

Focused proteomic analysis of cell factories (proChassis)

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Ville Santala (TUT), FinSynBio opening seminar, 23.1. 2014

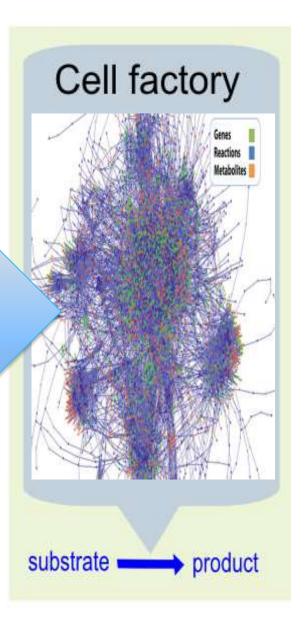
Synthetic biology

...engineering principles to biology...

Abstraction
Modularity
Standardization
Designing and Modeling

"The central premise: standardization of reusable biological components is the most efficient and effective way to engineering biology"

Gardner & Hawkins, 2013



Biological components

DNA

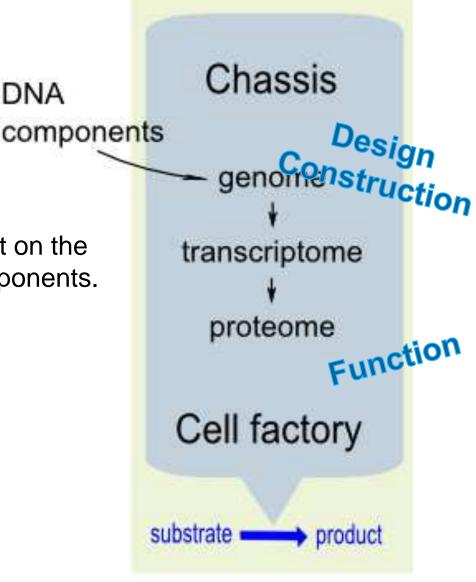
DNA components

Much research effort has been put on the characterization of individual components.

⇒ <u>Standardization</u>

A gap between components and the endogenous systems

Understanding of interactions!

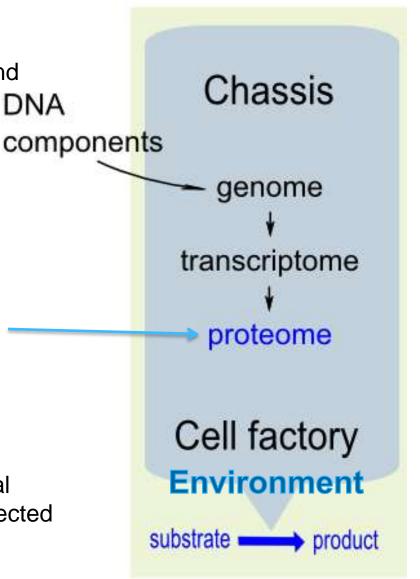




Native targets are used as general indicators (i.e. independent from the specific product formation pathway) for tracking different kind of perturbations and flux changes of the chassis.

NOVEL TOOLS
for quantification of
key proteins of chassis
AND components
in changing conditions.

Non-native targets encodes essential activities for the product and are selected from a well-characterized DNA components library.



$$x_i = \sum a_j r_j$$
 $(i = 1, 2, ..., m; j = 1, 2, ..., n)$

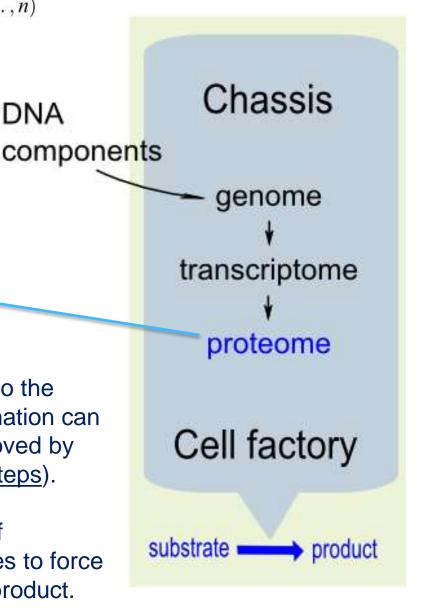
MODEL:
$$C_i^J = \frac{\mathrm{d}J}{\mathrm{d}E_i} \left(\frac{E_i}{J}\right)$$

DNA comp

quantification of key proteins of chassis AND components

By integrating quantitative informations to the model, factors affecting the product formation can be identified and the yields can be improved by removing bottlenecks (eg. <u>rate limiting steps</u>).

Eg. by changing the expression levels of components or knocking out native genes to force the flux from the substrate towards the product.





Chassis?

DNA components Acinetobacter baylyi ADP1

- 1) Easily genome-engineerable (allows high-throughput engineering)
 - Natural transformation & homologous recombination
- 2) **No IPR-issues** (allows open source policy)

With traditional strains, excessive IPR stacking can be problematic

3) Similarity to Escherichia coli

genome transcriptome proteome Cell factory substrate • product



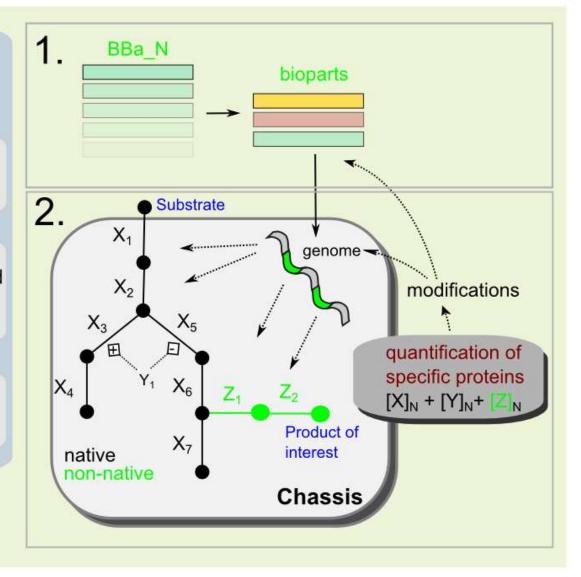
proChassis toolset

easily genome-engineerable chassis

quantitative monitoring of the expression of integrated bioparts and the key components of the host

computational approaches for system design and analysis

substrate product





System biology

proChassis

Synthetic biology

Metabolic engineering

Sustainable biofuel production