



# Next Generation Digital Microfluidic Technology for Efficient & Automated Transfection Experiments

**Do Jin Im**

ATP (Advanced Transport Phenomena) Lab

Department of Chemical Engineering

Pukyong National University, Korea

## Accidental Observation in 2005

A 300 nL water droplet (1 mm diameter) under 3 kV/cm



100 cSt silicone oil

## Electrophoresis of a **C**harged **D**roplet (**ECD**)

- Simple and Straightforward

- Easy to design electrodes
- High degree of freedom in chip design

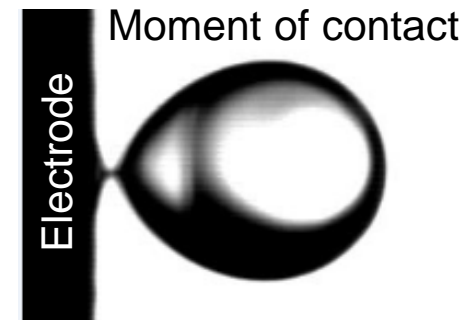
$$\mathbf{F} = Q\mathbf{E}$$

- Minimal solid surface contact

- Suitable for biological applications
- High reliability microfluidic device

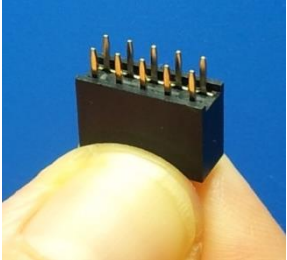
- Fast droplet velocity

- Fast and consistent droplet actuation
- High analytical efficiency



2011 National Futuristic Technology:  
NTIS NO. 1345100479

# ECD based Digital Microfluidic System



Pin header socket



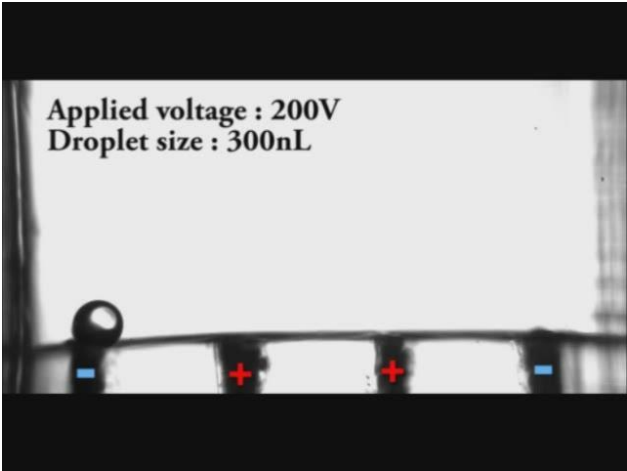
Assembly of electrodes



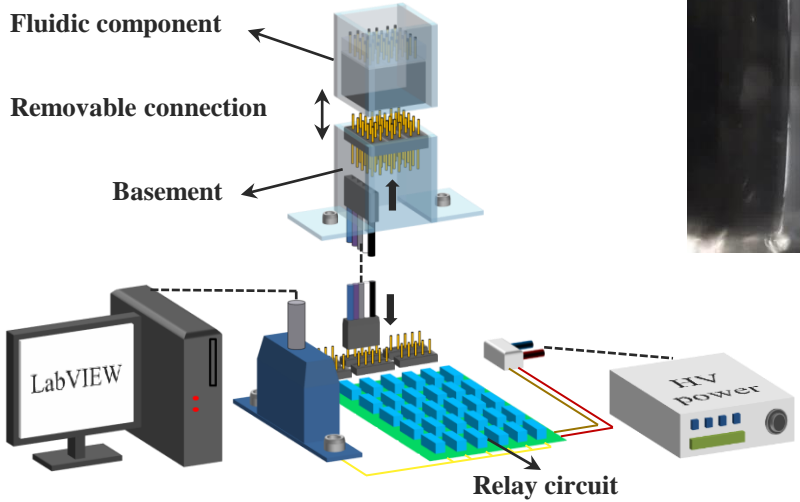
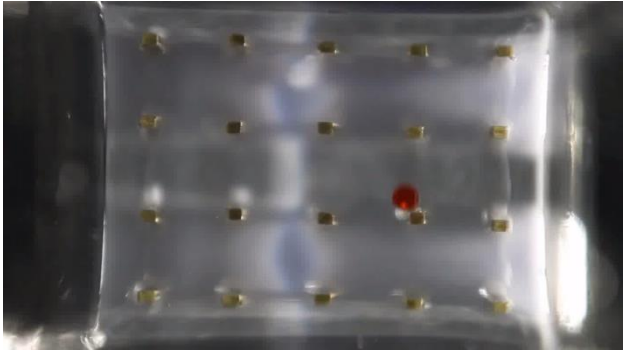
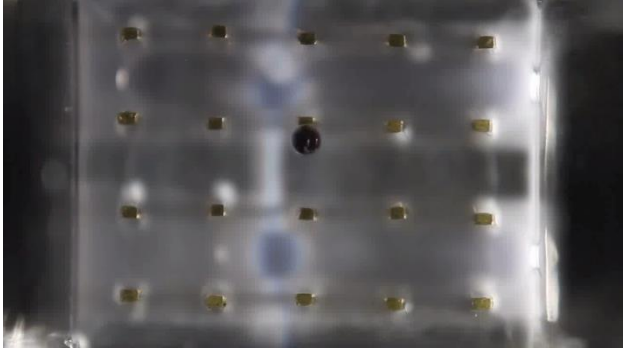
Fluidic component



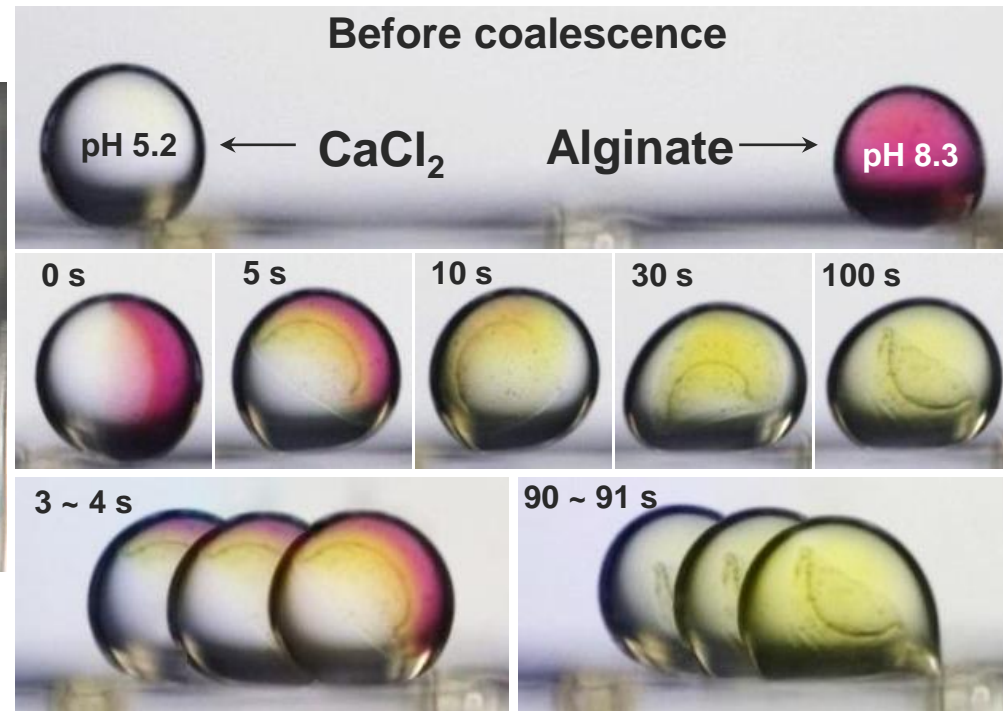
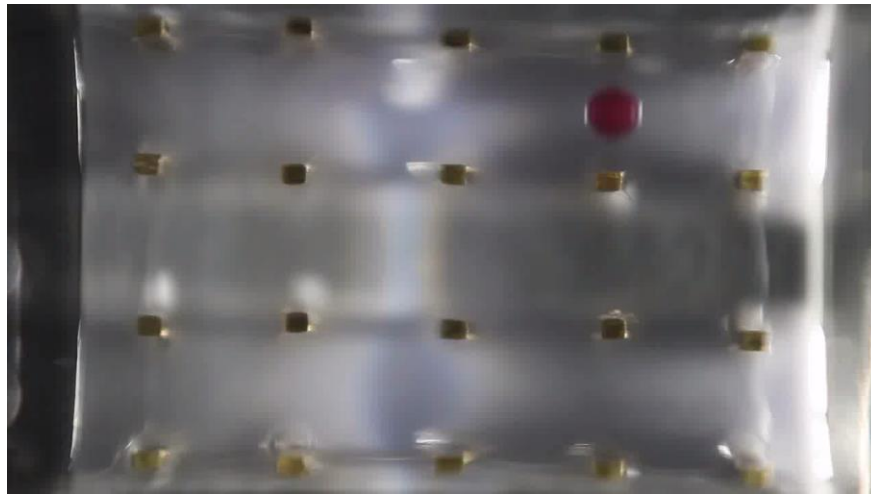
## 1D Translation



## 2D Manipulation



## Independent controls on the two droplets



✓ The interfacial tension changes as the gelation reaction proceeds!

*Anal. Chem.* 2013, **85**, 4038–4044



## ✓ Synthetic DNA delivery systems

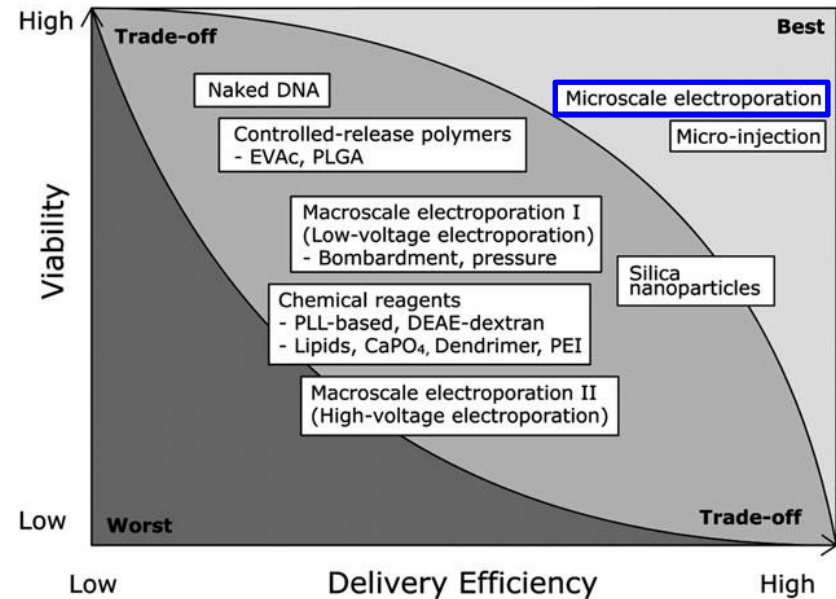
- Non-viral DNA delivery (transfection)
- Mechanical, Electrical, Chemical

## ✓ Electroporation (EP)

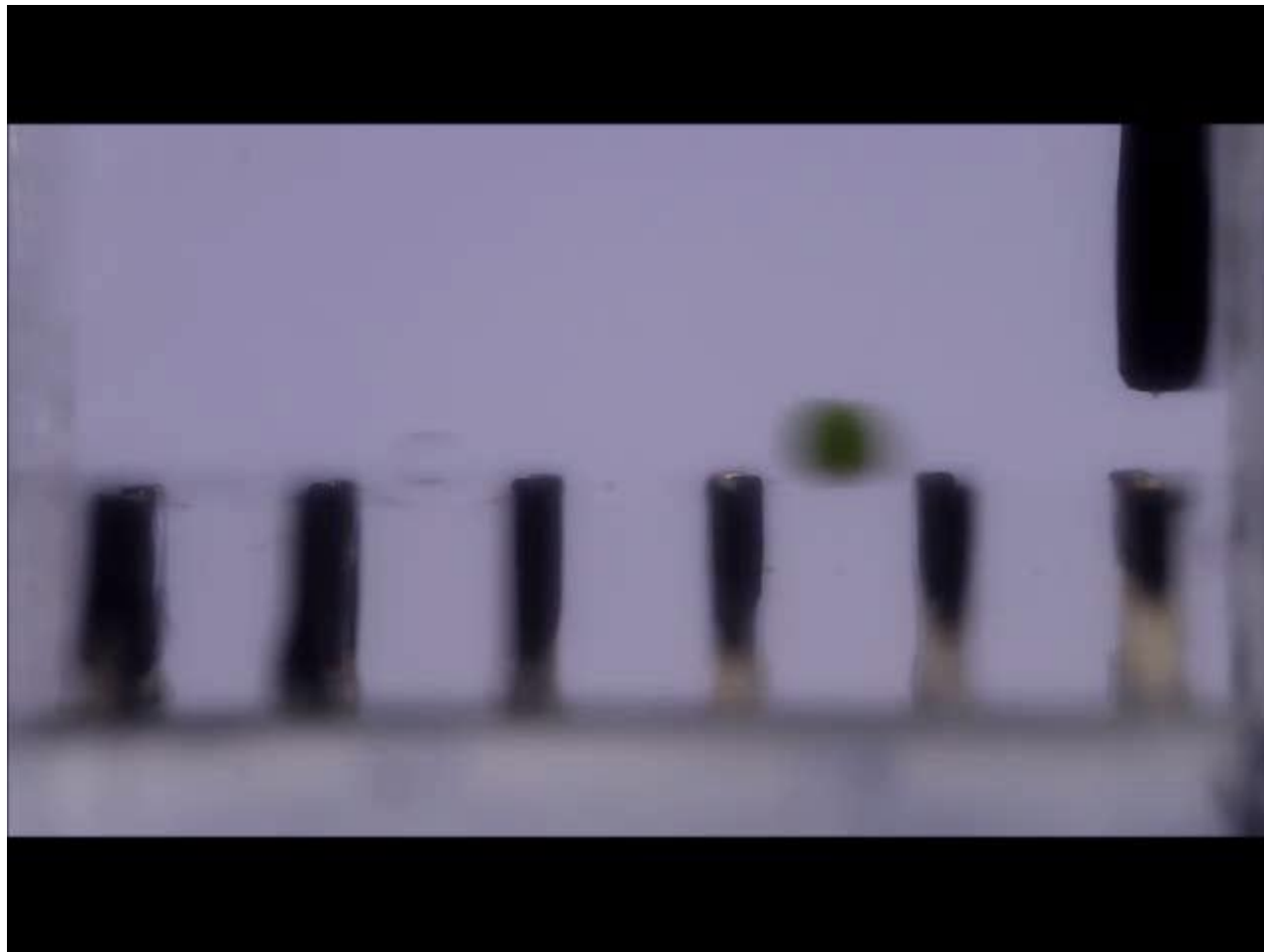
- Widely used, Good productivity
- Drawbacks of conventional EP  
: **Low Viability, Contamination**

## ✓ Microfluidic Electroporation

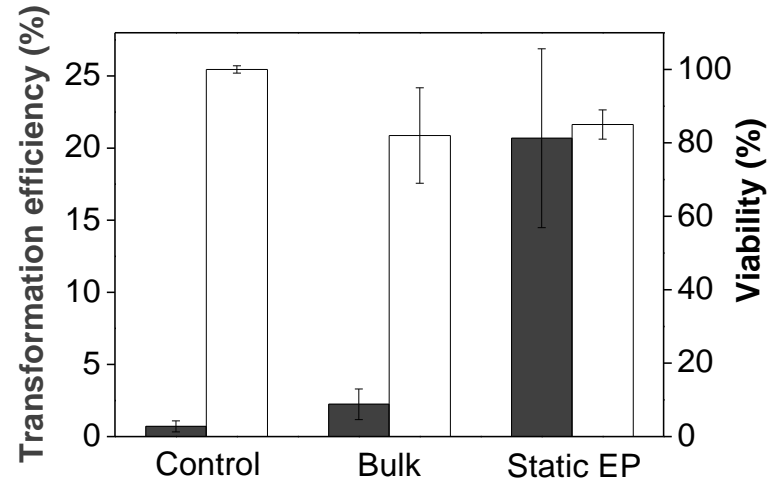
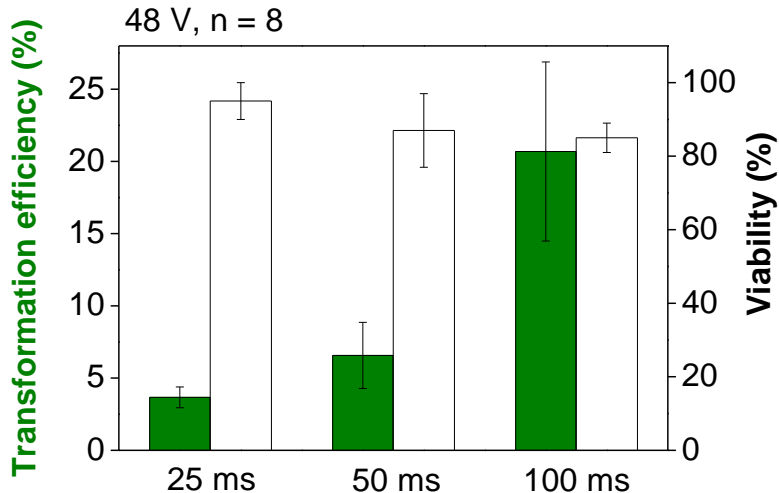
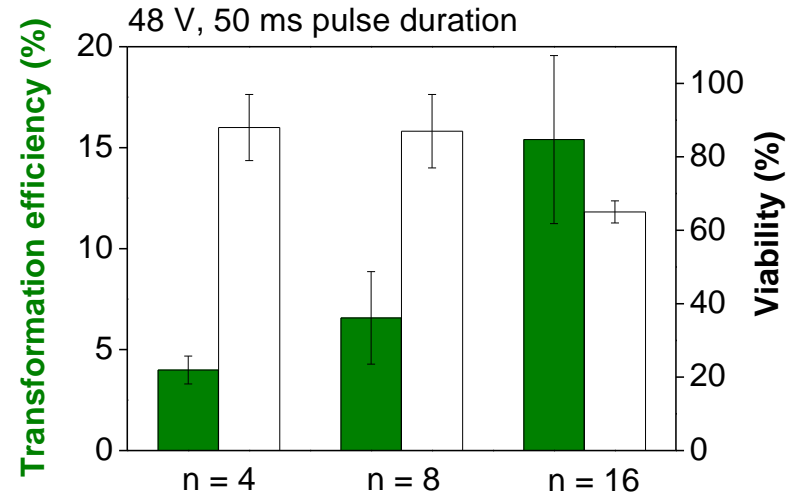
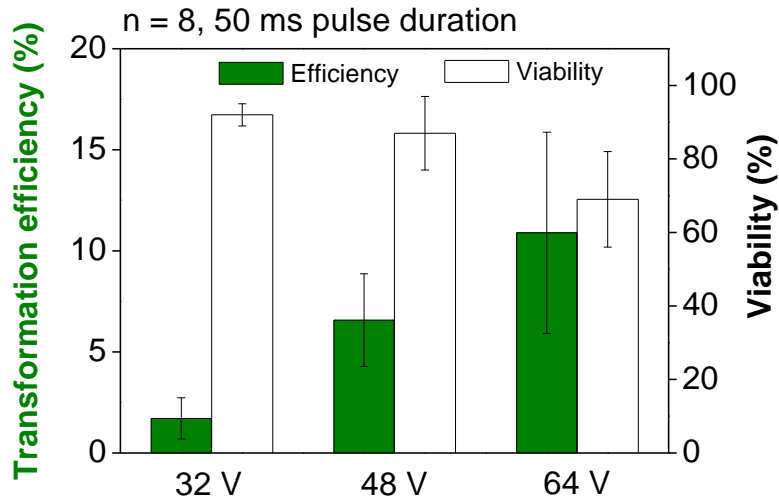
- High cell viability and delivery efficiency
- Drawbacks of current microfluidic EP systems  
: **Lack of usability, Low productivity**



*Integr. Biol.*, **2009**, 1, 242–251



# Algae transformation by Digital EP





# Integrated Cell Engineering Platform



**Input port/On-demand Dispensing Unit**

**Output port/ Splitting Unit**

**Optimal Electrode Design**

**Display Panel**

**HV CMOS Relay**

**HV Boost Chip : 5 V → 500 V**

**Rechargeable Battery**

**USB Port for External Power & Circuit Control**

**External USB Power/Battery Switching Module**

**Analyst 2013, 138, 7362–7368**

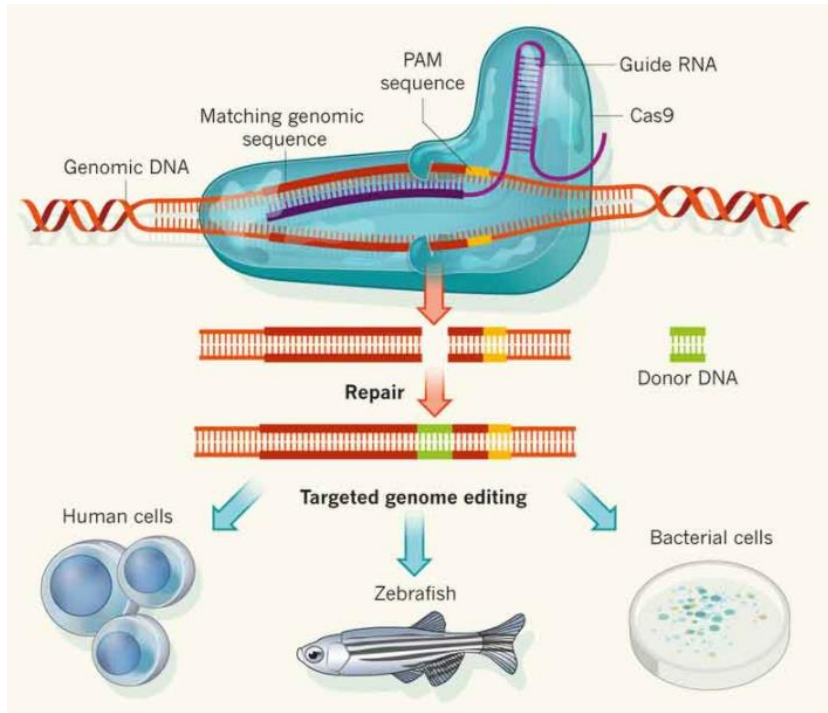
**Figure 1: Optimal Electrode Design**

The graph plots the ratio of the volume of the electrode to the volume of the droplet ( $Q/Q_{th}$ ) on the y-axis (ranging from 0 to 6) against the ratio of the electrode radius to the droplet radius (Electrode R/Droplet R) on the x-axis (logarithmic scale from 0.1 to 100). The data points are categorized by volume (0.1, 0.3, 0.7, 1.0  $\mu\text{L}$ ) and electrode type (N for needle, Pin for pin, Planar for planar). The legend indicates: 0.1  $\mu\text{L}$  (N) [solid blue line], 0.3  $\mu\text{L}$  (N) [dashed red line], 0.7  $\mu\text{L}$  (N) [dotted green line], 1.0  $\mu\text{L}$  (N) [dash-dot black line], 0.1  $\mu\text{L}$  (Pin) [solid blue square], 0.3  $\mu\text{L}$  (Pin) [dashed red circle], 0.7  $\mu\text{L}$  (Pin) [dotted green triangle], 1.0  $\mu\text{L}$  (Pin) [dash-dot black diamond], 0.1  $\mu\text{L}$  (Planar) [solid blue square], 0.3  $\mu\text{L}$  (Planar) [dashed red circle], 0.7  $\mu\text{L}$  (Planar) [dotted green triangle], 1.0  $\mu\text{L}$  (Planar) [dash-dot black diamond]. The graph shows that the optimal design for a given volume and electrode type is achieved at a specific Electrode R/Droplet R ratio, with the ratio generally increasing as the volume and electrode size increase.

# Transfection Technology Trend



## CRISPR-CAS9 Gene Editing



Guid RNA + CAS9 Protein  
→ Delivered by **Electroporation!**

\* CRISPR (Clustered Regularly Interspaced Short Palindromic Repeats)



## Cell Engineering & Culture in a Droplet

### ✓ Merits of droplet-in-oil environment

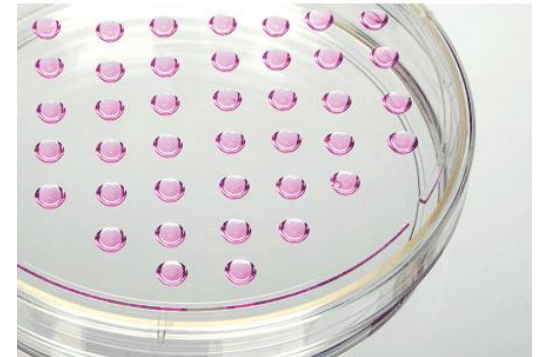
- Small resource consumption
- Low contamination
- Increased gas exchange

### ✓ 3D cell culture

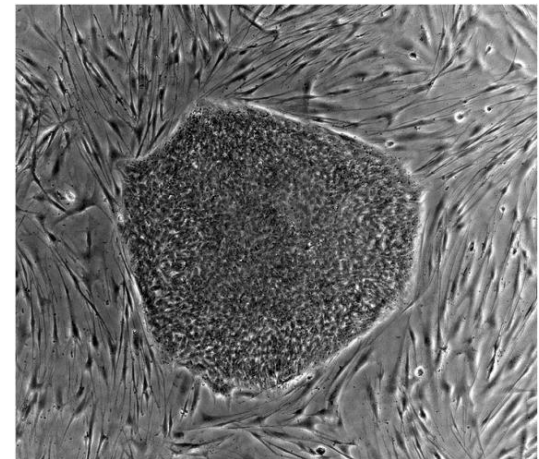
- New paradigm of cell culture
- Close to real biological system

### ✓ Digital EP for cancer & stem cell research

- PI - NRF Project (No. 2013R1A1A2010483)
- Korean, PCT patents are pending



Hanging drop culture of embryonic stem cells  
*Nature Protocols* 2011, **6** (7) Cover Page



Stem cell culture

## Microfluidics + Electronics + IT Tech

### → Bio/Chemical Digital Microfluidic Applications

