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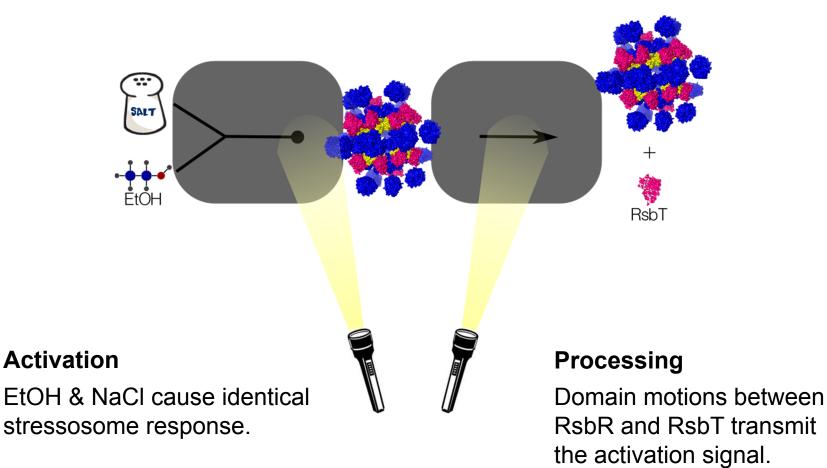
# **Conclusion of the Stressosome Models** Thomas Millat Dept. of Systems Biology & Bioinformatics 10<sup>th</sup> October 201 www.sbi.uni-rostock.de







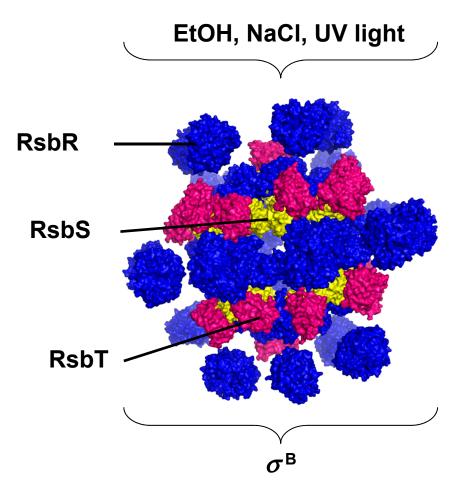
#### **Illuminating Processes of Stressosome Activation**







## Structure and Function of the Stressosome



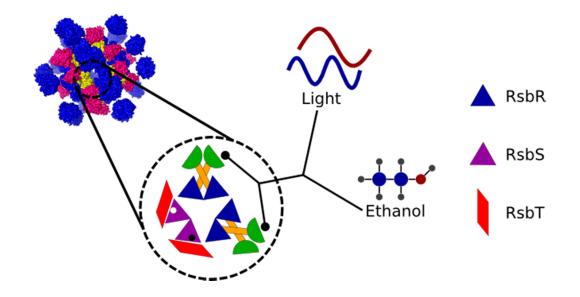
- Responsive to various stimuli
- Icosahedron with 3 proteins
- > Cytoplasmic RsbT activates  $\sigma^{\text{B}}$



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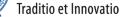


#### **Information Transmission**



- Signal reception: RsbR N-terminus
- Stimulus conversion: phosphorylations and interactions
- Information transmission: domain motions
- Effect: dissociation of RsbT







## **Open Questions**

- 1. How are different stimuli perceived?
  - Dose-Response measurements for different stimuli to test the effect of protein interactions.

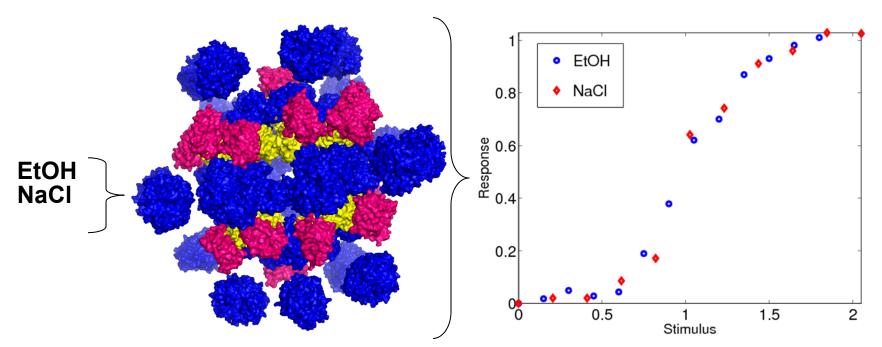
- 2. How is the signal transmitted from the sensor to the effector?
  - Geometric models of the icosahedral structure to find general rules of motion.





#### **Explaining Identical Dose-Response**

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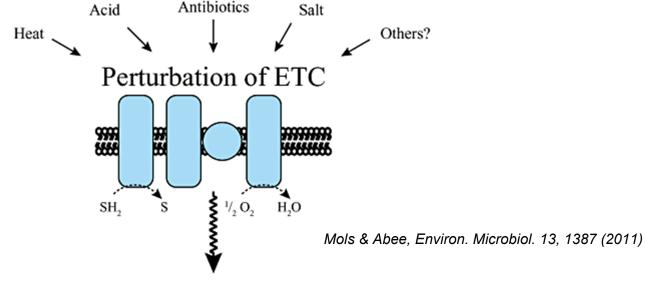
EtOH & NaCl are processed identically, because:

- 1. they cause identical modifications of the sensor.
- 2. they induce a common secondary signal transducer.





## Support for a Secondary Signal Transducer



-

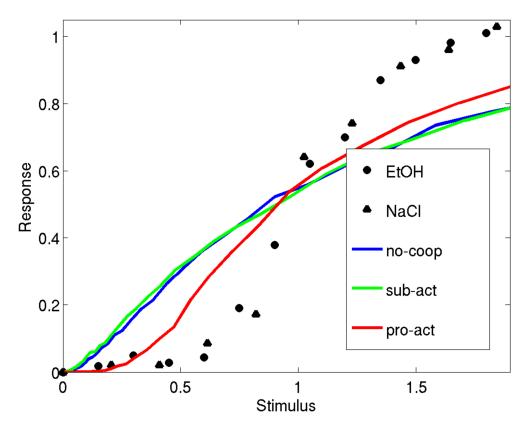
- Perturbation of the electron transport chain (ETC) generates reactive oxygen species (ROS)
- ROS have an intimate relation with the general stress response:
  - ROS are important inducers of the general stress response.
  - Majority of response proteins protect against or repair oxidative stress.

Höper et al., J. Bacteriol. 187, 2810 (2005); Reder et al., J. Bacteriol. 194, 3601 (2012)





#### In silico Reproduction of Dose-Response Curve



Sigmoidal Dose-Response Curve reproduced if:

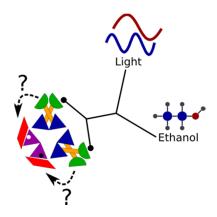
RsbR-P stimulates RsbT kinase activity.

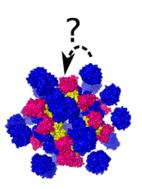
Liebal et al., Submitted for publication (2012)



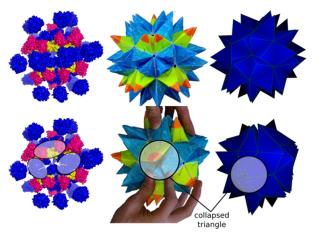


## Domain Motions can Transmit the Activation Signal





- No direct contact between signal receiver RsbR and effector RsbT.
  - How is the information transmitted?
- Geometric models of icosahedra show a concerted motion of domains.
- Collapse of a triangle of dimers possible with geometric distances preserved.
- YtvA performs scissor-like motions upon light stimulation.



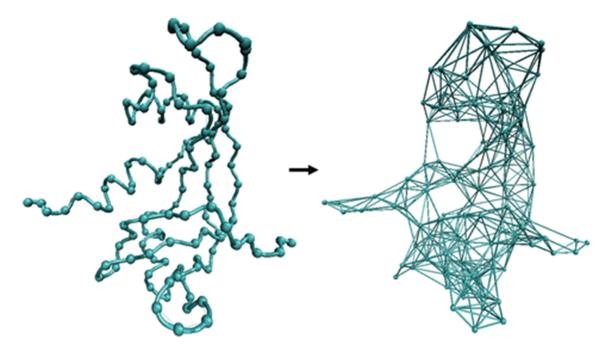






# Outlook: Moleculardynamic Modelling of Domain Motions

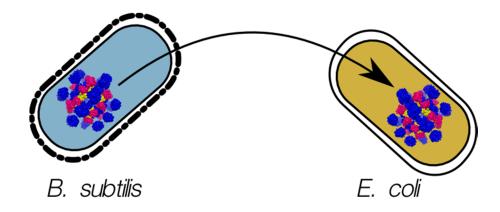
- Constructing an elastic network model of the stressosome.
- > Approximating structure and interactions by a low resolution grid.
- Analysing possible motions of domains.
- Cooperation with Rebecca Wade in Heidelberg, SysmoLAB.







## Vision: Light-Controlled Expression in E. coli



- Stressosome controlled σ<sup>B</sup> was transplanted to *E. coli*. Scott et al., Biochem. Biophys. Res. Commun. 257, 106 (1999)
- Fully light-sensitive stressosome was constructed in *B. subtilis*. van der Steen, J. et al. J Bacteriol, 2012, 194, 1708
- Light-sensitive YtvA was used for light induced expression in *E. coli*. Ohlendorf, R. et al. J Mol Biol, 2012, 416, 534



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stressosome structure & function

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