Controlled Olfactory Delivery using Magnetophoretic Guidance

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Introduction: Direct nose-to-brain drug delivery has many advantages over the intravenous delivery approach. The major challenge in achieving clinically significant nose-to-brain drug delivery is delivering drugs to the olfactory region. We propose to increase the olfactory delivery efficiency by using magnetic guidance.

Limitations of normal magnetic control:

- 1. Field and gradient decay quickly in space.
- 2. Unstable: too weak at far, too large when near.

New thought and Rationale:

A nonmagnetic particle within a ferrofluid will experience a repulsive force; Stable control

Analogy: a lighter object experience a buoyance force in a heavier fluid.

The idea is inspired by several seemingly irrelevant phenomena when they met and fused together. Figure 1 speaks a thousand words on how they can connect.



Figure 1. Building Magnetic track for olfactory drug delivery.

Study Design:

- 1. Test magnetic track in a 2-D channel.
- 2. Test magnetic track in human nose cavity.

Methods:

- 1. Four COMSOL modules used: fluids, electromagnetic, particle tracking, optimization.
- 2. Parameter of interest: magnet layout and strength, drug resale position, particle diameter, geometrical complexity.
- 3. Nose model reconstructed from MRI scans.
- 4. Ferromagnetic nanoparticles suspending in the nasal cavity to form the Ferro fluids ($\varepsilon r > 1$).

Results:

2-D channel:

- 1. With magnetic guidance, particles travel though L-shaped channel contact-free.
- 2. All deposit at the corner without magnet.

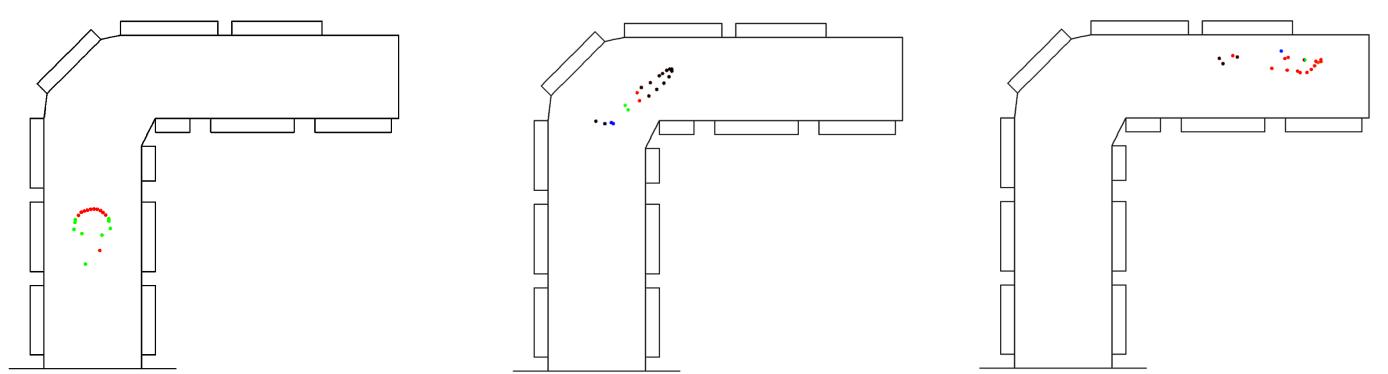


Figure 2. Contact-free particle transport in 2-D channel

Image-based nose model:

✓ Magnetic guidance improves olfactory dose

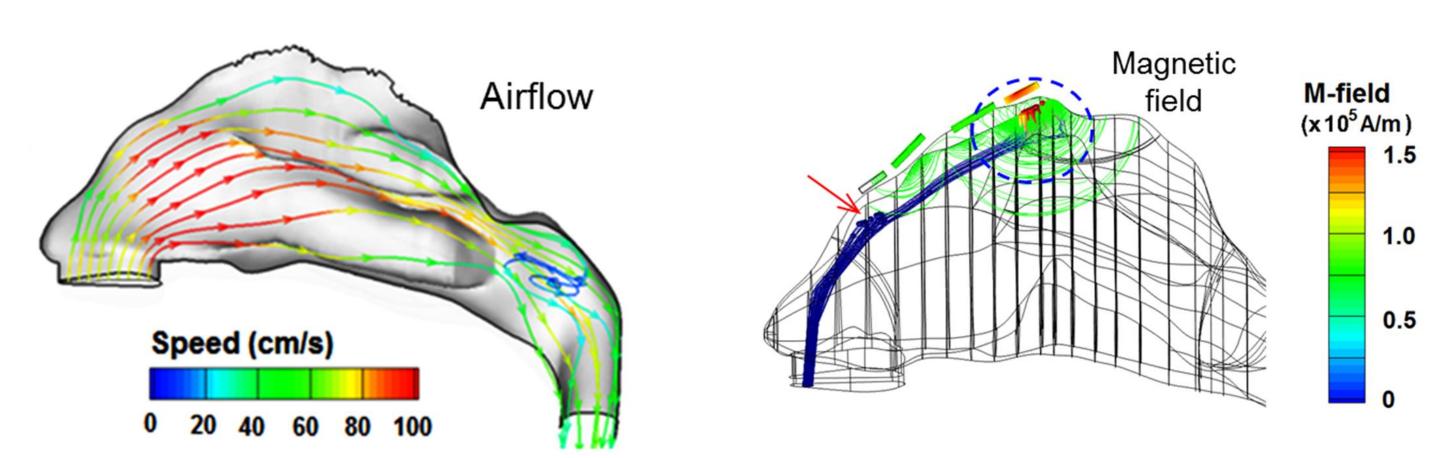


Figure 3. Airflow and Magnetic transport

✓ Optimization conducted on point-release

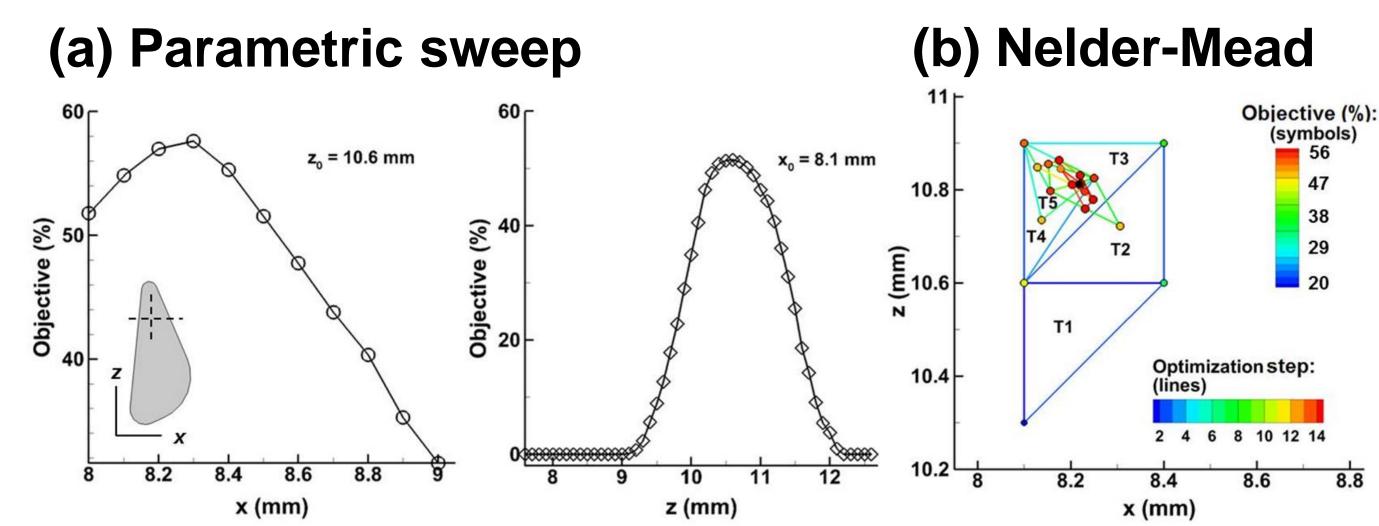
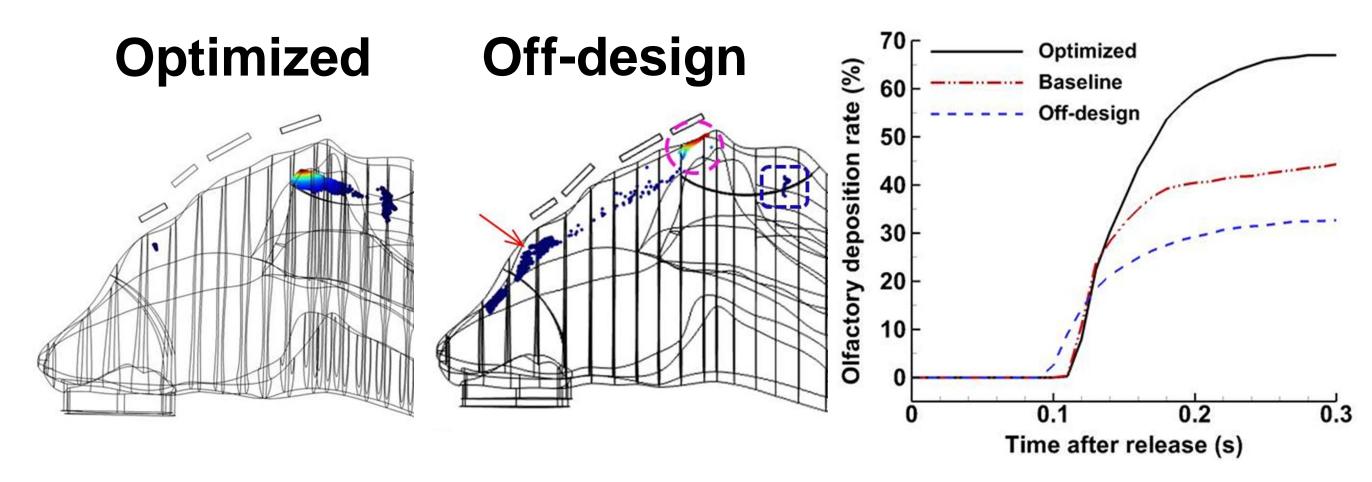


Figure 4. Optimization of point-release position



Conclusions:

- 1. With magnetic-guidance and point-release, improved olfactory dosing is feasible.
- 2. Topical delivery efficiency can be 70%.

References:

1. J. Xi, Z. Zhang, X. Si, Improving intranasal delivery of neurological nanomedicine to the olfactory region using magnetophoretic guidance of microsphere carriers, Int. J. Nanomedicine, 10, 1211-1222, (2015).