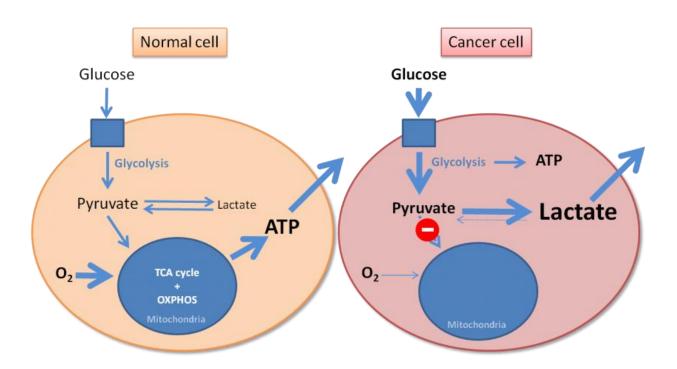
Biological background: Cancer metabolism – Warburg effect

- Healthy cells primarily produce energy through mitochondrial ox. phos.
- Most cancer cells have high rate of glycolysis followed by lactic acid fermentation even in presence of abundant oxygen
- Aerobic glycolysis (the Warburg effect)
- Can we simulate the same behaviour using Genome-Scale Metabolic Models?



Outline of practical

- 1. We will use the Recon 2.2 genome scale metabolic model (Swainston et al. 2016) and the COBRA package in Matlab (https://opencobra.github.io/cobratoolbox/stable/)
- 2. For all simulations, we will treat the model as 'cells growing in a medium' by virtually feeding it with RPMI medium
- 3. First, we will assess the difference between how much energy can be created in this model under aerobic and anaerobic conditions
- 4. Next, we will assess where the main differences in reactions fluxes occur
- 5. Finally, we will create a cancer specific model and repeat the steps. What are the differences between a metabolic model of a healthy cell culture and a cancer cell culture?

We don't have the time for a complete programming course in Matlab, the focus of the practical is on interpretation. However, there is 'cheat sheet' here with the most common Matlab commands.

Short instructions

- 1. You can find all the required materials on Canvas
- 2. We will use Matlab for the practical
- 3. The installation process is as follows:
 - 1. Save **GEM_Practical_2021.zip** to: 'SYSTEM(C:) \rightarrow 'Users' \rightarrow 'Public' \rightarrow 'Public Downloads'
 - 2. Unzip **GEM_Practical_2021.zip** (right-click and use e.g. 7-Zip)
 - 3. Open Matlab
 - 4. Open the *FBA_normal.mlx* file in Matlab
 - Go through the script line by line (use the 'Step' button on top)
- 4. After completing the code and questions from *FBA_normal.mlx*, open the *FBA_cancer.mlx* file and go through it step by step as well
- 5. All instructions, including screenshots and weblinks with additional info, are included in the *mlx* files.
- 6. You may write your answers to the questions (also contained in the *mlx* files) in the separate Word file provided on Canvas.