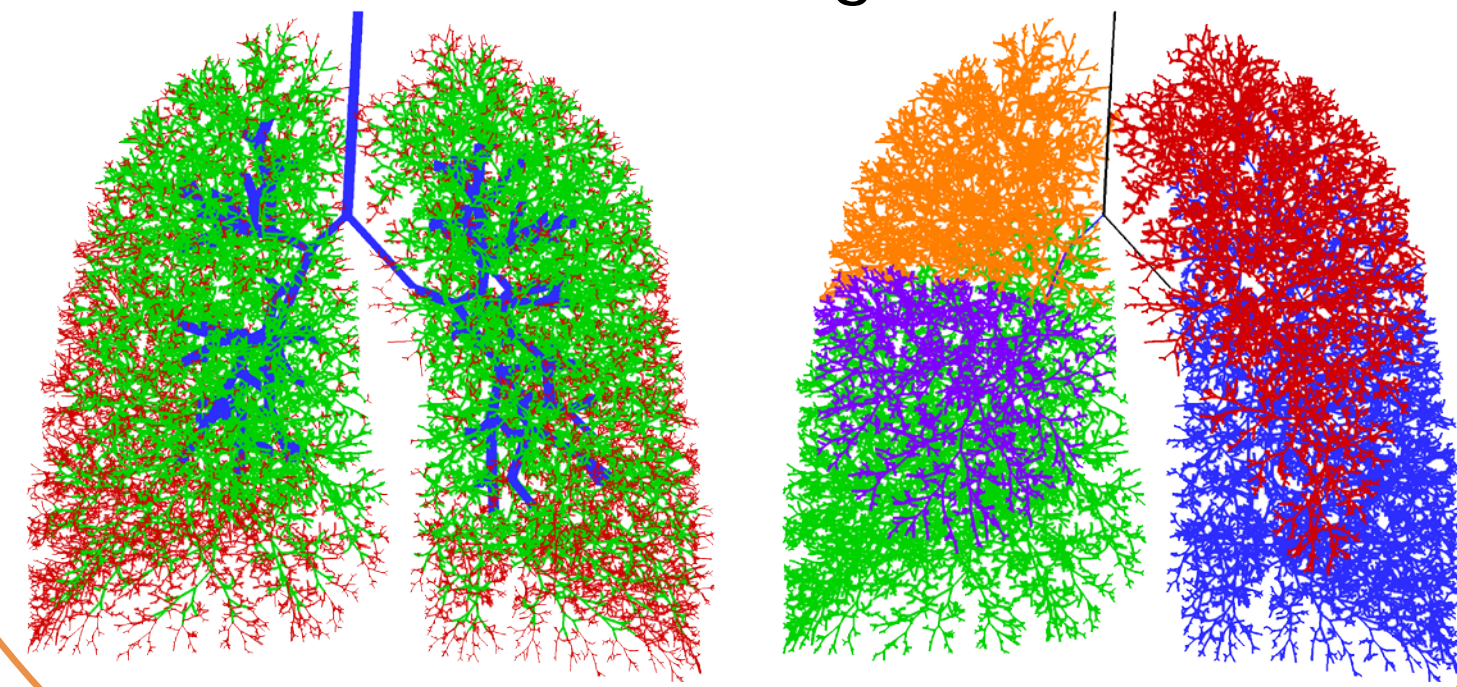


1. Effective Drug Delivery

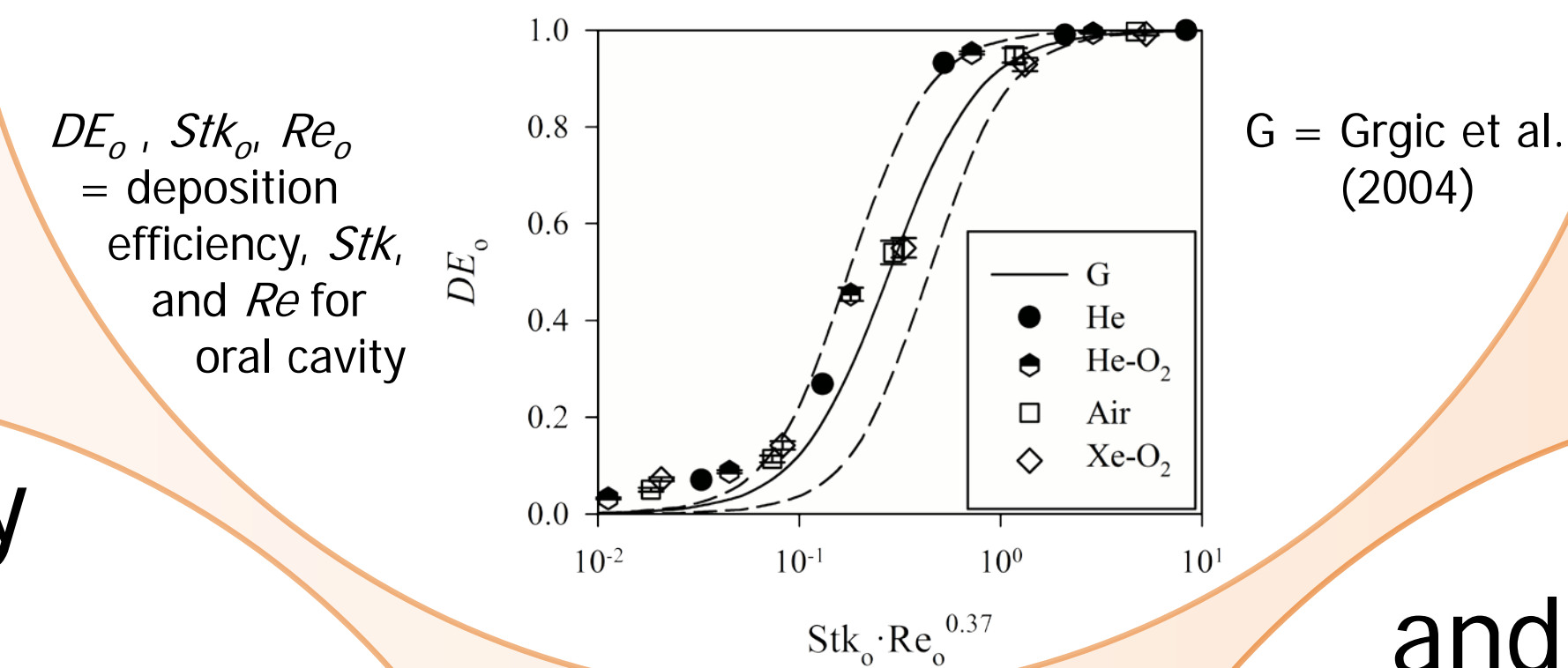
The mechanisms of regional aerosol distribution are investigated to maximize therapeutic drug effect, minimize unwanted side effects, and reduce treatment cost by effective delivery of inhaled drugs to different regions of interest in the lung.



Serial targeting Parallel targeting

3. Numerical Simulation

- Gas flow simulation (CFD)
 - Large eddy simulation (LES) (3-D, unsteady, eddy-resolving)
 - Gas property: He, He-O₂, air, Xe-O₂
 - Steady inspiratory flow: 342 mL/s
- Aerosol simulation (Lagrangian particle tracking)
 - Unsteady flow field for turbulent flow
 - Particle size = 2.5 μm



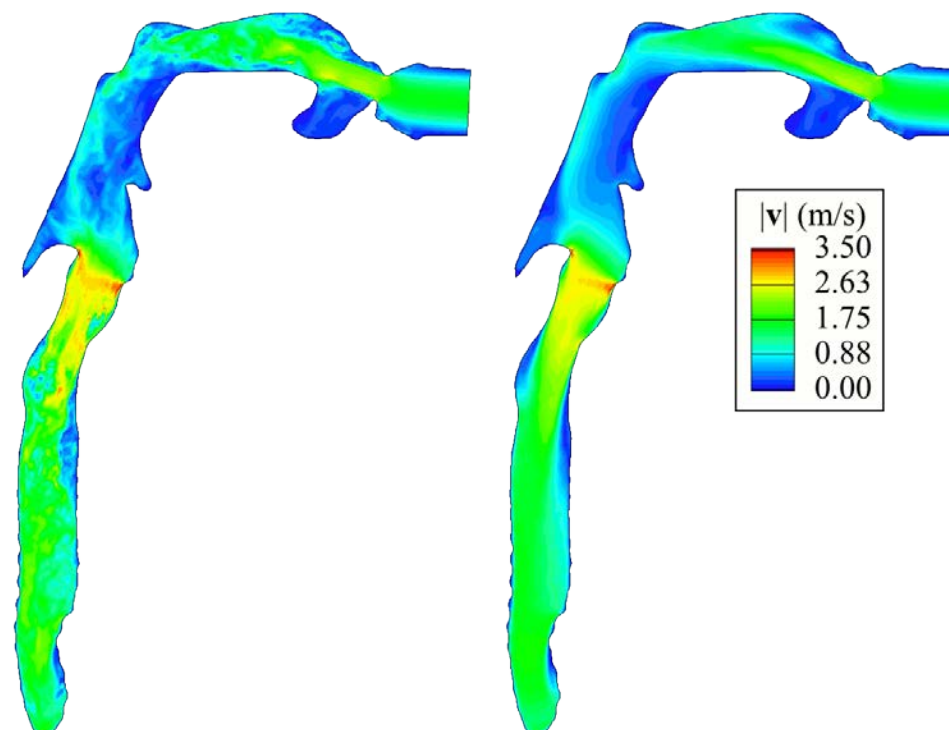
2. Unsteady Simulation

In turbulent flows ($Re_{trachea} > 400 \sim 600$)
(Time-averaged flow + α + particle)
 \neq

Time-averaged (flow + particle)

- Non-linear nature of fluid and particle motion
- Different temporal and spatial scales of turbulent flow structures

Instantaneous flow field
- anisotropic turbulence
- oscillation of jets



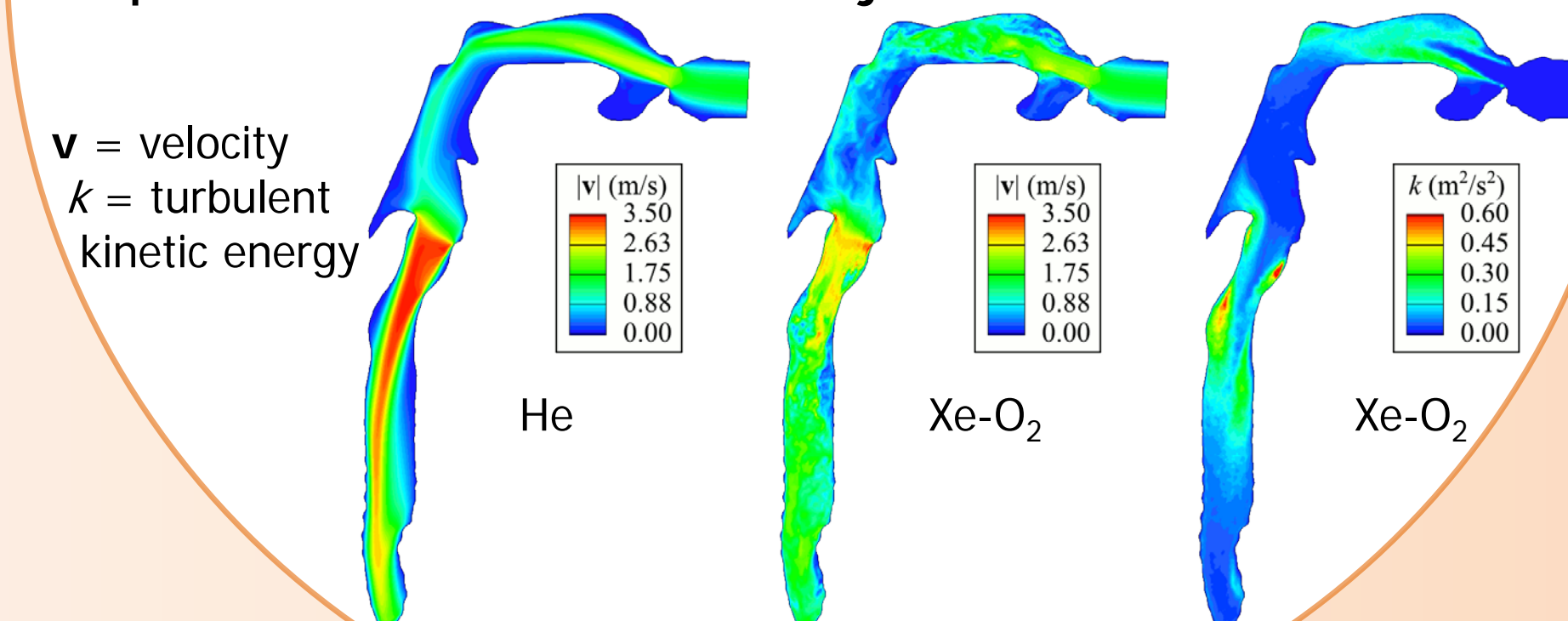
Time-averaged flow field + α
- isotropic turbulence
- steady jets

4. Laminar and Turbulent Flows

He ($Re_t \sim 190$): Laminar flow, flow field does not vary in time (stable).

He-O₂ ($Re_t \sim 460$): Transitional flow, low-frequency oscillation of jets in the oral cavity and trachea. Critical $Re_t \sim 430$.

Air, Xe-O₂ ($Re_t \sim 1300, 2800$): Turbulent flow, dispersed flow and shorter jets.



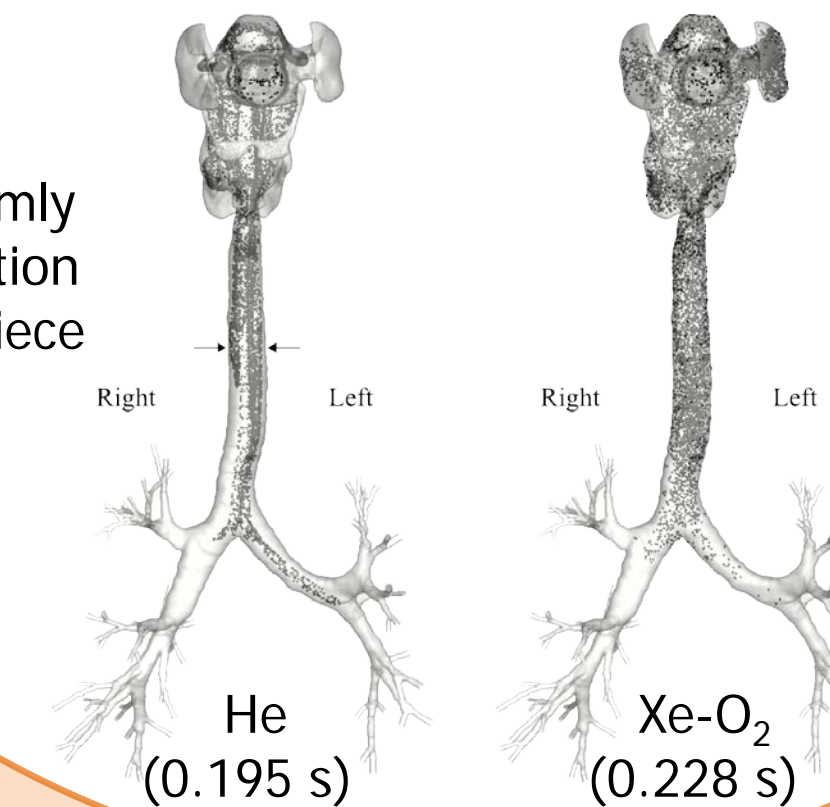
5. Aerosol Distribution

He: Small particles basically follow the steady streamlines of the flow, so the particles are not well mixed.

He-O₂: Particles are mixed by the jets oscillating with low-frequency in the oral cavity and trachea.

Air, Xe-O₂: Particles are well mixed by the turbulence in the oral cavity and trachea.

The aerosols are distributed uniformly in the cross-section at the mouthpiece



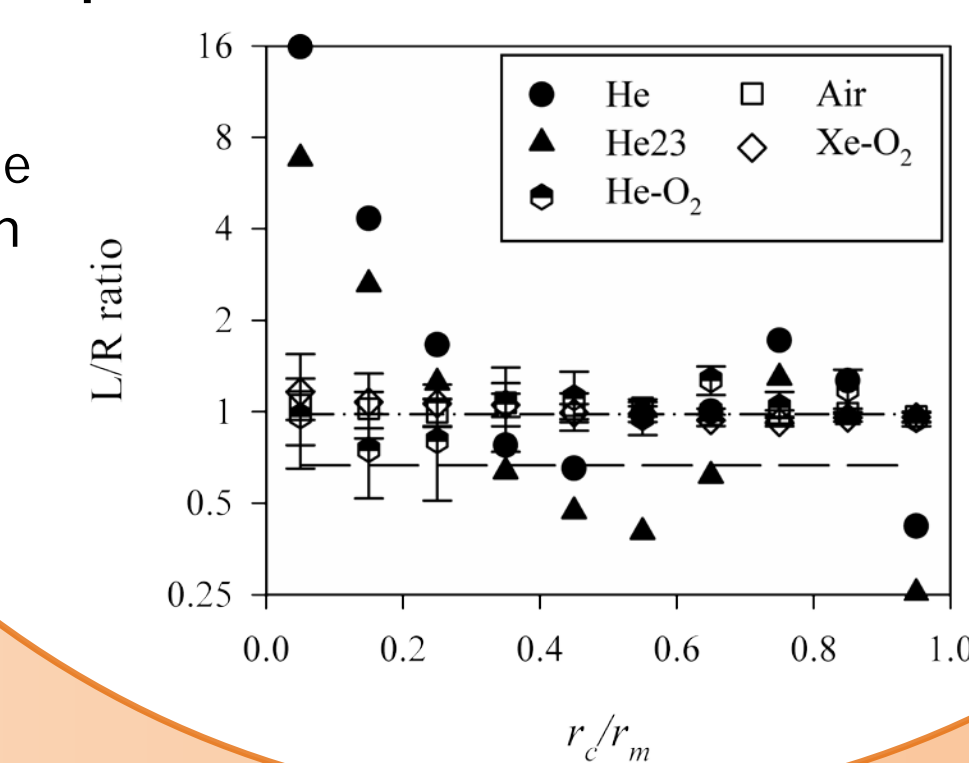
6. Parallel Targeting

He: L/R particle distribution of small particles depends on the particle location at the mouthpiece. Large deviation of L/R particle distribution ratio from the L/R ventilation ratio.

He-O₂: Large uncertainty due to the low-frequency oscillation of the jets.

Air, Xe-O₂: L/R particle distribution of small particles depends on the L/R ventilation ratio.

L/R ratio = L/R particle distribution ratio



r_m = radius of mouthpiece
 r_c = radial coordinate of particles at mouthpiece

7. Conclusions

The regional distribution of micro-particles is sensitive to the airway geometry and the flow structure in laminar flows, while regional ventilation determines the regional particle distribution in turbulent flows. Laminar flows could potentially be used for parallel targeting with low density gas and/or low flow rate.

Reference

Miyawaki, S., M. H. Tawhai, E. A. Hoffman, and C. L. Lin. Effect of Carrier Gas Properties on Aerosol Distribution in a CT-based Human Airway Numerical Model. *Ann. Biomed. Eng.* (2012), DOI: 10.1007/s10439-011-0503-2

Acknowledgments:

This work was supported by NIH grants R01-HL094315, R01-HL064368, R01-EB005823, and S10-RR022421. The authors thank the San Diego Supercomputer Center (SDSC), the Texas Advanced Computing Center (TACC), and XSEDE sponsored by NSF for the computer time.