Switching between magnetotactic and aerotactic sensory-based motion of MC-1 bacterial nanorobots

#### Sylvain Martel

NanoRobotics Laboratory - Polytechnique Montréal Montréal, CANADA Canada Research Chair (CRC) in Medical Nanorobotics

> sylvain.martel@polymtl.ca 2016

## Physiological Pathways to a Tumor

- Narrowest blood vessels (diam. 4 micrometers)
- Interstitial spaces
- The intercellular openings of typically less than 2 µm in diameter between endothelial cells\* responsible for much of the vessel leakiness in solid tumors.

\*McDonald D.M. & Baluk P. Significance of blood vessel leakiness in cancer. *Cancer Res.* **62**, 5381-5385 (2002).

# This suggests a diameter not exceeding 2 micrometers



#### MC-1 Magnetotactic Bacterium

 Since an artificial implementation is far beyond technological feasibility, a potential strategy was to identify a microorganism that has all these specifications and to harness it to act as a "nanorobot" for drug delivery applications



Pettaquamscutt Estuary in Rhode Island, USA





#### Harnessing What Nature Already Provides







S. MARTEL - NanoRobotics Laboratory, Polytechnique Montréal

# MC-1 Strain Magnetococcus Marinus Natural Migration Behavior

- Contains a chain of magnetic iron-oxide (Fe<sub>3</sub>O<sub>4</sub>) nano-crystals enclosed in membranes known as magnetosomes acting like a nano-compass needle.
- Downward migration along geomagnetic field lines in conjunction with aerotaxis to efficiently migrate to and maintain position at their preferred low oxygen concentrations.
- Such magnetically-assisted aerotaxis known as magnetoaerotaxis results in the formation of microaerophilic bands of MC-1 cells at O<sub>2</sub> concentrations equivalent to the ones observed in the hypoxic regions of solid tumors.







#### **Experimental Magnetotaxis Platform**



Magnetic Pole



	A	0	T	2T	3T	4T	5T	6T
	X-Left							
agnet	X-Right							
ctrom	Y-Bottom							
m elec	Ү-Тор	_						
Syste	Z-Back	_						
	Z-Front	_						
	Time							
	ВΓ	0	1T	2T	3T	4T	5T	6T
	X-Left							

Time

System electromagnet	X-Left	
	X-Right	
	Y-Bottom	
	Ү-Тор	
	Z-Back	
	Z-Front	

	$\sim$	Time								
	Uμ	0	1T	2T	3T	4T	5T	6T		
	X-Left									
agnet	X-Right									
System electrom	Y-Bottom									
	Ү-Тор									
	Z-Back									
	Z-Front	-								

#### **Basic Principle**











#### **Oxygen Sensors – Microaerophilic Behavior** (Objective: Autonomous Targeting of Tumor Hypoxic Zones)



## Switched Taxis-based Directional Control



#### Reference: S. Martel, "Bacterial Microsystems and Microrobots," *Biomed. Microdevices*, 2013



# Acknowledgements

- Several funding organisms are acknowledged for their support and vision by providing the resources required to pursue the development of these technologies. The main supporting organizations include Polytechnique Montreal, Univalor, Consortium Québécois sur la Découverte du Médicaments (CQDM), CMC Microsystems, ReSMIQ, GRSTB, Canada Research Chair (CRC) Program, Natural Sciences and Engineering Research Council of Canada (NSERC), Government of Québec, Canadian Funds for Innovation (CFI), National Institute of Health (NIH), and several other organisms that contributed financially to our partners and students.
- The active participation of many individuals besides the staff, the graduate students and the technicians is worth
  mentioning and the list of participants in such highly interdisciplinary projects to make it accessible clinically in the future
  is growing.
- In alphabetical order (current immediate collaborators in medical applications only):
- G. Batist Oncologist, McGill University
- N. Beauchemin Biochemist, McGill University
- G. Beaudoin Medical Physics and MRI sequencing, University of Montréal
- F. Cheriet Medical imaging, Polytechnique Montréal
- L. Gaboury Pathologist, University of Montréal
- H. Girouard Pharmaceutical, University of Montréal
- S. Kadoury Medical image registration, Polytechnique Montréal
- M. Lafleur Chemist, University of Montréal
- M. Mohammadi Biologist, bacterial culture, Polytechnique Montréal
- D. Radzioch Immunologist, McGill University
- G. Soulez Interventional Radiologist, University of Montréal
- M. Tabrizian Biomaterials and Bio-interfaces, McGill University
- T. Vuong Radio-oncologist, McGill University
- Etc...

#### Sylvain Martel, P. Eng., Ph.D., FIEEE, FCAE

NanoRobotics Laboratory Polytechnique Montréal, CANADA

Contact: sylvain.martel@polymtl.ca

