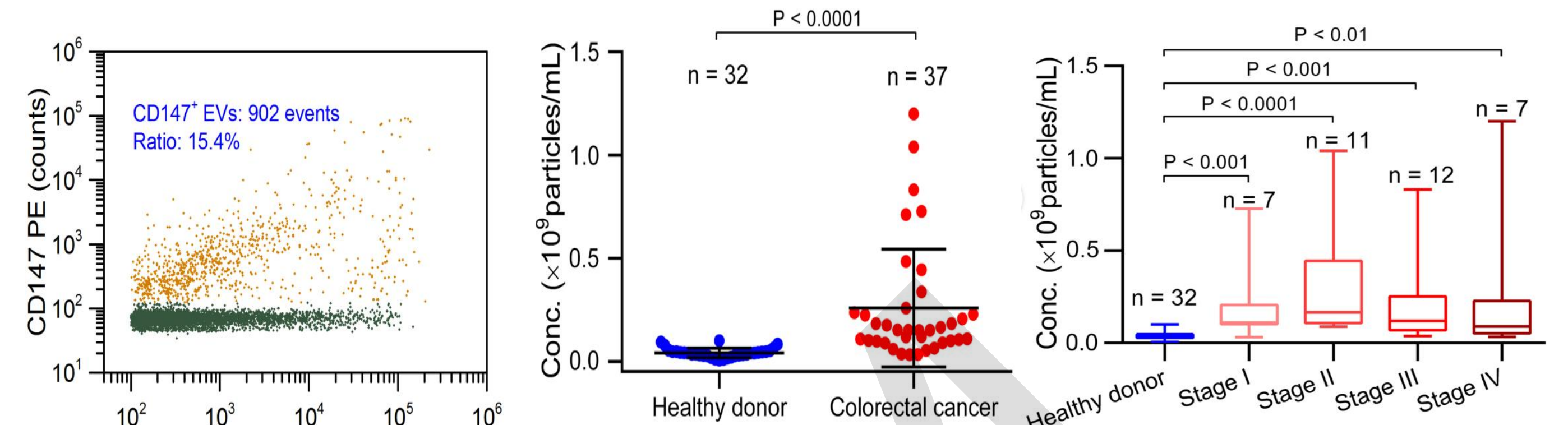
# Application of Extracellular Vesicles

## Early Diagnosis of Cancer

### CD147+ EVs in Colorectal Cancer Cell Lines



### CD147+ EVs in Clinical Blood Samples

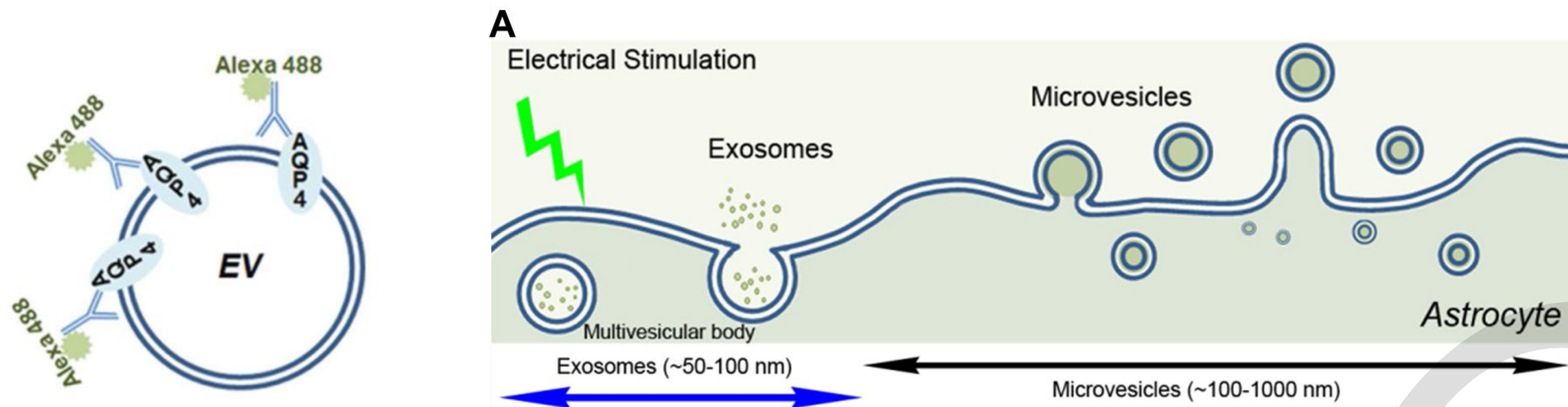


CD147 expression is analyzed quantitatively at single EV level by NanoFCM. Moreover, NanoFCM allows correlating the protein abundance with vesicle size at the single-particle level, CD147-positive EVs exhibit a range of sizes depending on their cell origin.

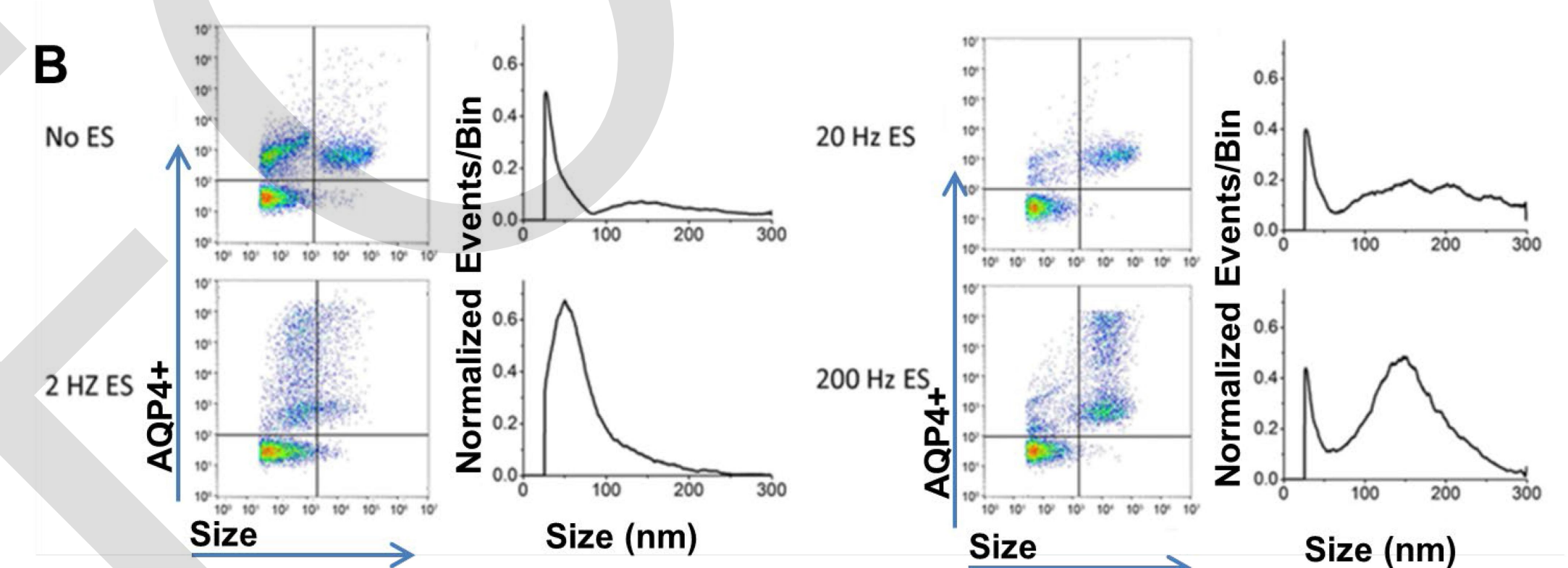
*ACS Nano* 2018, 12, 671-680

NanoFCM is able to identify the elevated level of CD147 positive EVs for patients at all the cancer stages, even stage I. Moreover, this strategy can be used to track the level of CD147 expression after surgical resection.

## Programmable Modulation for EVs

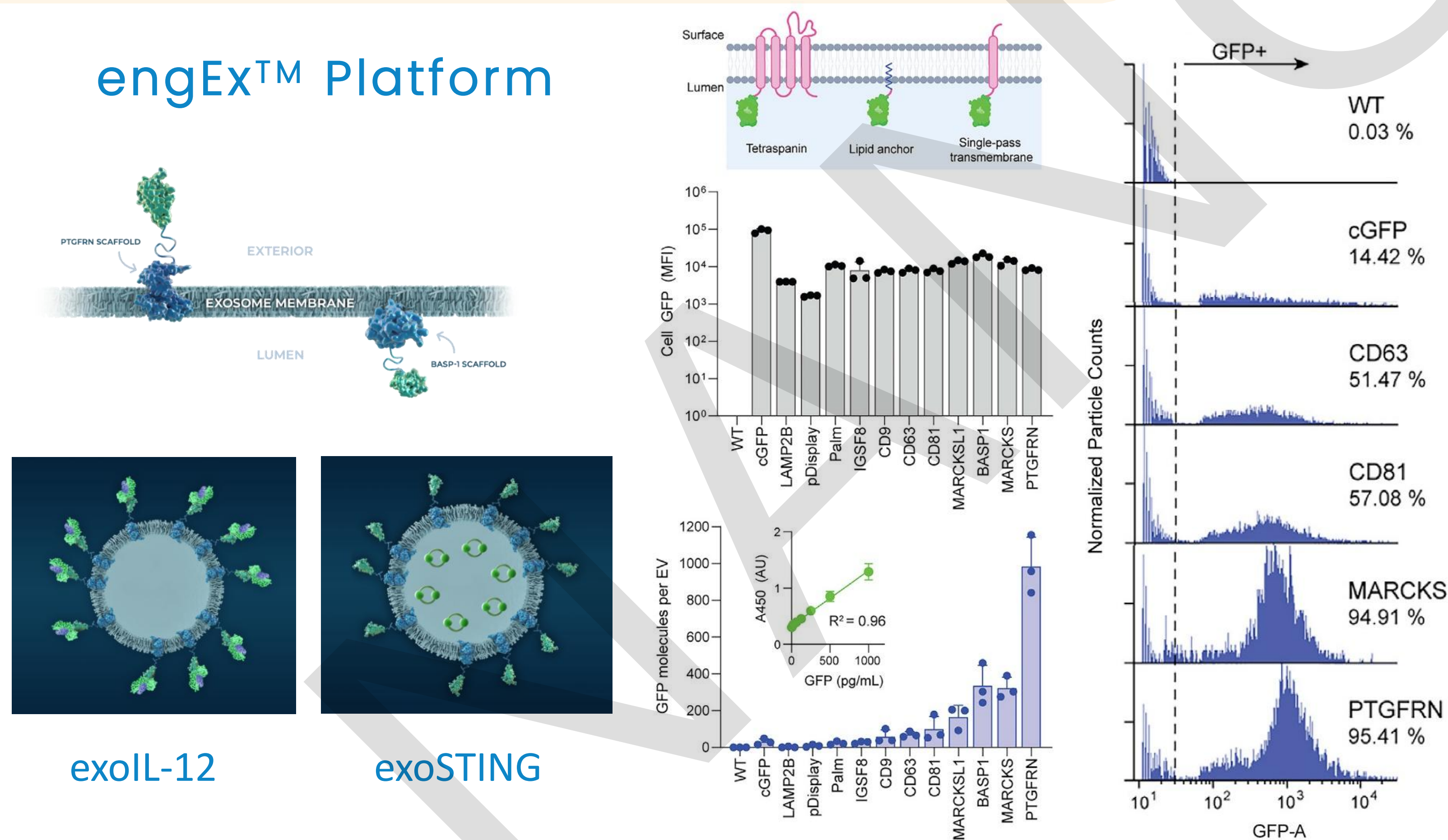


### Distributions of EVs after Electrical Stimulation



DOI: <http://dx.doi.org/10.1101/566448>

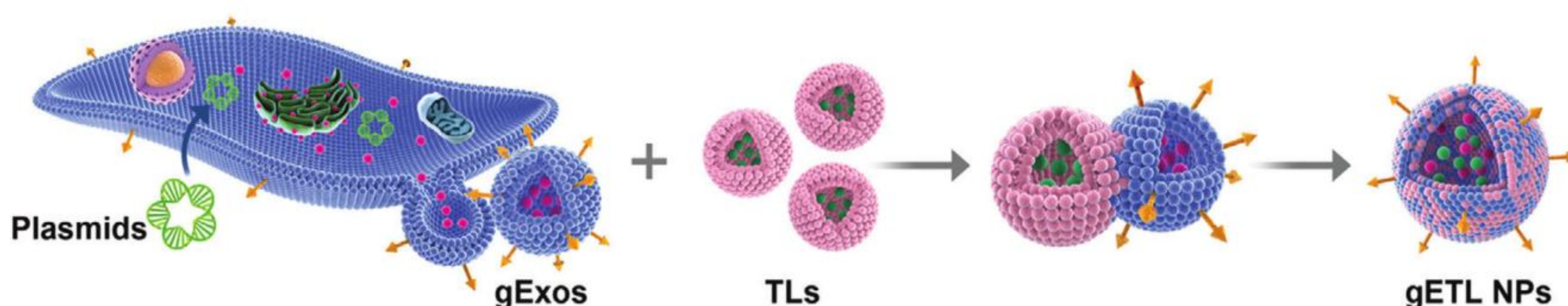
## Identification of A Versatile Platform



engEx™ Platform is based on the discovery of two scaffold proteins. Flow cytometry and ELISA were used to measure cellular and exosome-associated protein expression, however, these methods fail to determine whether overexpressed scaffold proteins were uniformly distributed among EVs or enriched in subsets. By analyzing EVs at single particle level, the data suggest that overexpression of the candidate scaffolds results in abundant, uniform distribution across EVs.

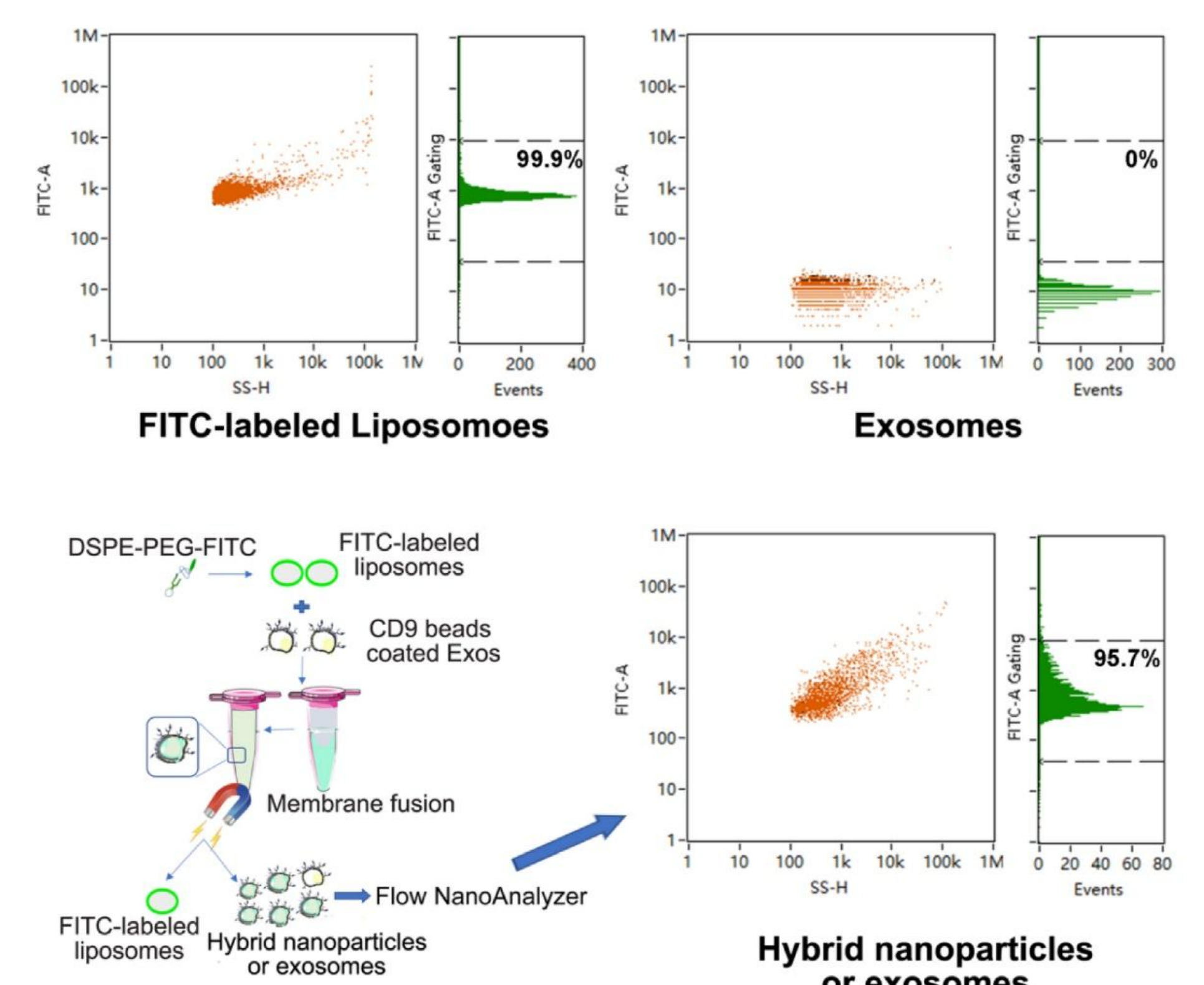
*Mol Ther.* 2021 May 5; 29(5):1729-1743

## Exosome-Liposome Hybrid Nanoparticles



Thermosensitive liposomes were fused with genetically engineered exosomes, the resulting exosome-liposome hybrid NPs display CD47 on the surface and bear thermosensitive agents inside. NanoFCM allows the analysis of liposomes and exosomes at single particle level, and the fusion efficiency is also determined.

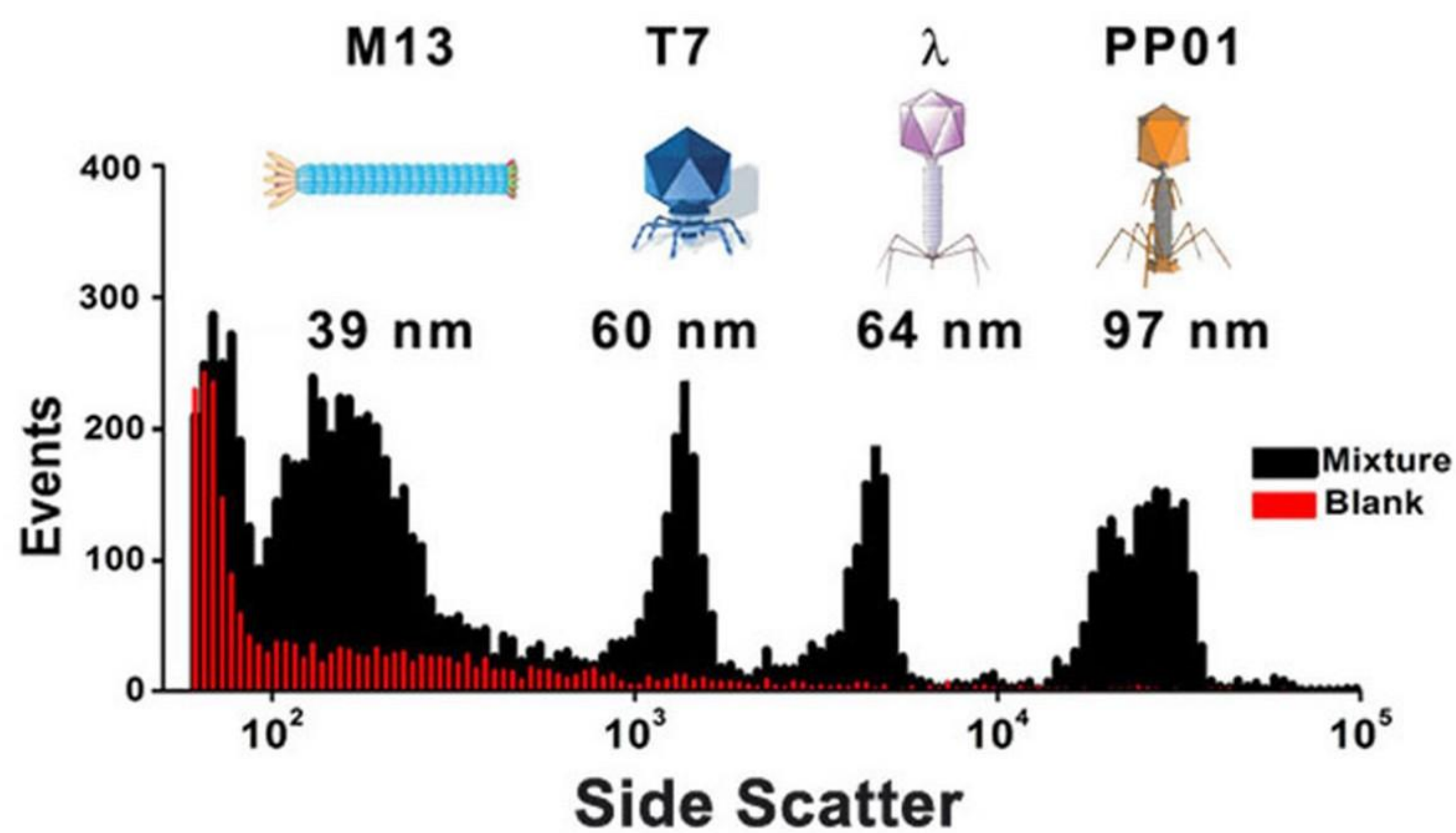
*Adv. Sci.* 2020, 7, 2000515



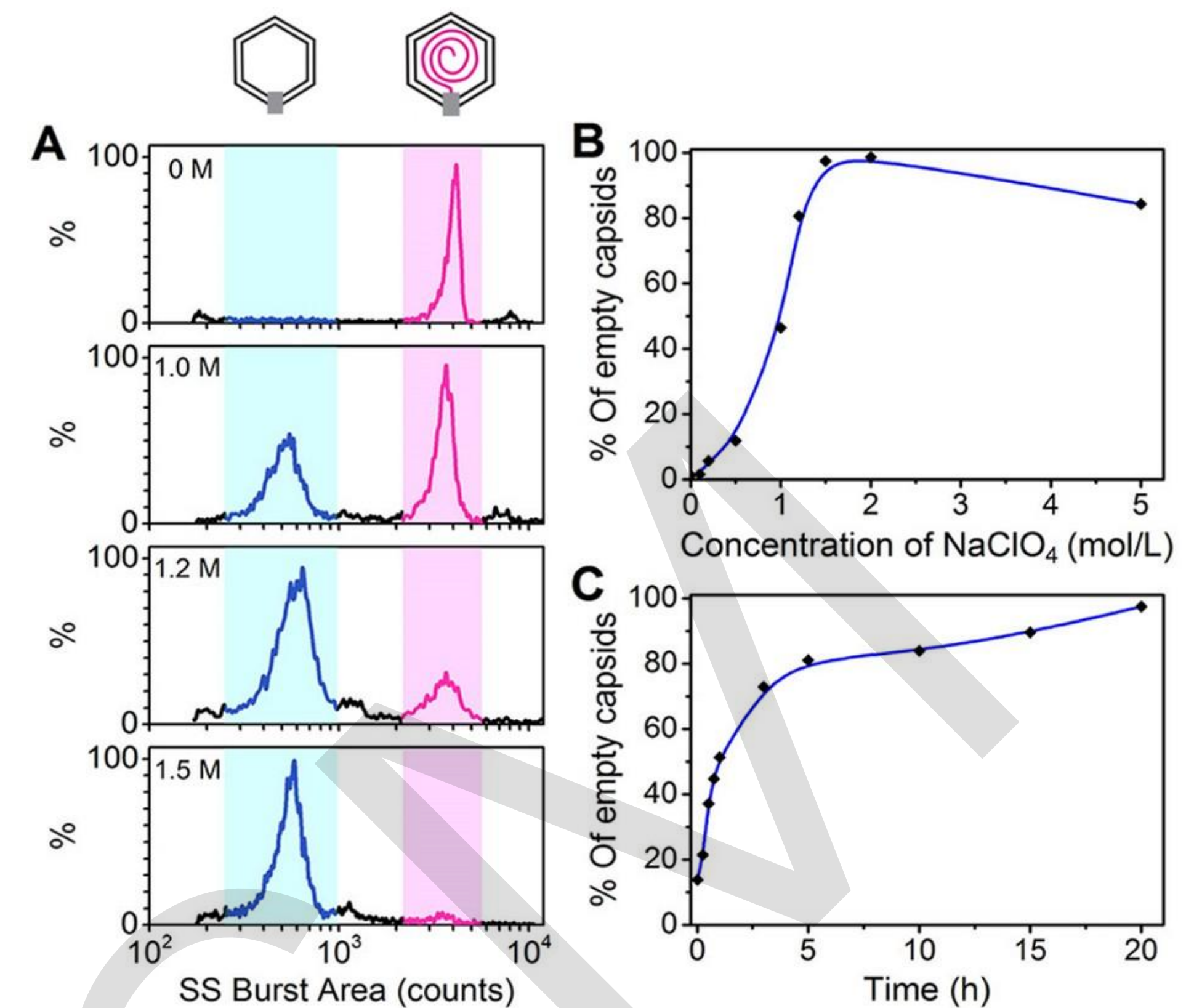


# Application of Viruses

## Size Differentiation of A Virus Mixture

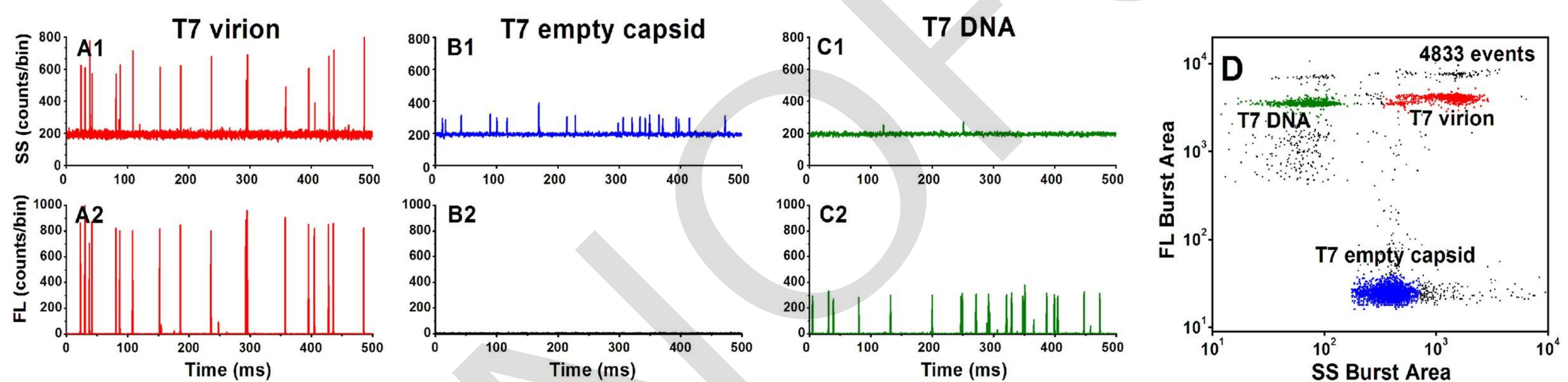


## Dynamic Monitoring of Viral DNA Release



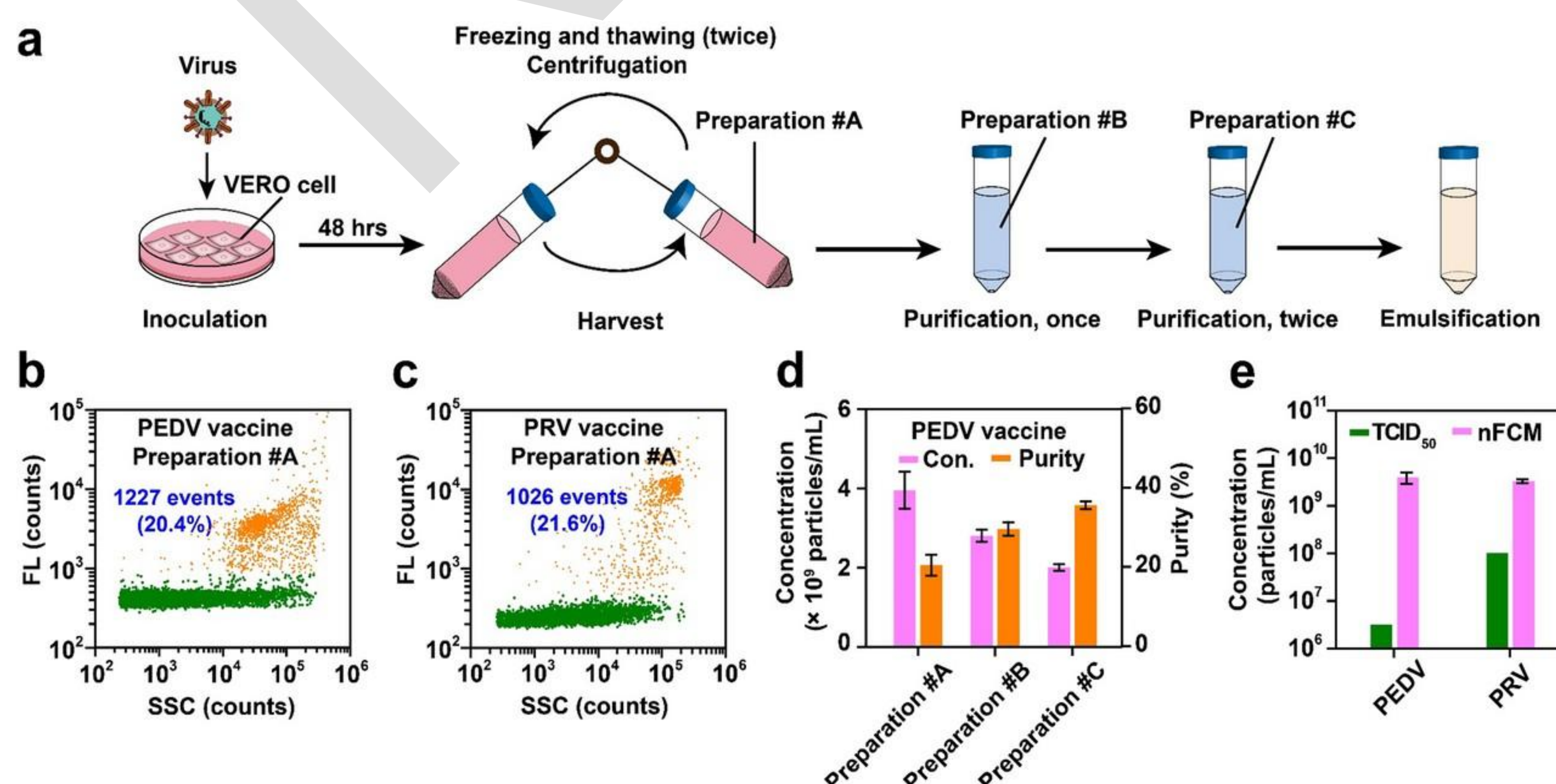
Angew. Chem. Int. Ed. 2016, 55, 10239-10243

## Virus-Mediated Drug Delivery Vectors



Viruses can be considered as nature's nanotechnology, serving as nanoscale vehicles for the delivery of nucleic-acid cargos into host cells. Through nucleic acid staining, nano-flow cytometry is able to discriminate individual virions from empty viral capsid and free viral DNA, which is helpful to determine the drug loading efficiency, the loading content, and the effective ratio.

## Characterisation of Virus-Based Vaccines



The main steps of producing virus-based vaccines include virus inoculation, harvest, purification, and emulsification. Through nucleic acid staining, NanoFCM allows the fast evaluation of virus products at different steps. With further purification upon precipitation, the concentration of intact viruses decreases, while the purity increases.

Angew. Chem. Int. Ed. 2021, 60, 9351-9356



# Technical Specifications

| NanoAnalyzer         | Model Number                         | N30  |        | U30          |              |
|----------------------|--------------------------------------|--|--------|--------------|--------------|
| Detectors            | Laser                                | 488 nm   | 528 nm | 488 & 638 nm | 528 & 638 nm |
|                      | SSC                                  | SPCM   | SPCM   | SPCM         | SPCM         |
|                      | 525/40 nm                            | SPCM   | —      | SPCM         | —            |
|                      | 580/40 nm                            | —  | SPCM   | —            | SPCM         |
|                      | >650 nm                              | SPCM   | SPCM   | SPCM         | SPCM         |
|                      | SPCM: Single Photon Counting Module; |  |        |              |              |
| Optics               | Laser Configuration                  | 6 μm × 24 μm elliptical spot                               |        |              |              |
|                      | Flow Cell                            | 250 × 250 μm rectangular quartz flow cell                  |        |              |              |
|                      | SSC Sensitivity                      | < 30 nm  |        |              |              |
|                      | SSC Resolution                       | 40/50 nm   |        |              |              |
|                      | Particle Size                        | 7 - 1000 nm  |        |              |              |
|                      | Fluorescence Sensitivity             | AF488 <10, PE< 1   |        |              |              |
|                      | Fluorescence Resolution              | 42/133 ERF   |        |              |              |
|                      | Filters                              | User Exchangeable  |        |              |              |
| Fluidics             | Sample Acquisition Rate              | 10,000 events/min  |        |              |              |
|                      | Sample Flow Rate                     | 2 - 60 nL/min  |        |              |              |
|                      | Sheath Flow Rate                     | 10 - 40 μL/min   |        |              |              |
|                      | Sample Volume                        | 10 - 100 μL  |        |              |              |
|                      | Fluid Container Capacity             | 1 L sheath, 1 L waste, 50 mL cleaning                      |        |              |              |
|                      | Fluidics Maintenance                 | Automated startup, cleaning, decontamination and shutdown  |        |              |              |
| Data Processing      | Parameters                           | Peak Height, Area and Width for all Channels               |        |              |              |
|                      | Output Data Files                    | NFA; FCS 3.0   |        |              |              |
|                      | Software                             | NF Profession 3.0  |        |              |              |
| Sampling             | Manual Sample Loading                | 0.6 mL EP tube   |        |              |              |
| Operating Conditions | Instrument Dimension (W × D × H)     | 50.6 cm × 34.6 cm × 29.0 cm<br>19.9 in × 13.6 in × 11.4 in |        |              |              |
|                      | Instrument Weight                    | 51.6 lb (23.4 kg)  |        |              |              |
|                      | Power Requirements                   | 110-240 VAC, 50-60 Hz                                      |        |              |              |
|                      | Environment Requirements             | Temperature: 15-35°C; Relative Humidity: 80% maximum       |        |              |              |

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