

Allometry

Biomechanics – 16.12.19

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BIOREACTORS

DESIGN AND
REALIZATION

COMPUTATIONAL
MODELS

SENSING AND
ACTUATION



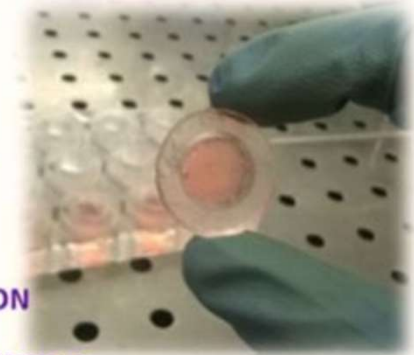
BIOMECHANICS & BIOMATERIALS

HYDROGELS &
BIOPRINTING

TISSUE-DERIVED
SCAFFOLDS

MECHANICAL
CHARACTERIZATION

CELL MECHANOSENSING

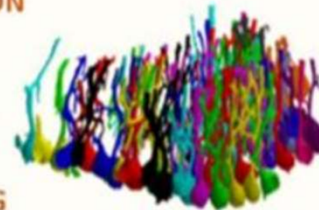


CELL IMAGING

TISSUE DELIPIDATION

CELL
MORPHOMETRICS

IMAGE PROCESSING

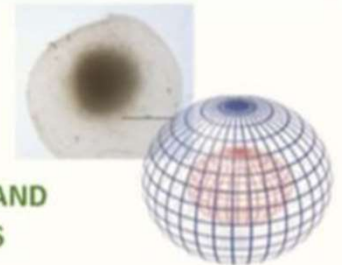


LIVER AND BRAIN ORGANOID

ALLOMETRIC SCALING

NANOTOXICOLOGY

NUTRIENT TRANSPORT AND
CONSUMPTION MODELS



Allometry: scaling property emerging in all living organisms about characteristic physiological parameters (e.g. metabolic rates), which are related to body size (*i.e.* mass) through power laws.

$$Y = aM^b$$

$$\log Y = \log a + b \log M$$

- a normalization constant (depending on Y and on taxonomic class)
- b scaling exponent (depending on Y)

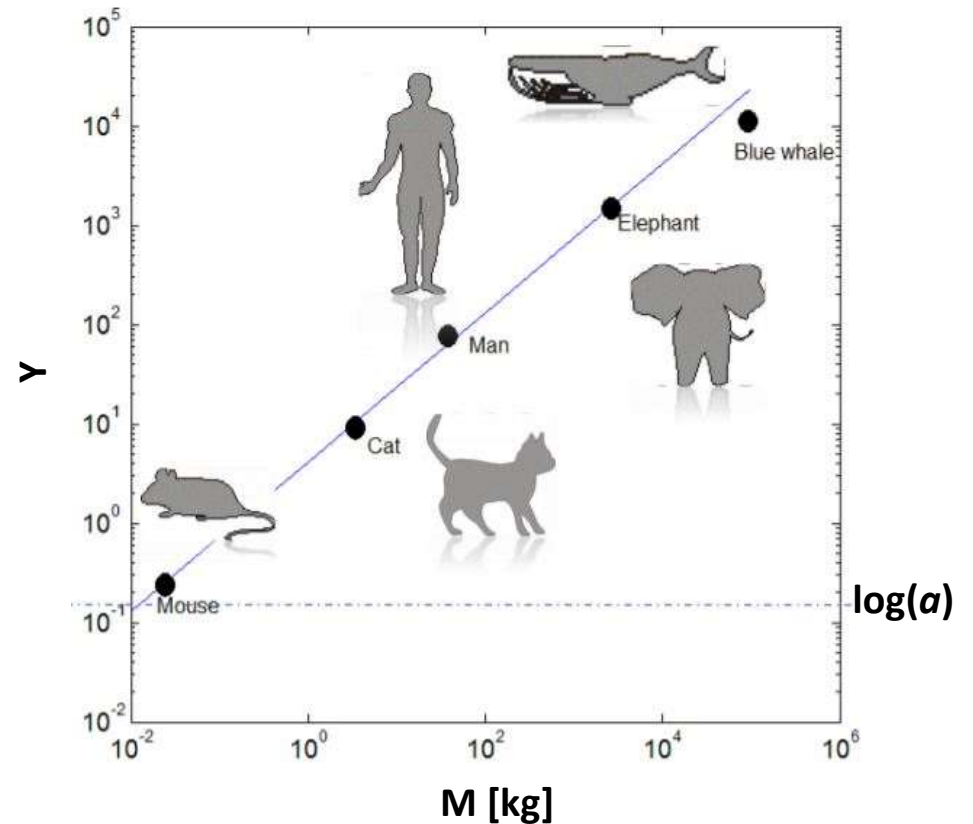


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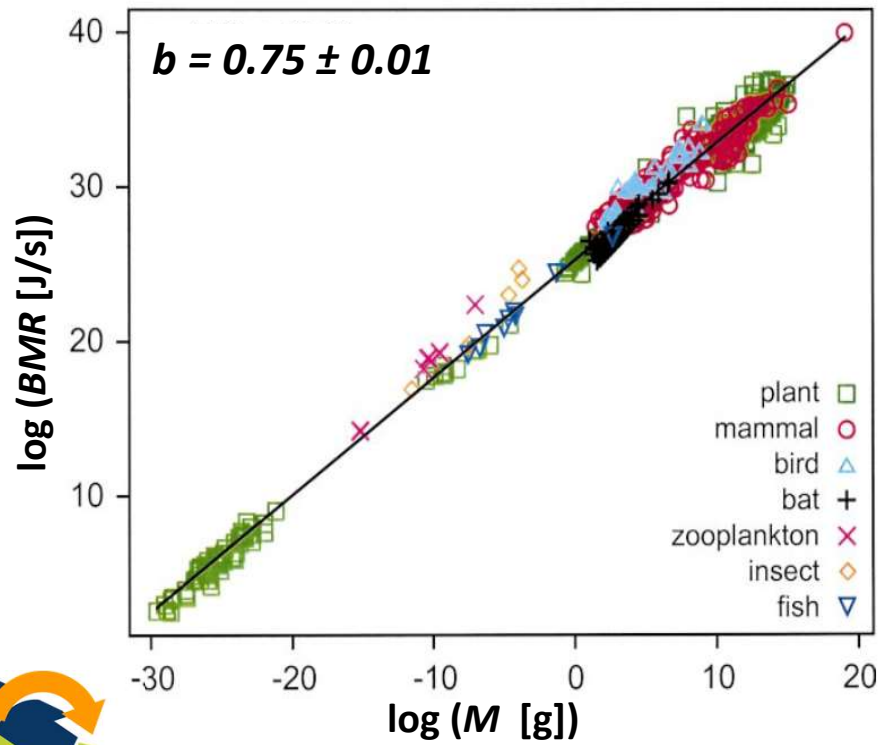


PARAMETER	EXPONENT VALUE	MEANING
Cells size [m] Blood velocity [m/s] Pressure gradients [Pa]	$b = 0$	Parameter and body mass are independent
Volumes (bone, blood...) [m ³]	$b = 1$	Parameter and body mass are directly proportional (isometric scaling)
Metabolic rates [J/s] Flow rates (haematic, respiratory...) [m ³ /s]	$b = 3/4$	Parameter increases slower than body mass
Radii of aorta and trachea [m]	$b = 3/8$	Parameter increases slower than body mass
Frequencies (cardiac, respiratory...) [Hz]	$b = - 1/4$	Parameter decreases when body mass increases
Bone mass [kg]	$b = 4/3$	Parameter increases faster than body mass



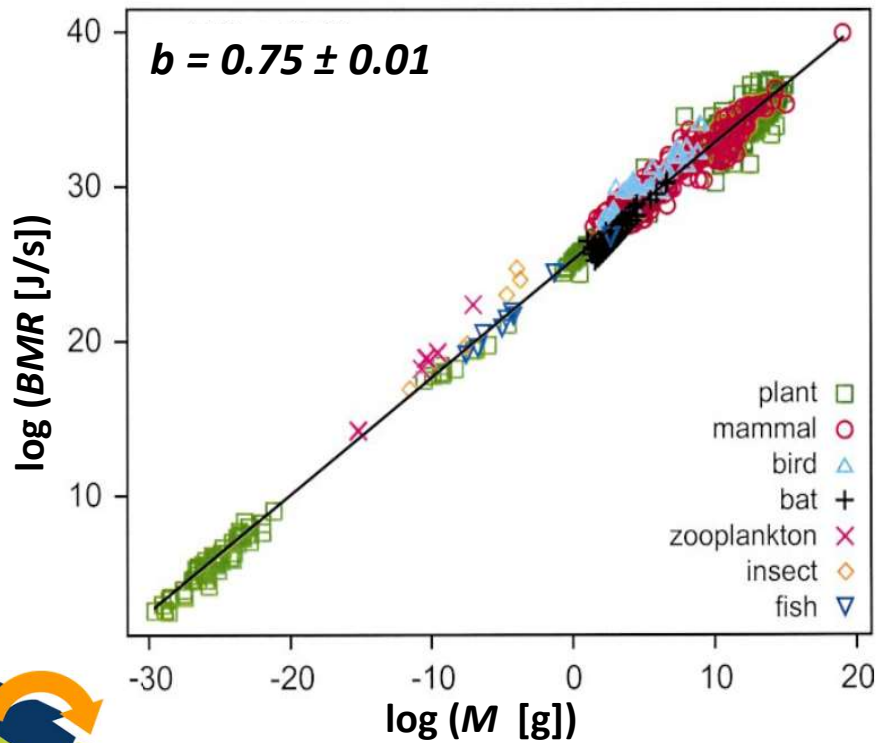
Kleiber law (KL)

$$BMR = aM^{3/4}$$

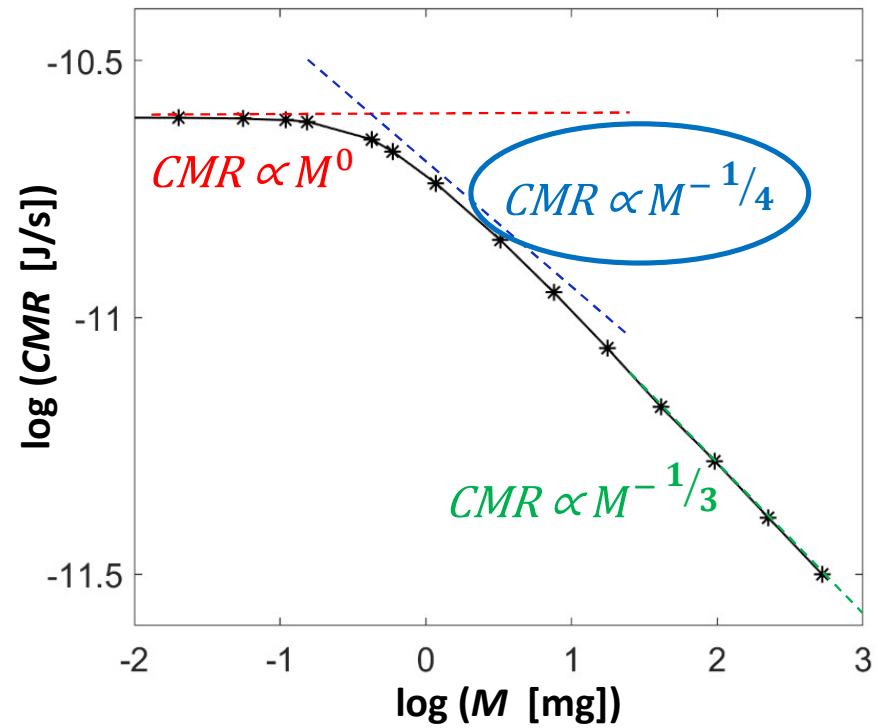


Kleiber law (KL)

$$BMR = aM^{3/4}$$



$$CMR = a'M^{-1/4}$$



Origin of quarter-power scaling

➤ Geometric scaling

$$\left\{ \begin{array}{l} A \propto l^2 \\ V \propto l^3 \\ V \propto M \end{array} \right.$$



$$\begin{array}{l} l \propto M^{1/3} \\ A \propto M^{2/3} \end{array}$$



Origin of quarter-power scaling

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Why allometric exponents are not multiples of 1/3?



Origin of quarter-power scaling

➤ Geometric scaling

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➤ Biological scaling

- Stoichiometric constraints in biochemical processes
- Integrated optimization of interdependent sub-systems
- Self-similar structure of nutrients supply networks



Origin of quarter-power scaling

➤ Geometric scaling

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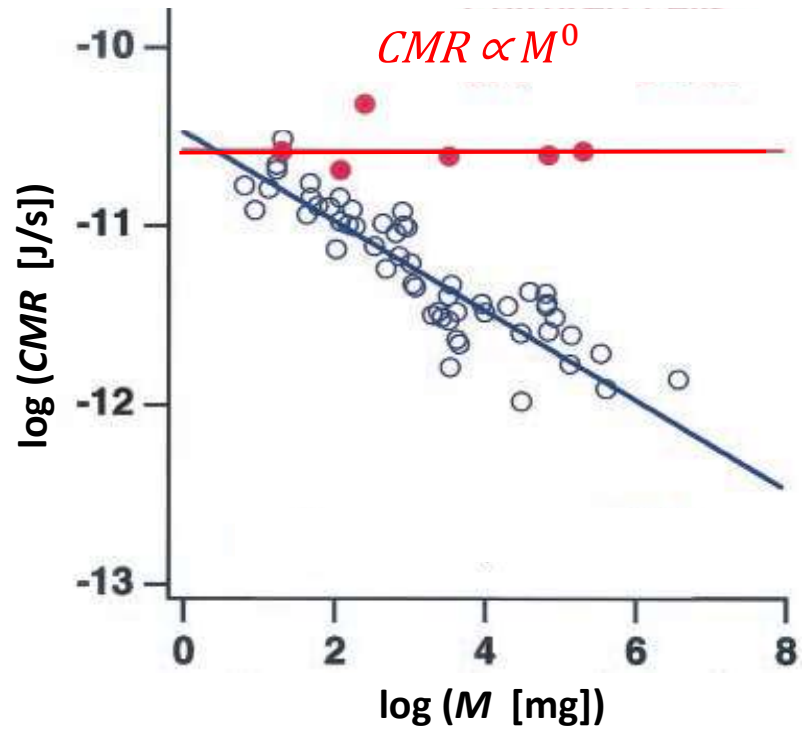
➤ Biological scaling

- Stoichiometric constraints in biochemical processes
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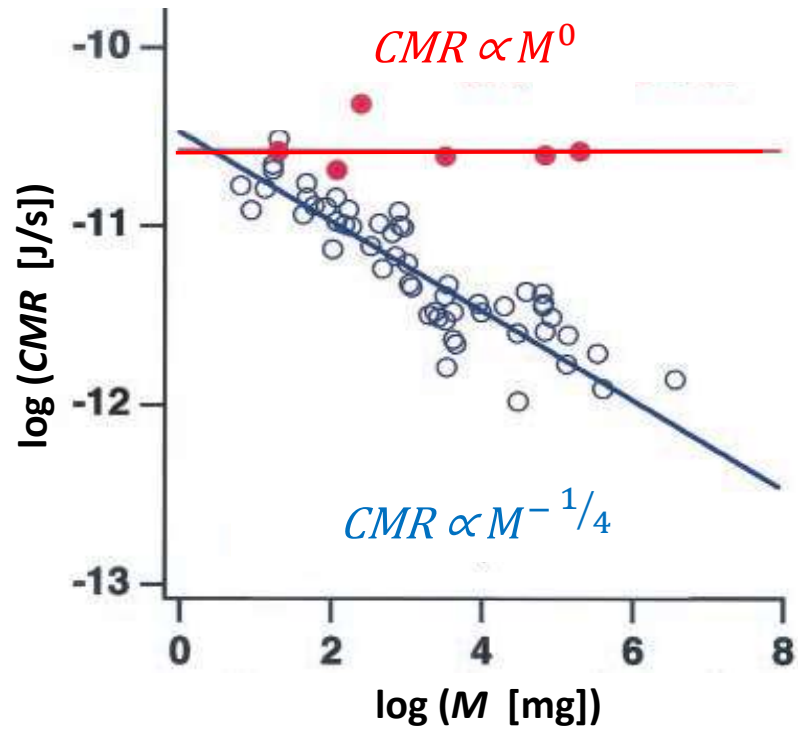
Since **metabolic rates** (per cell, *i.e.* CMR) **underlie all physiological processes** and scale with $M^{-1/4}$, allometry is described by quarter-power scaling



Why allometry in bioengineering?



Why allometry in bioengineering?

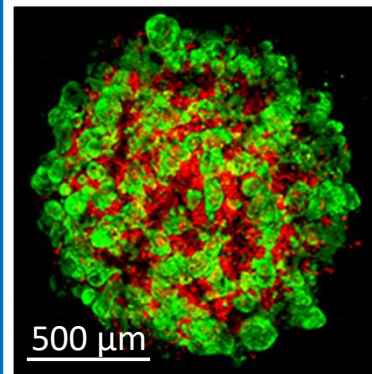


In vitro monolayers

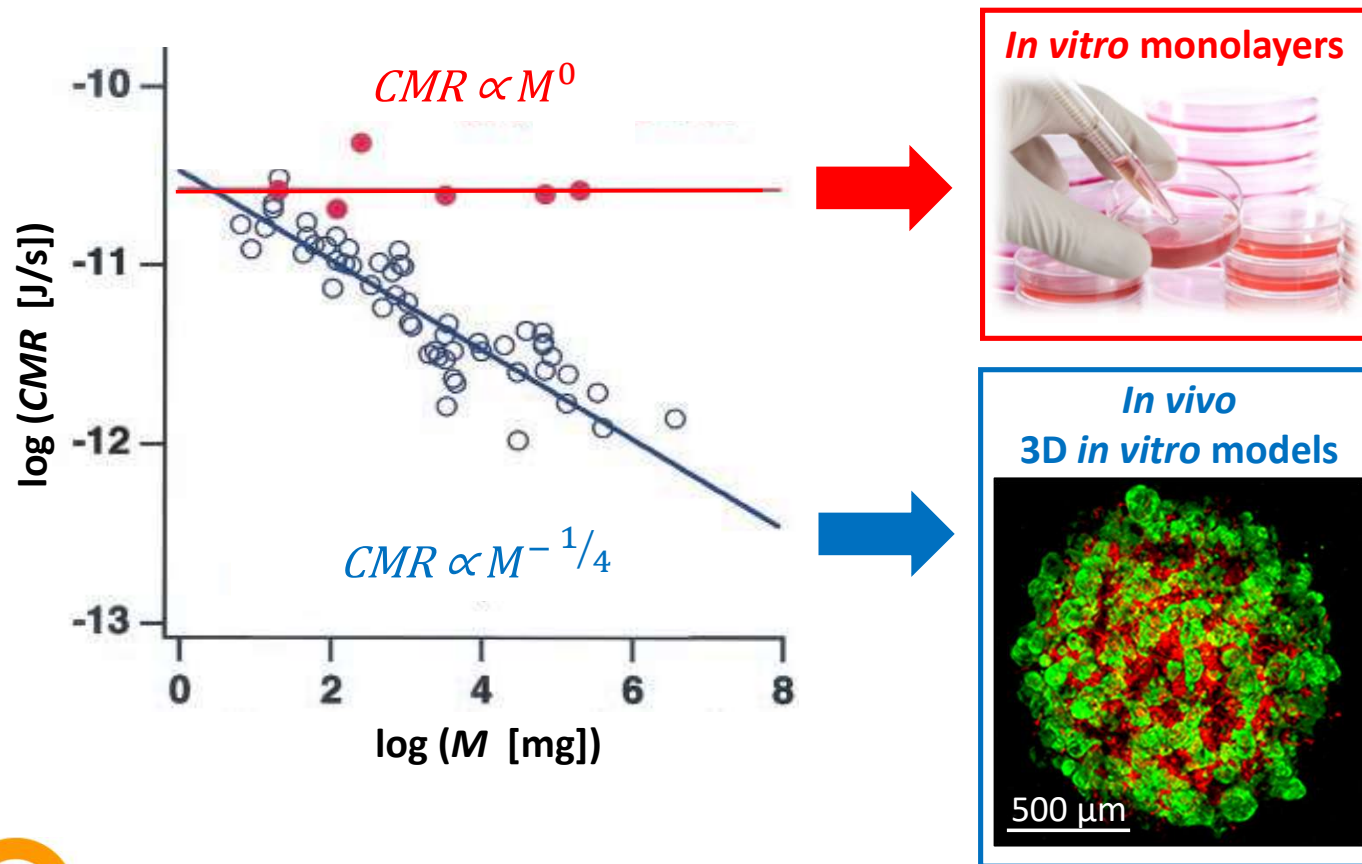


In vivo

3D *in vitro* models



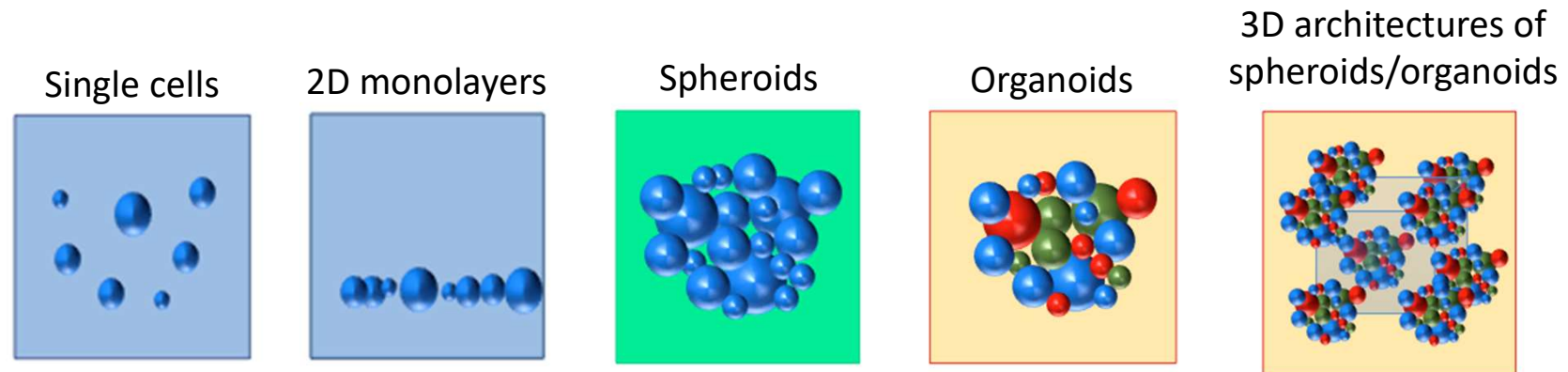
Why allometry in bioengineering?



Since **KL** is a **universal law for biological systems**, we need to account for it to design **predictive and physiologically relevant *in vitro* models!**



Why allometry in bioengineering?



INCREASING COMPLEXITY

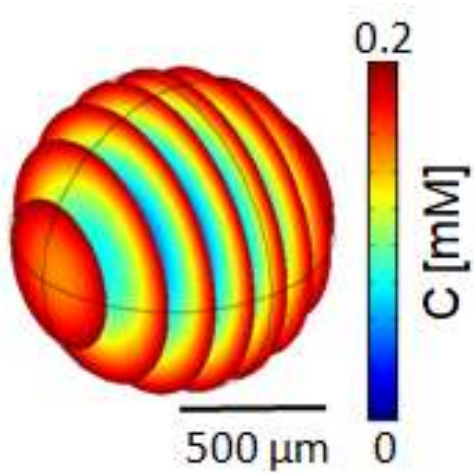
Which size level does allometric scaling start from?
Which is the size range allowing allometric scaling to emerge?



Case of study: liver and brain *in vitro* models

In silico modelling: diffusion and consumption

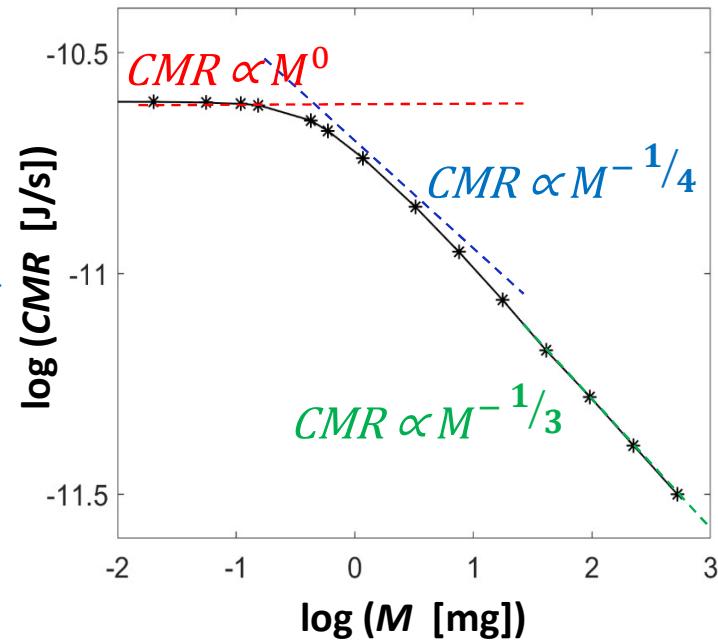
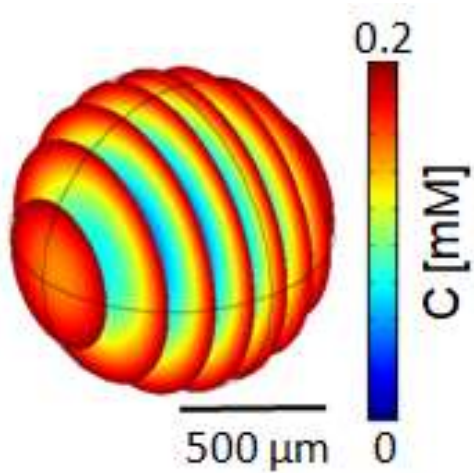
$$\frac{\partial c_{O_2}}{\partial t} = \nabla^2(D_{O_2}c_{O_2}) + R = \nabla^2(D_{O_2}c_{O_2}) - \frac{V_{max}c_{O_2}}{k_M + c_{O_2}}$$



Case of study: liver and brain *in vitro* models

In silico modelling: diffusion and consumption

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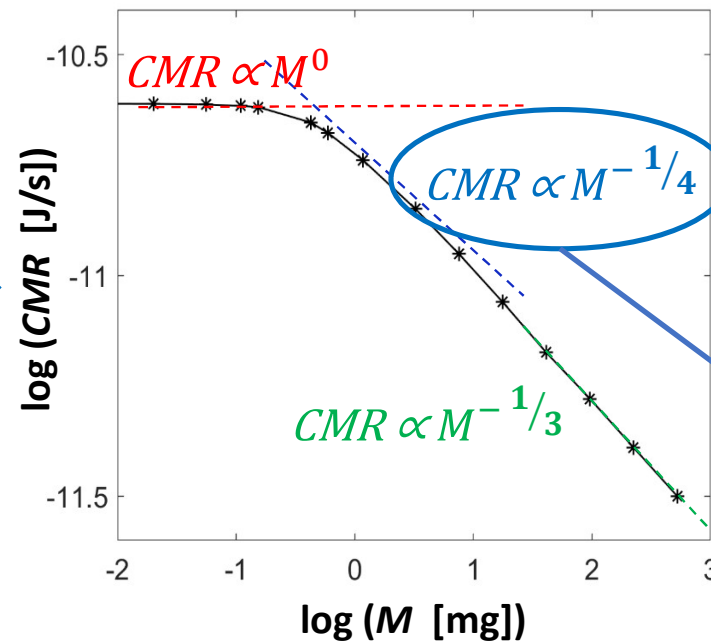
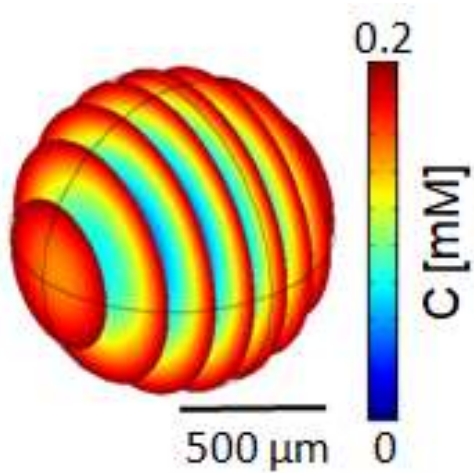
$$B_{O_2} = -\Delta H \iint \underline{J}_{O_2} \cdot \partial \underline{A} = -\Delta H \iint \underline{\nabla}(D_{O_2}c_{O_2}) \cdot \partial \underline{A}$$



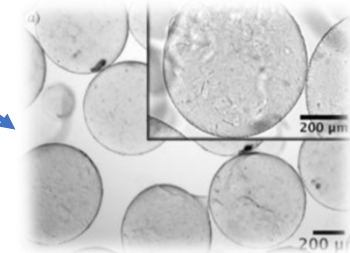
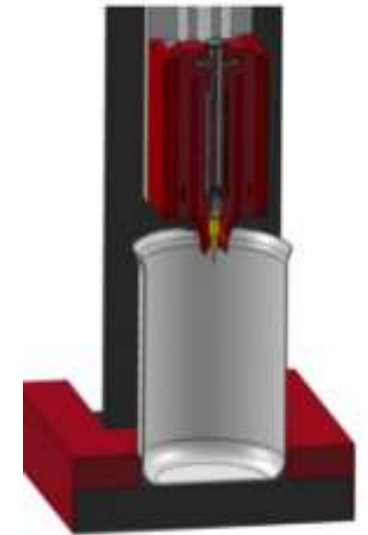
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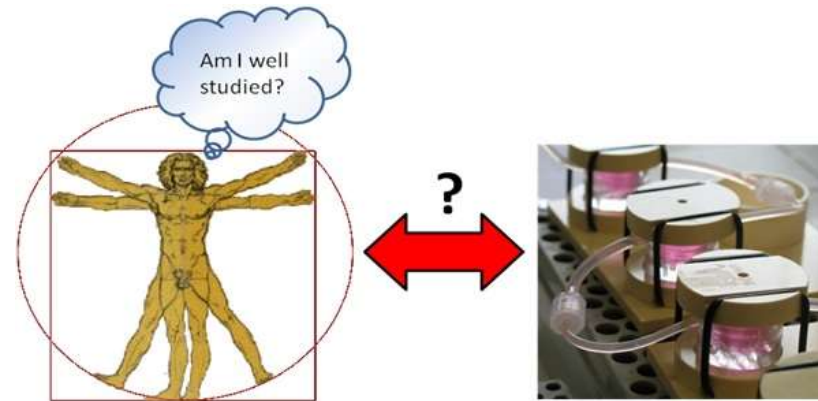
Fabrication, *in vitro* O₂ measurements and validation



$$B_{O_2} = -\Delta H \iint \underline{J}_{O_2} \cdot \partial \underline{A} = -\Delta H \iint \underline{\nabla}(D_{O_2}c_{O_2}) \cdot \partial \underline{A}$$



Thesis are available on these topics!



If you are interested in:

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