7th October, 2025 Yerevan, Armenia



Role of *ion gradients* and redox potential during anaerobic and aerobic conditions

in E. coli and S. cerevisiae

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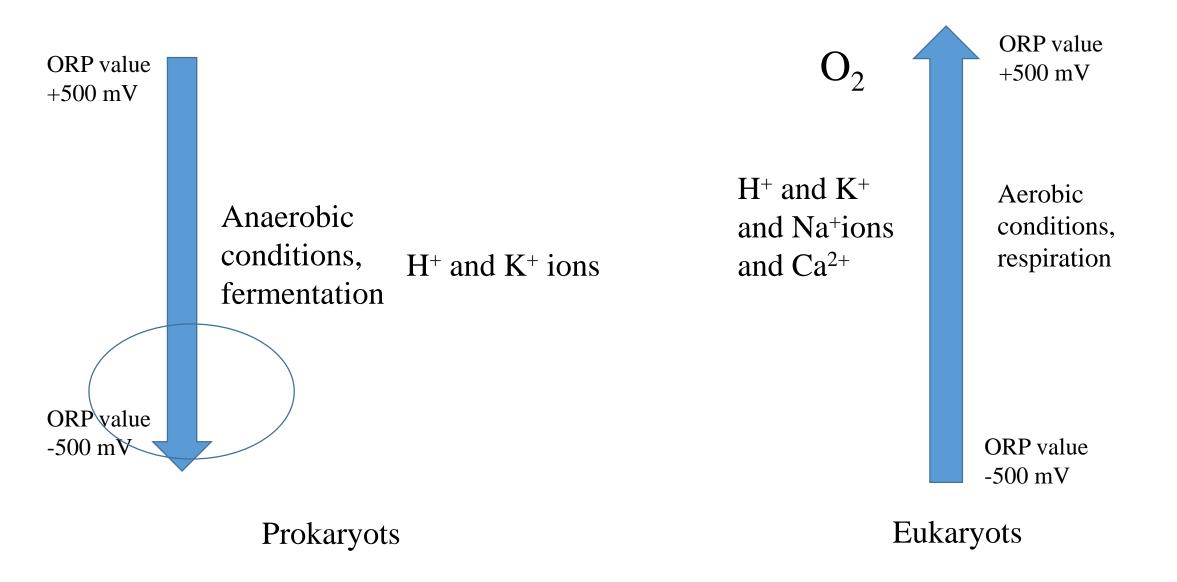
Movement or motion

Our nature consists in motion; complete rest is death.

Blaise Pascal

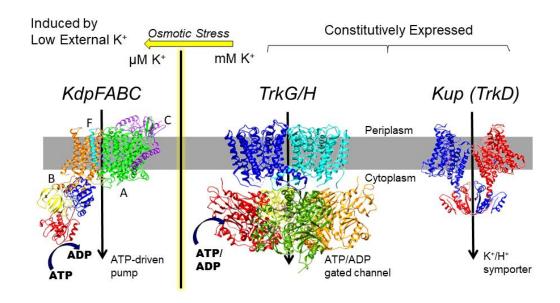


Ion fluxes, redox potential, oxygen

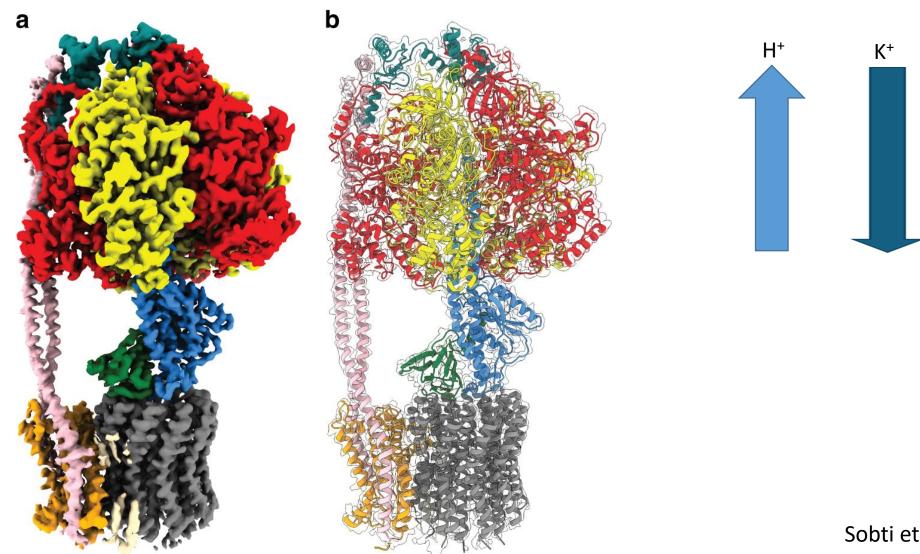


Potassium ions concentration in the cells

- ➤In sea water (ocean) 10 mM
- ➤In bacteria/yeasts 100 600 mM
- ➤In mammalian cells 100-150 mM
- ►In blood plasma ~5 mM

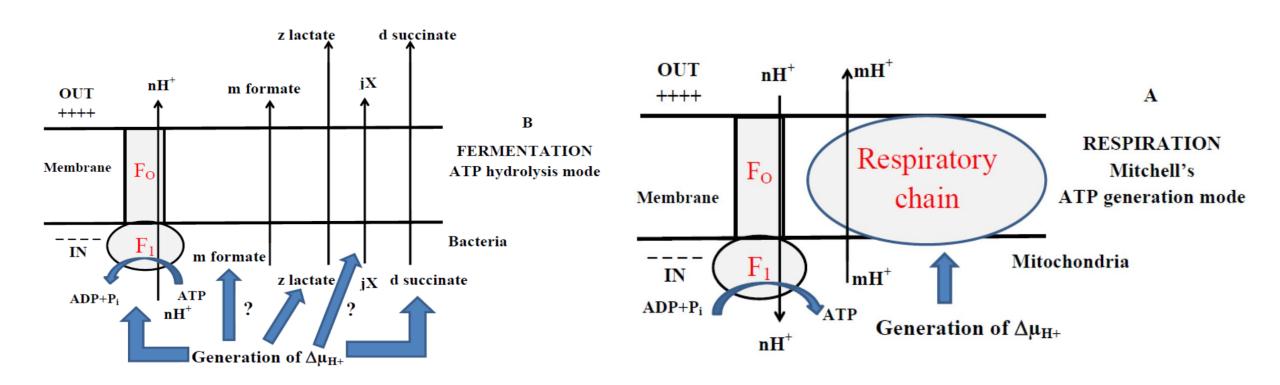


F₀F₁ ATP synthase

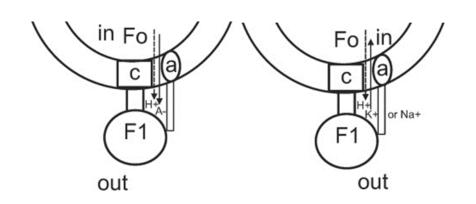


Sobti et al., 2020, Nature communications

Fermentation and respiration



Nath's two ion theory

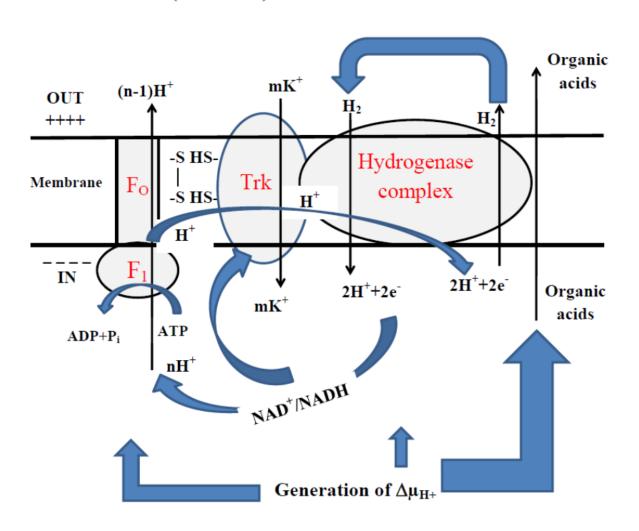




Nath's torsional mechanism was quantitative and contained mathematical equations that described the overall driving force of ATP synthesis arising from discrete proton and *anion/countercation* translocations in the membrane-bound F_O portion of F_1F_O -ATP synthase. These translocations occurred by a dynamically electrogenic but overall electroneutral mode of ion transport in the F_O portion of the ATP synthase that did not violate overall electroneutrality.

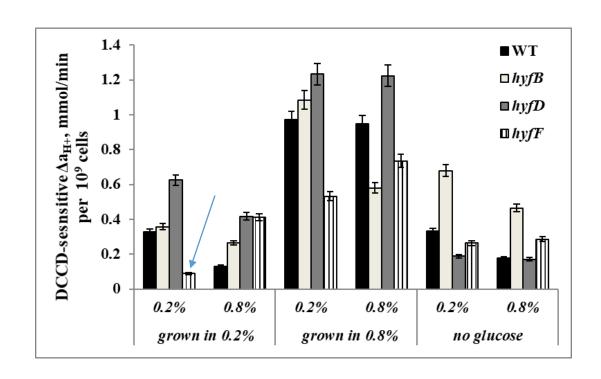
Ion gradients, F₀F₁, hydrogenases

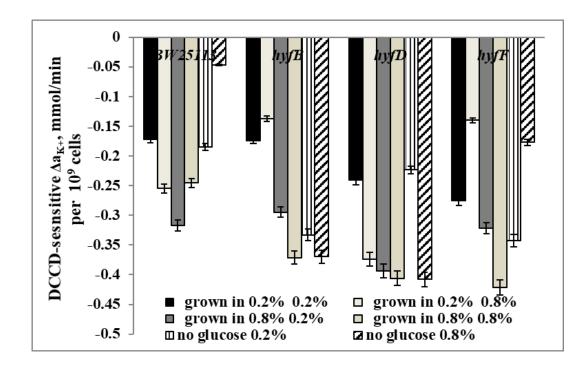
SUGAR (GLUCOSE) FERMENTATION



Trends in Biochemical Sciences, 2019, 44, 391-400

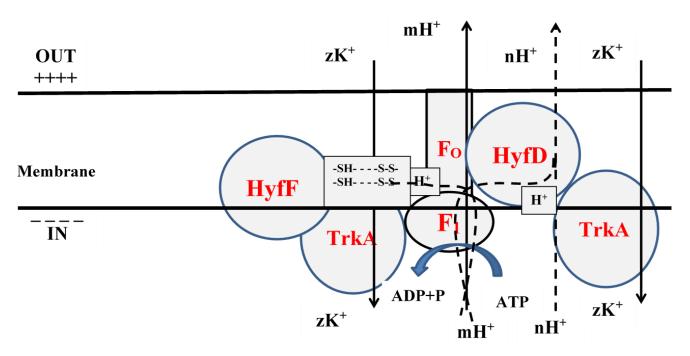
DCCD sensitive proton and potassium fluxes in Hyd-4 mutants





➤ In *hyfF* mutant grown on 2 g L⁻¹ DCCD sensitive proton flux was absent and this depends on glucose concentration

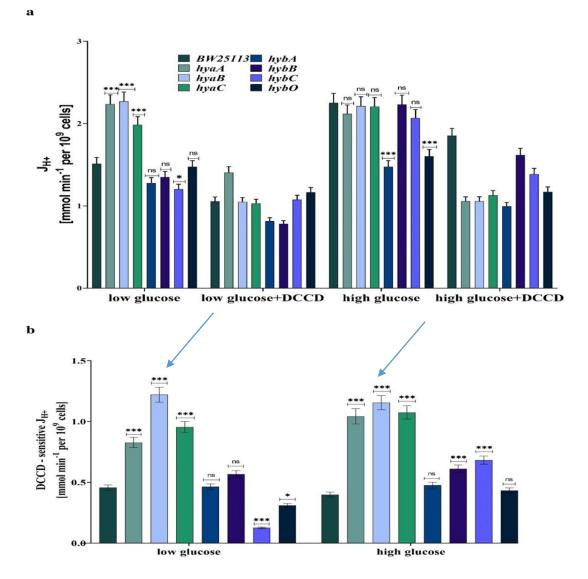
Schematic working model of Hyd-4 subunits



Strain	No glucose		Grown on 2 g L-1 glucose		Grown on 8 g L-1 glucose	
	2 g L ⁻¹	8 g L ⁻¹	2 g L ⁻¹	8 g L ⁻¹	2 g L ⁻¹	$8~\mathrm{g~L^{1}}$
WT	1.8	3.73	1.9	0.51	3.06	3.85
hyfB	2	1.25	2.1	1.93	3.75	1.48
hyfD	0.84	0.42	2.3	1.11	4.2	3.01
hyfF	0.77	1.6	0.32	2.94	1.66	1.74

• Vanyan and Trchounian, 2022 Journal of Bioenergetics and Biomembranes, 54, 69-79

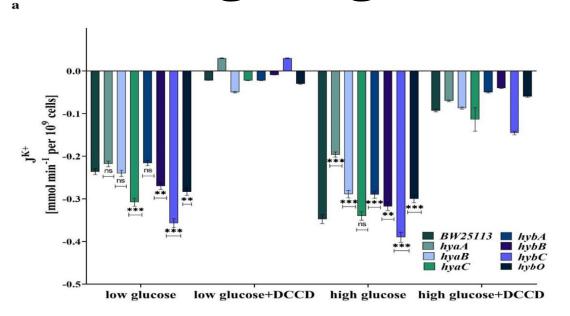
Total and DCCD sensitive H^+ ion fluxes with $2\ g\ L^{-1}$ glucose



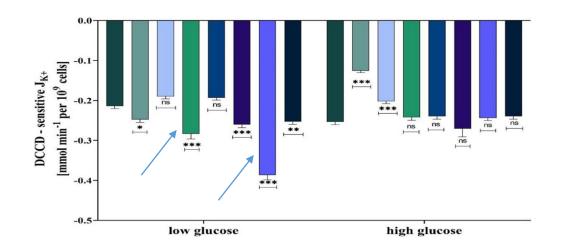
➤ In *hyaB* (Hyd-1) mutants DCCD sensitive H⁺ flux is increased ~ twice

• Vanyan and Trchounian, 2024, Biochimie, 227,205-216

Total and DCCD sensitive K^+ ion fluxes with $2\ g\ L^{-1}$ glucose

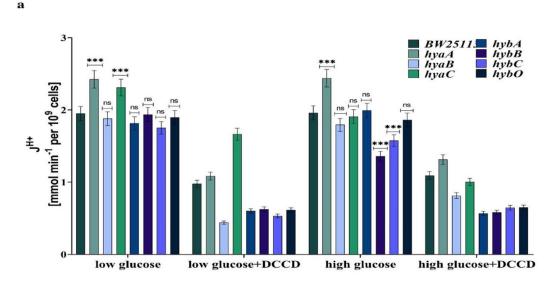


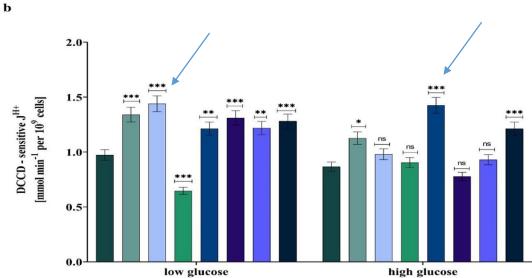
➤ In *hybC* mutants DCCD sensitive K⁺ flux is increased ~ twice



b

Total and DCCD sensitive H^+ ion fluxes with 8 g L^{-1} glucose

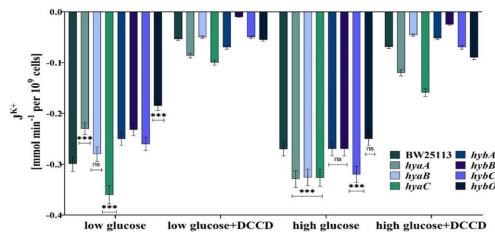


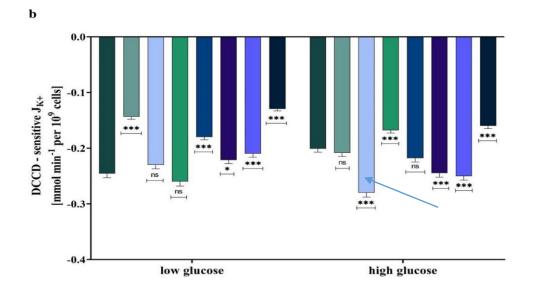


- ➤ In *hyaB* mutants DCCD sensitive H⁺ flux is increased ~ 50% (low glucose assays)
- ➤ In *hybC* mutants DCCD sensitive H⁺ flux is increased ~ 50% (high glucose assays)

Total and DCCD sensitive K^+ ion fluxes with 8 g L^{-1} glucose

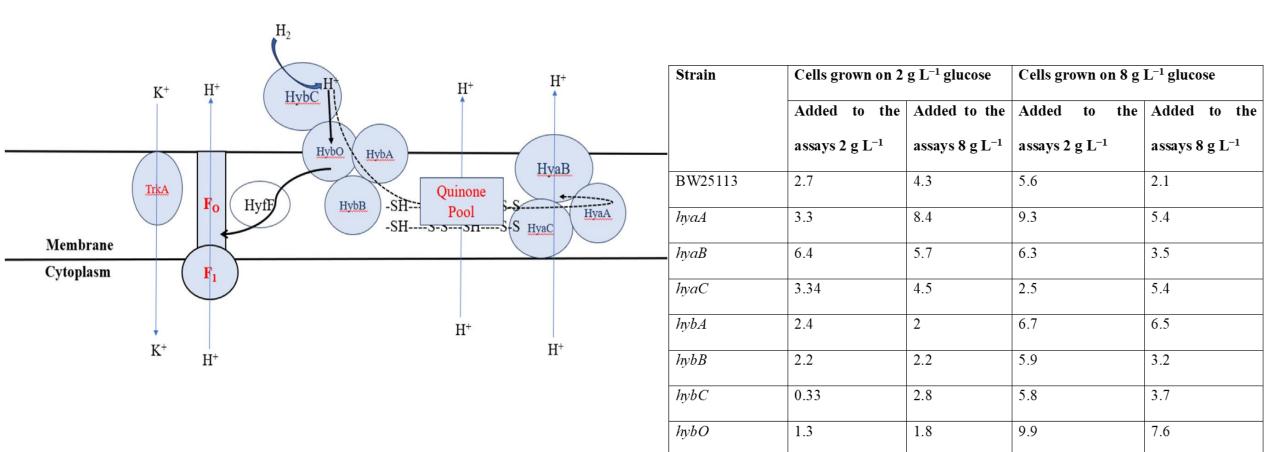




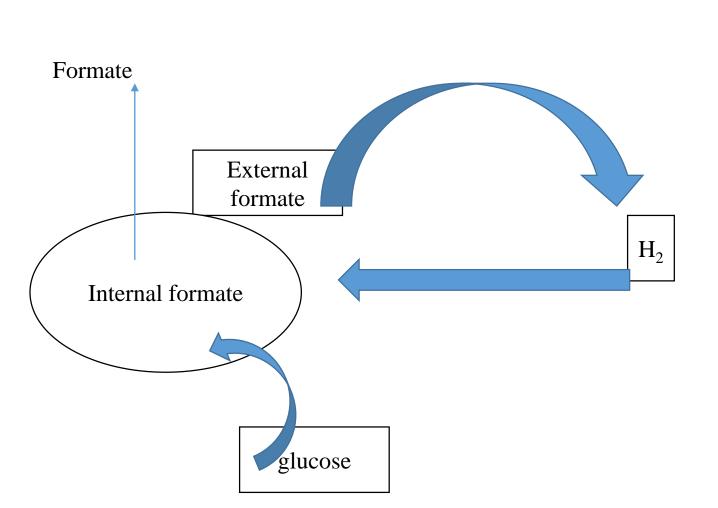


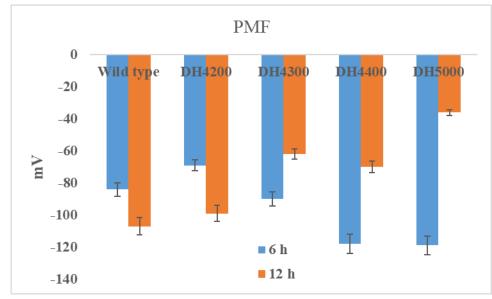
➤ In *hyaB* mutant DCCD sensitive K⁺ flux is increased ~ 50%

Schematic working model of Hyd-1 and Hyd-2



Internal/External formate





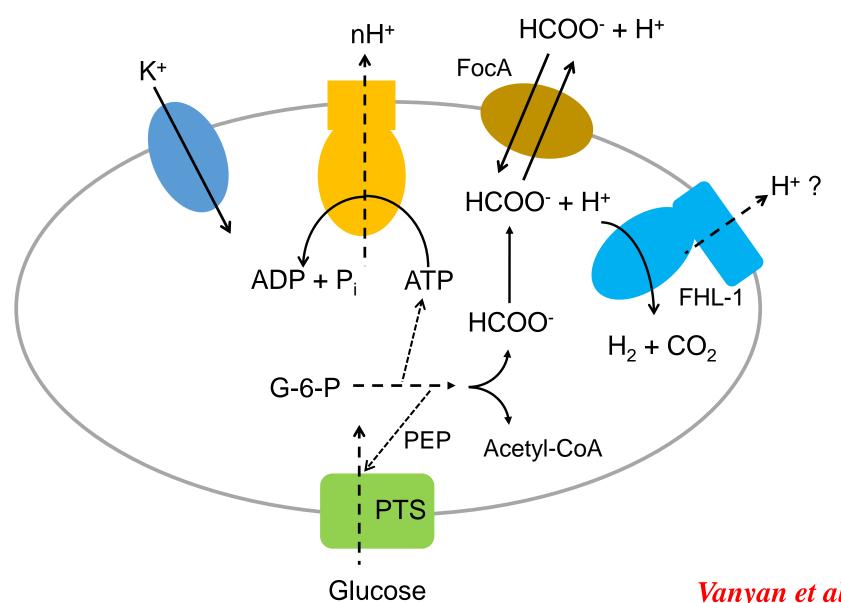
DH 4200 – all formate is effluxed out

DH 4300 – formate is not exported

DH 4400 – export but not import

Charge Balance

pH Balance

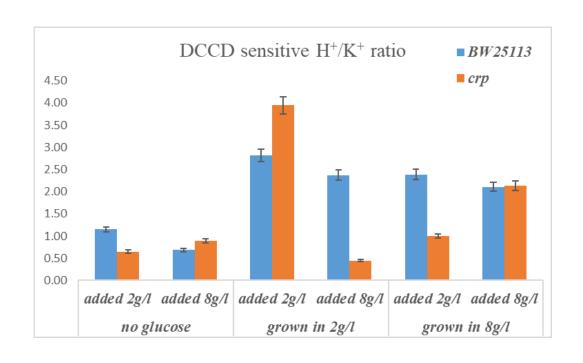


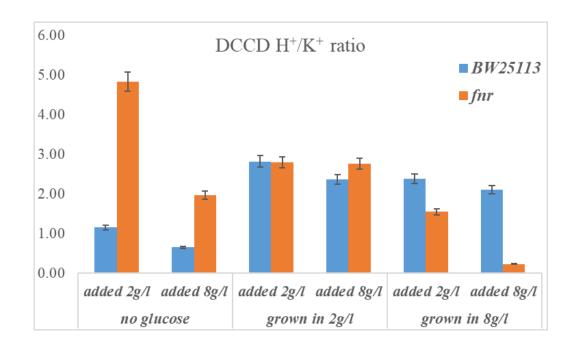


Prof. Gary Sawers (Germany)

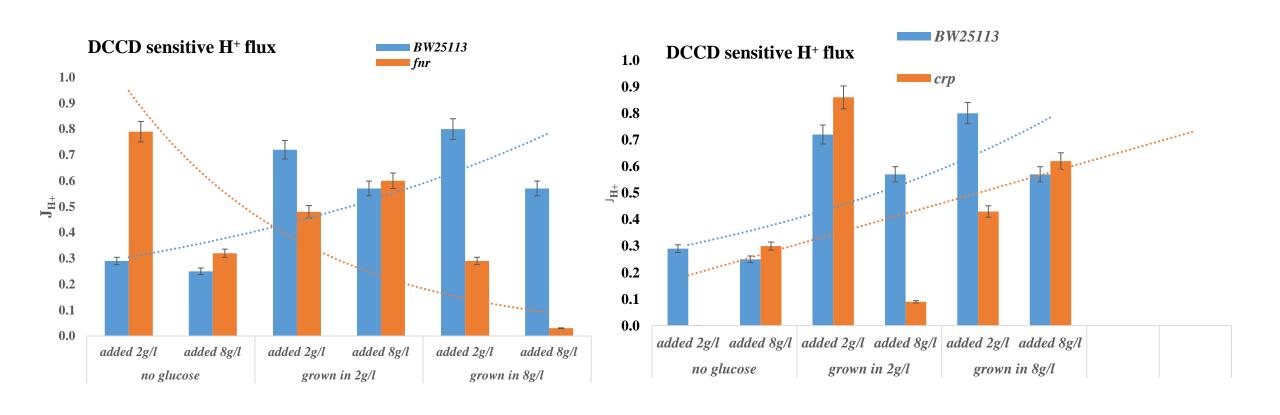
Vanyan et al., (2024). Archives of Biochemistry and Biophysics, 752, 109877.

Role of FNR and CRP on glucose dependent ion fluxes

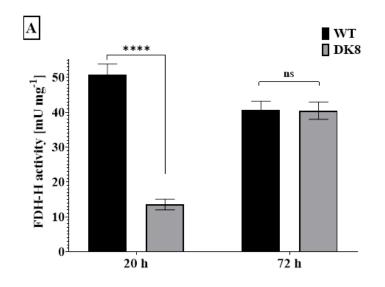


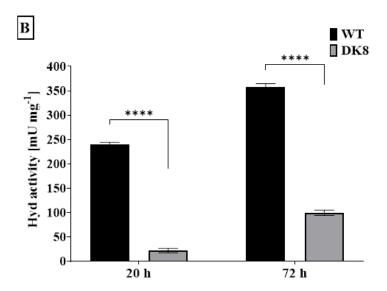


DCCD sensitive H⁺ flux in FNR and CRP mutants

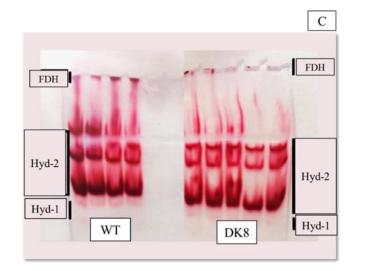


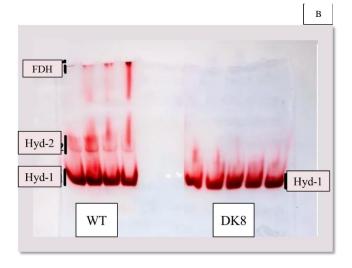
FDH-H and Hyd activities in DK8 mutant



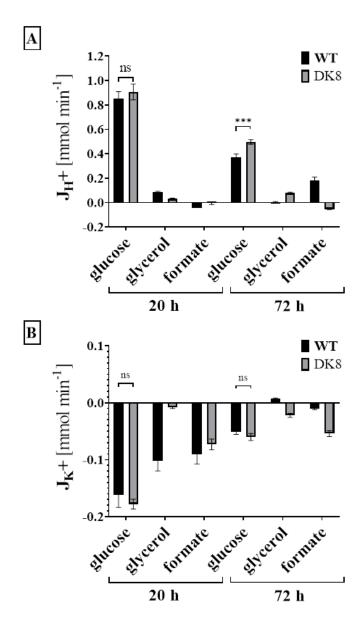


- ✓ Relationship between F_OF₁ and FDH-H at 20h but 72 h grown cells
- \checkmark Hyd activity depends on active $F_{O}F_{1}$ on both growth phases



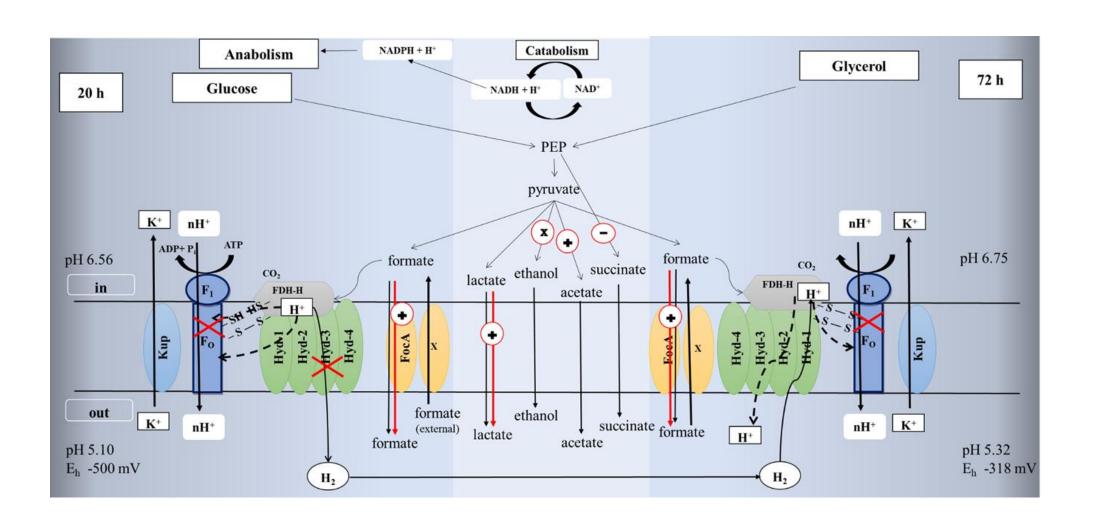


Ion fluxes in DK8 mutant

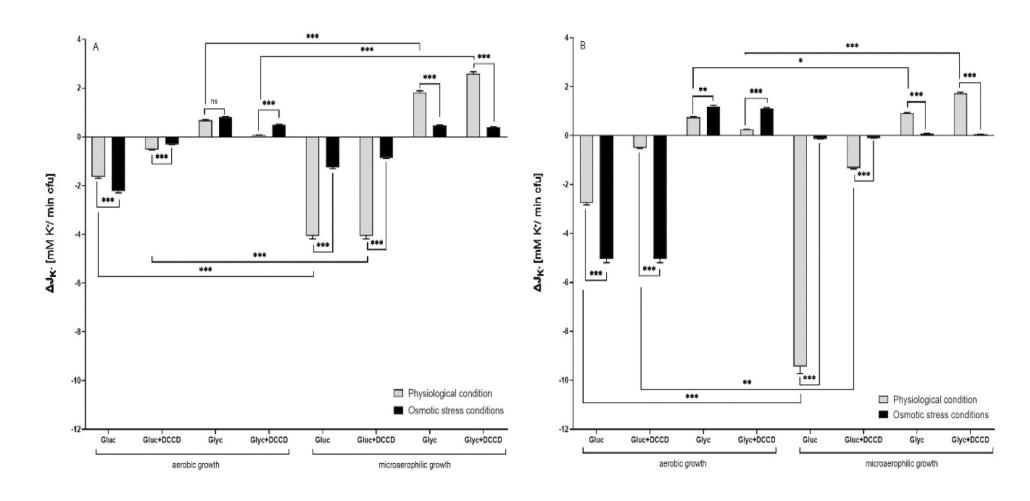


- ✓ Relationship between F_OF₁ and FDH-H at 20h but 72 h grown cells
- \checkmark Hyd activity depends on active F_0F_1 on both growth phases

Schematic working model of the cells grown 20 h and 72 h

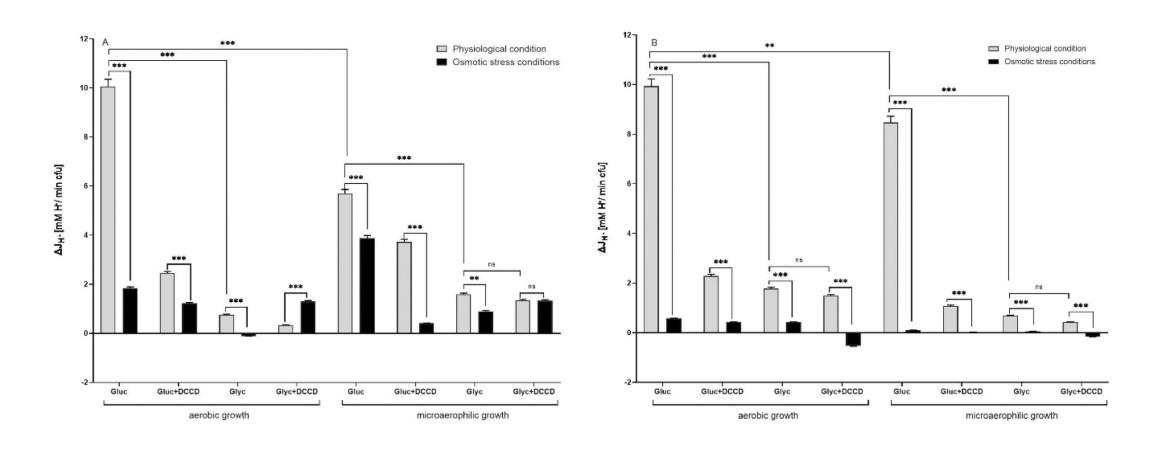


Potassium transport in yeasts



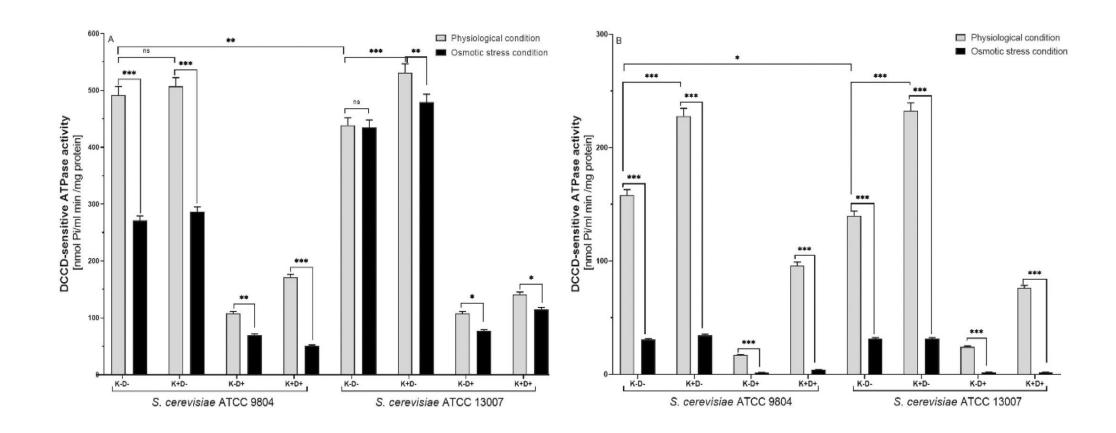
Shirvanyan, A., Trchounian, K. *Sci Rep* **14**, 23982 (2024).

Proton transport in yeasts



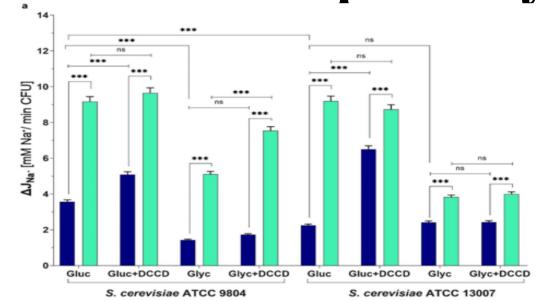
Shirvanyan, et al., Proc. Biochem. 133, 309-318 (2023).

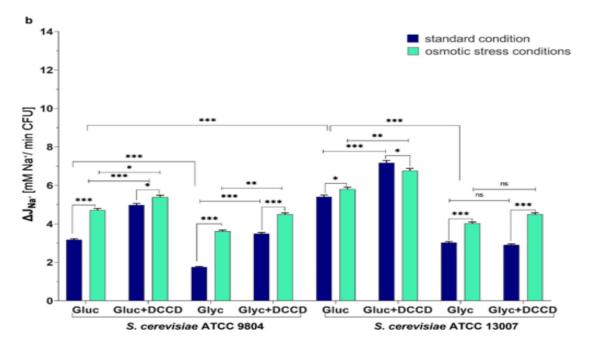
ATPase activity in yeasts



Shirvanyan, et al., Proc. Biochem. 133, 309-318

Sodium transport in yeasts



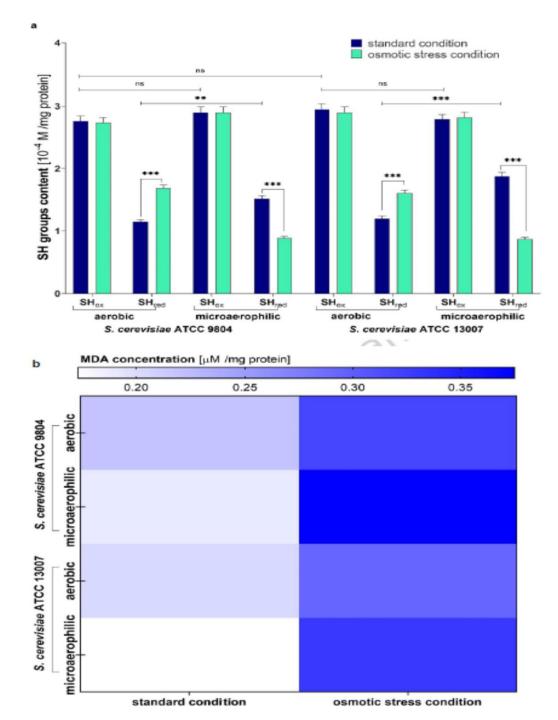


Shirvanyan, A., Trchounian, K. *Sci Rep* **14**, 23982 (2024).

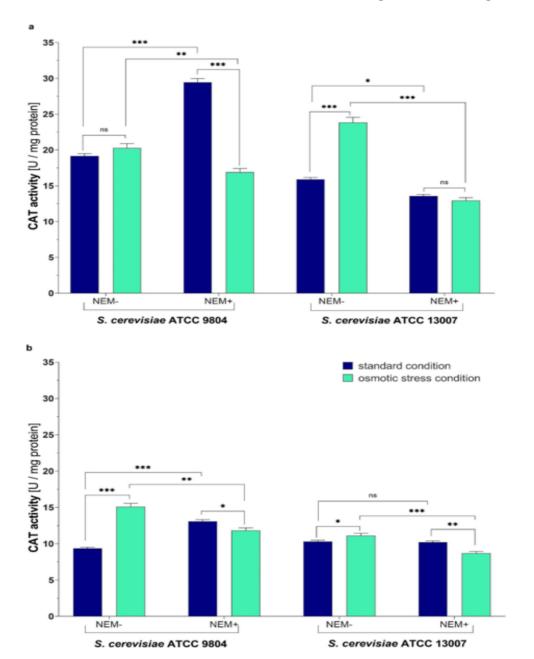
Thiol groups in yeasts

> Thiol groups are important for overcoming stress

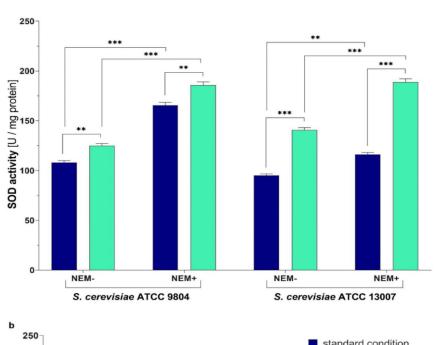
Shirvanyan, A., Trchounian, K. *Sci Rep* **14**, 23982 (2024).

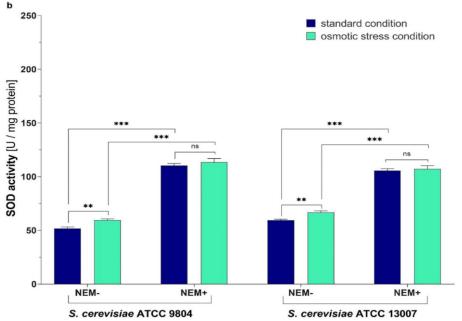


Catalase activity in yeasts



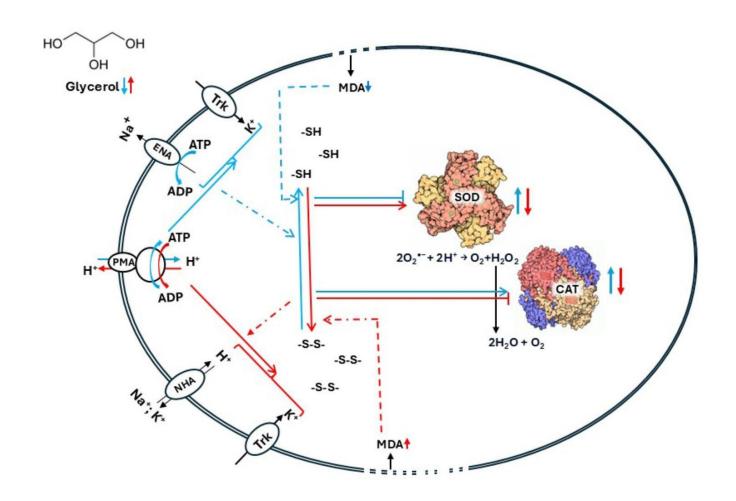
SOD activity in yeasts





Shirvanyan, A., Trchounian, K. *Sci Rep* **14**, 23982 (2024).

Schematic working model suggested for th yeasts



Questions to think for and revisit

- > Do we know much about adding glucose as media component for cell growth or?
- ➤ Are the main principles during respiration applicable for fermentation? Is it possible to apply the classical bioenergetics knowledge for anaerobic fermentative conditions?

- > Anaerobic hydrogenases are mimicking aerobic respiratory complexes?
- > Bacteria and yeast have similar ion gradient regulatory mechanisms for pH homeostasis?

Dr. Heghine Gevorgyan

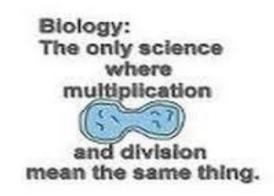
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