

EMERGENCE: A foundation for Synthetic Biology in Europe

WP1: General Networking activities

Fostering a community of knowledge

Vitor Martins dos Santos

**Systems and Synthetic Biology Group
Division of Microbiology
Helmholtz Centre for Infection Research
Braunschweig, Germany**

Consolidating the bases for a Synthetic Biology community

Foundations:

- Intellectual (concepts, abstraction, semantics...)
- Methodological (design, standardization, experimental / computational,...)
- Technological (IT, gene synthesis, microfluidics,...)

By:

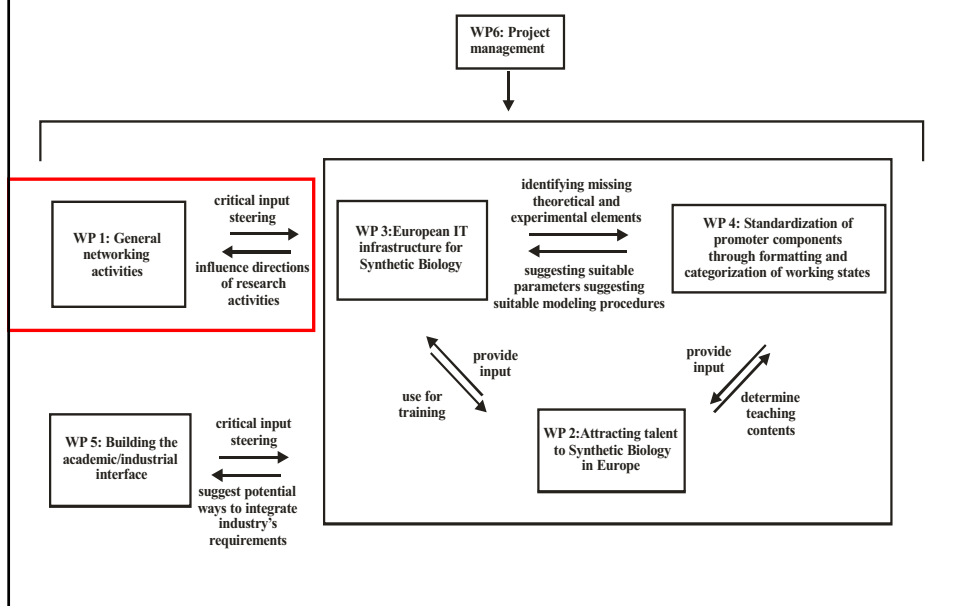
- recruiting the required competences from (not so) neighboring disciplines;
- exploiting synergies (competences, expertise, complementarity...);
- fostering transnational/transcontinental communication & cooperation
- promoting education at various stages (school, undergraduate, ..)
- embedding early developments into a meaningful societal / economical context

WP1: General Networking Activities

Objectives:

- a) To establish a networking platform for current and future synthetic biology projects
- b) To rapidly organize workshops for urgent issues in European synthetic biology
- c) To implement a Europe-wide, cross-disciplinary framework for discussion on the possibilities, needs, limitations, and implications of synthetic biology.
- d) To foster interactions with extra-European initiatives, with special emphasis on US, the Mid-East and Asia: Global knowledge space

Embedding WP1 in EMERGENCE



Description of Tasks I

Task 1: *Developing, maintaining, and evaluating a standardized meeting structure* that allows efficient review and distribution of the conclusions obtained at individual meetings.

Overarching, jointly with WP Management

Task 2: *Hosting workshops on development of the European IT infrastructure for synthetic biology, design tools for synthetic biology, and/or standardization of biological parts.*

Jointly with WP3 (IT infrastructure), WP4 (Design tools and Biological parts), Standardisation Issues (Overarching)

Description of Tasks II

Task 3: *Establishment of study groups on specific subjects relevant to synthetic biology*

“Foundational” technologies, including e.g. high-throughput genome minimization,
DNA synthesis),
potential of genetic circuits, modularity in proteins, handling noise & error
propagation in biological systems,
robustness in biological systems,
transferability of engineering foundations
.....

Description of Tasks II

Task 4: Platform for organizing thematic workshops/courses/meetings, resulting from maturation of study groups into specific workshops, courses, or small scientific meetings, or from initiatives from members of the advisory board or the steering committee.

Task 5: EMERGENCE will promote exchange and training visits between European and overseas participants, in particular with the Middle East and Asia, including:

- invitations for a number of leading scientists in the field to participate in study groups;
- seeking actively to participate in similar initiatives in those countries; and inclusion of Middle Eastern/Asian researchers in the EMERGENCE
- communication and dissemination pipelines.
- The participation of senior European synthetic biology scientists in Asian meetings will be particularly encouraged.

Deliverables Month 1-18

D1.1: Material and rules for standardized meeting structure in place for the first time (month 3). Responsible: HZI

D1.2: Report on the first workshop on development of the European IT infrastructure for synthetic biology (month 8) Responsible: HZI

D1.3: Report on the first workshop for design tools for synthetic biology (month 4) Responsible: CNIO

D1.4. Report on recommendations of the intra-consortium expert group on suitable promoter standardization formats (month 12) Responsible: CNB

Deliverables 18-36 month

D1.5: Updated material for the appropriate section in the quarterly Synthetic Biology Newsletter regarding tasks 2, 3, and 4 (months 3, 6, 9, 12, etc):

Responsible ETH

D1.6. Report on workshop on foundations of measurement statistics in synthetic biology (month 24)

D1.7. Document identifying “common European-Asian interests and ways to develop them” or similar document in place and signed by extra-European and European groups/organizations involved in synthetic biology (month 32)

Milestones and expected results

M1.1. Recommendations for the European IT infrastructure for synthetic biology are discussed and recommendations issued (month 3)

M1.2. Recommendations for design tools on the IT infrastructure are discussed and recommendations issued (month 4)

M1.3 First experiences with the study group format are reviewed by the steering committee after 6 months and by advisory board and steering committee after 12 months and the format is adapted, if necessary (month 6, 12)

M1.4. Recommendations on standardization of biological parts are discussed (month 11)

M1.5. Recommendations on measurement systems in synthetic biology are discussed (month 24)

M1.6. Steering committee and advisory board decide whether the **critical mass in Europe-Asian relations in synthetic biology** has been reached and drafting a “common interests” document is going to be useful (month 24)

D1.1 - Material and rules for standardized meeting structure

Web-based template document:

Definition of the theme and Scope

The need for the SynBio community and goals

Implementation plan (size, mode, participants)

Timeline

Financing possibilities

Process:

Submission to Steering committee (WP-leaders, Coordinator)

Eg. IT: A. Valencia; Teaching: Sven P.; INdustry: L. Pasamontes

D1.1 - Material and rules for standardized meeting structure: examples themes

context-independent biological systems/modules

microfluidics technologies / single cell measurements

minimal genomes / minimal systems

what to measure / how to measure?

**design concepts
how can we handle "systems"
(made of parts)?**

D1.2 -Report on the first workshop on development of the European IT infrastructure for synthetic biology

Workshop Computational Infrastructure and Methods for Synthetic Biology The Eighth Annual BioPathways Meeting

Vítor Martins dos Santos
Vincent Schachter
Vincent Danos
Joanne Luciano
Aviv Regev
Eric Neumann

July 19-20, 2007
Satellite Meeting ISMB-ECCB 2007
Vienna, Austria

| | | |
|--|---|---|
| 7:30 – 8:45 | Registration | |
| 9:00-9:10 | Vincent Schachter Genoscope, Evry & BioPathways Consortium | Opening remarks |
| Session 1: Computational Methods and Infrastructure for Synthetic Biology | | |
| Chairman: Vítor Martins dos Santos | | |
| 9:10-09:30 | Vitor Martins dos Santos, Helmholtz Center for Infection Research, Braunschweig | EMERGENCE: a Foundation for Synthetic Biology in Europe |
| 9:30-10:00 | Alfonso Valencia, CNIO, Madrid | Bioinformatics tools to help in the design of biological systems |
| 10:00-10:30 | Jörg Stelling, ETH, Zürich | Formal tools for Model-Based Synthetic Biology |
| 10:30-11:00 | Coffee Break | |
| 11:00-11:30 | Randy Rettberg, MIT, Cambridge | The MIT registry of parts and devices |
| 11:30-12:00 | Alfonso Jaramillo, Ecole Polytechnique, Paris | Model-based design of genetic circuitry |
| 12:00-13:00 | Lunch | |
| Session 2: Network Reconstruction & Analysis (part 1) | | |
| Chairman: Vincent Schachter | | |
| 13:00-13:45 | Florence d'Alche-Buc, University of Evry | Supervised Inference of Protein-Protein Interaction Networks |
| 13:45-14:30 | Jason Ernst, Carnegie Mellon University | Reconstructing Dynamic Regulatory Maps |
| 14:30-14:50 | Tijana Milenkovic and Natasa Przulj, Irvine, University of California | Uncovering Biological Network Function via Graphlet Degree Signatures |
| 14:50-15:10 | Kam Dahlquist, Loyola Marymount University | Mathematical Modeling of the Transcriptional Network Controlling the Environmental Stress Response in <i>Saccharomyces cerevisiae</i> |
| Session 3: Databases & Software Tools | | |
| Chairman: Joanne Luciano | | |
| 15:10-15:30 | Ozgun Babur, Bilkent University | PATIKWeb Components for Microarray Data Analysis & Advanced Graph-Theoretic Querying |
| 15:30-16:00 | Coffee Break | |
| 16:00-16:20 | Richard Adams, University of Edinburgh | The Edinburgh Pathway Editor |
| 16:20-16:40 | Esther Schmidt, EBI, Cambridge | Reactome - a knowledgebase of biological pathways |
| 16:40-17:20 | Eric Neumann, Teranode Corp. | A Genome - Phenome Integrated Approach for Mining Disease-Causal Genes using Semantic Web |
| Round Table Discussion | | |
| 17:20-18:30 | IT Infrastructure & Computational Methods for Systems and Synthetic Biology | |

| Session 4 : Network Reconstruction & Analysis (part 2) | | |
|---|--|---|
| Chairman: Eric Neumann | | |
| 9:00-9:45 | Peter Karp, SRI International | Gene Regulation in EcoCyc and Pathway Tools |
| 9:45-10:30 | Jerzy Tiurny, University of Warsaw | Identification of functional modules from ancestral protein-protein interactions |
| 10:30-11:00 | Coffee Break | |
| 11:00-11:20 | Rainer Koenig, DFKZ, Heidelberg | Using gene expression data and network topology to detect substantial pathways, clusters and switches |
| 11:20-11:40 | Hanif Khalak | Microarray-based Class Modeling and Prediction using Set-Enrichment Analysis |
| 11:40-12:00 | Sol Efroni, NIH/NCI | Identification of Key Processes underlying Cancer Phenotypes using Biologic Pathway Analysis |
| 12:00-13:00 | Lunch | |
| 13:00-13:45 | Eytan Ruppin, Tel-Aviv University | Genome Scale Studies of Robustness and Annotation of the Yeast Metabolic Network |
| 13:45-14:30 | Fengzhu Sun, University of Southern California | Network motif identification in stochastic networks |
| Session 5: Evolution of pathways and networks | | |
| Chairman: Toni Gabaldón | | |
| 14:30-15:15 | Simon Lovell | Protein-protein interactions and their networks: can they tell us about biology? |
| 15:15-15:35 | Natalia Maltsev | Co-evolutionary analysis of Metabolic Pathways and Enzymes in PUMA2 and Chisel systems |
| 15:35-16:00 | Coffee Break | |
| 16:00-16:45 | Toni Gabaldón | Evolution of metabolic systems: insights from comparative genomics |
| 16:45-17:30 | Philip Kim | Relating three-dimensional structures to protein networks provides evolutionary insights |
| Round Table Discussion | | |
| 17:30-18:30 | Network Reconstruction and Evolution | |
| End of meeting | | |

D1.3 - Report on workshop for design tools for synthetic biology (CNB)

Satellite meeting to the ESF – EMBO on SynBio

November 2007

(Alfonso, Jörg, Randy, etc)

Report being drafted (CNIO)

D1.4 - Report on recommendations of the intra-consortium expert group on suitable promoter standardization formats (CNB)

VDL – Report in preparation

**Silva-Rocha R, de Lorenzo V.
Mining logic gates in prokaryotic transcriptional
regulation networks.
FEBS Lett. 2008 Apr 9;582(8):1237-44.**

**D1.4 -Updated material for the appropriate section
in the quarterly Synthetic Biology Newsletter
regarding tasks 2, 3, and 4**

Frauke Greve / Sven Panke

**Newsletters Dec 2006, June 2007, Dec 2008, June
2008**

**Includes list of conferences, research highlights,
press releases, funding activities**

Activities towards Task 4 (European Networking)



UNIVERSITAT DE BARCELONA



RESEARCH CONFERENCES

ESF-UB Conference in Biomedicine

European Conference on Synthetic Biology (ECSB): Design, Programming and Optimisation of Biological Systems

Hotel Eden Roc, Sant Feliu de Guixols • Spain
24-29 November 2007

Chair: **Alfonso Valencia**, CNIO Madrid, ES

Co-Chairs: **Natalio Krasnogor**, University of Nottingham, UK

- **Sven Panke**, ETH, Zürich Institute of Process Engineering, CH

- **Victor de Lorenzo**, Centro Nacional de Biotecnología, Madrid, ES

www.esf.org/conferences/07241

Activities towards Task 4 (European Networking)

Series of Workshops on different aspects of SynBio:

- Biofine (Tessy), Freiburg April 10, 2008
- Genopole (Jaramillo), 26-27 June, 2008
- IRGC Workshop Session on the Risk Governance of Synthetic Biology (26 & 27 June - Geneva, Switzerland)
- Stakeholder meeting Roadmap SynBio (Tessy), 10 June 2008
- ESF workshop on Minimal Systems (with A. Moya), in planning
- Etc.....

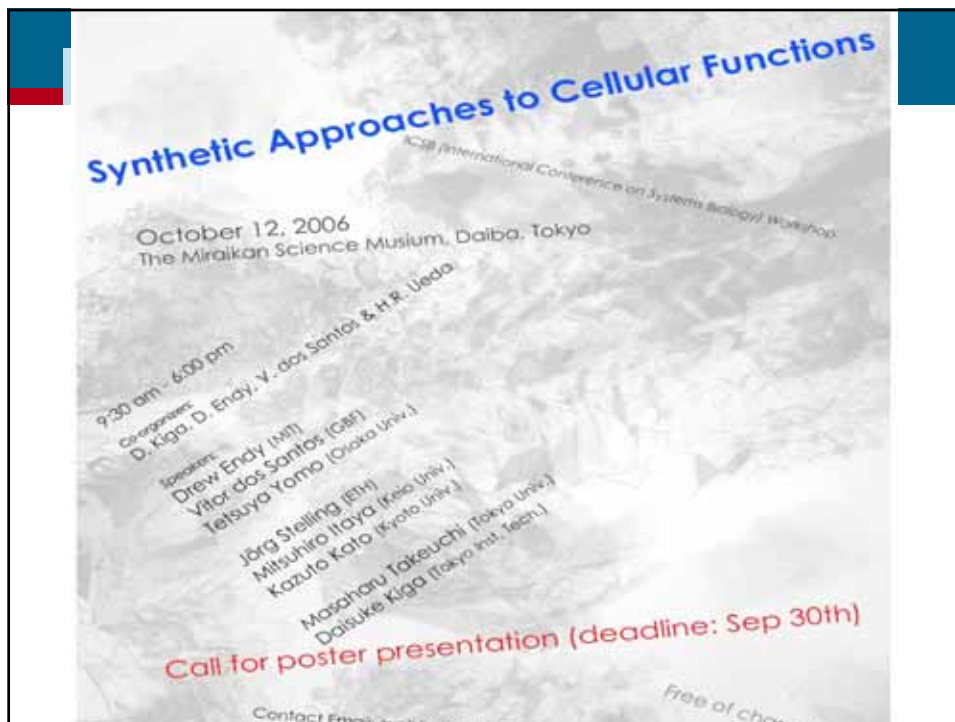
Activities towards Task 4 (Global Networking)

Workshop on:

Synthetic Approaches to Cellular Functions, Tokyo, 13 October 2006

Organised jointly by D. Kige (JP), H. Ueda (JP), D. Endy (US), Martins dos Santos („EU“)

About 120 worldwide attendants, 50+ posters. NEST-PATHFINDER SB projects presented. Overwhelming reaction



Future networking activities Asia

Sino-German Exploratory Workshop on Synthetic Biology, Hangzhou, China, 2008. Couple to Probiactys (EU) and perhaps other projects

To be organised jointly with Huanming Yang (Beijing Genome Institute, CN)

Exchange of students/ scientists:

China (2 students 7 month each plus 2 scientists 1 week in 2007)

India (2 Students 4 month each, plus scientist 1 week 2008)

Explorative project in Israel on digital evolving microbial communities

Indian - EU workshop on Synthetic Biology (January or September 2009)

ESF-JSPS Frontier Science Conference Series for Young Researchers
(Synbio tentative for 2009)

How shall we proceed?

Report on the identification of scientific & infrastructure bottlenecks in SB (jointly WP4 & WP3)

Study groups: bottom-up, priorital themes?

Possible themes:

context-independent biological systems/modules

microfluidics technologies / single cell measurements

minimal genomes / minimal systems

what to measure / how to measure?

design concepts

how can we handle "systems" (made of parts)?

.....

Thematic Workshops: IT and Standardization. Time plan?

Exchange visits?

| Future activities, other | |
|--------------------------|--|
| | |

Consolidating the bases for a Synthetic Biology community

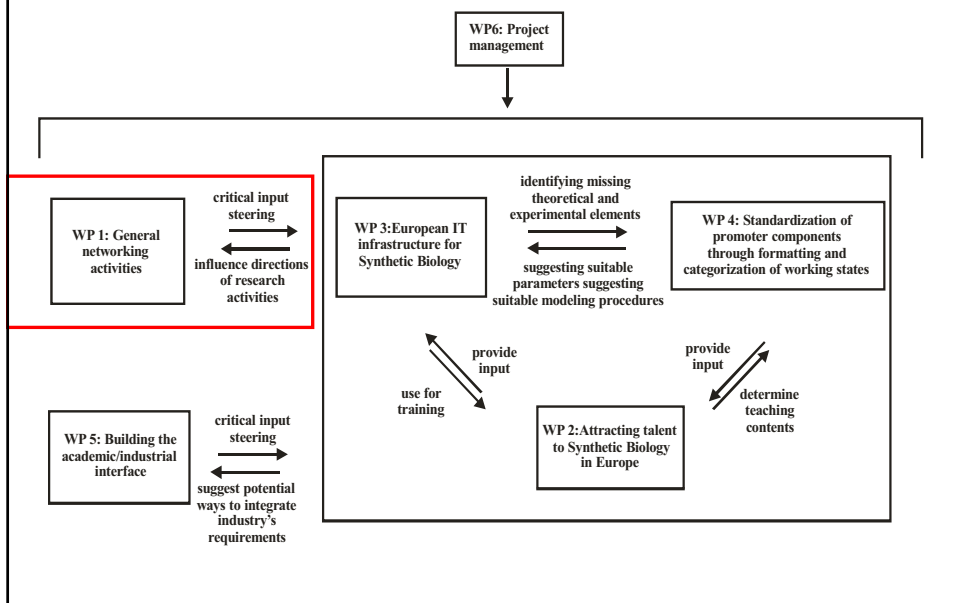
Foundations:

- Intellectual (concepts, abstraction, semantics...)
- Methodological (design, standardization, experimental / computational,...)
- Technological (IT, gene synthesis, microfluidics,...)

By:

- recruiting the required competences from (not so) neighboring disciplines;
- exploiting synergies (competences, expertise, complementarity...);
- fostering transnational/transcontinental communication & cooperation
- promoting education at various stages (school, undergraduate, ..)
- embedding early developments into a meaningful societal / economical context

Project Structure



Emergence WP3
A European IT infrastructure for Synthetic
Biology

M.A. Marchisio
Zurich, 28/05/08



Starting point: the *MIT Registry of Standard Biological Parts*

(<http://partsregistry.org>)

Goal: building a European mirror of the Registry containing computational tools

--> to retrieve biological information from other databases;

--> to design circuits made of the Standard Biological Parts.

Information integration

(CNIO – I. Cases, A. Valencia; MIT – R. Rettberg)

DAS (Distributed Annotation System) server solution:

- easier implementation;
- cooperation with other projects (Biosapiens NoE).

1st step: testing the Registry connectivity with other databases

Three prototype DAS servers implemented on the MIT site:

- reference (sequence and IDs);
- annotation1 (structure, functions, etc.);
- annotation2 (Uniprot as a reference server; gets Uniprot ID, returns part IDs).

2nd step: development of new tools to access and visualize content of biological databases

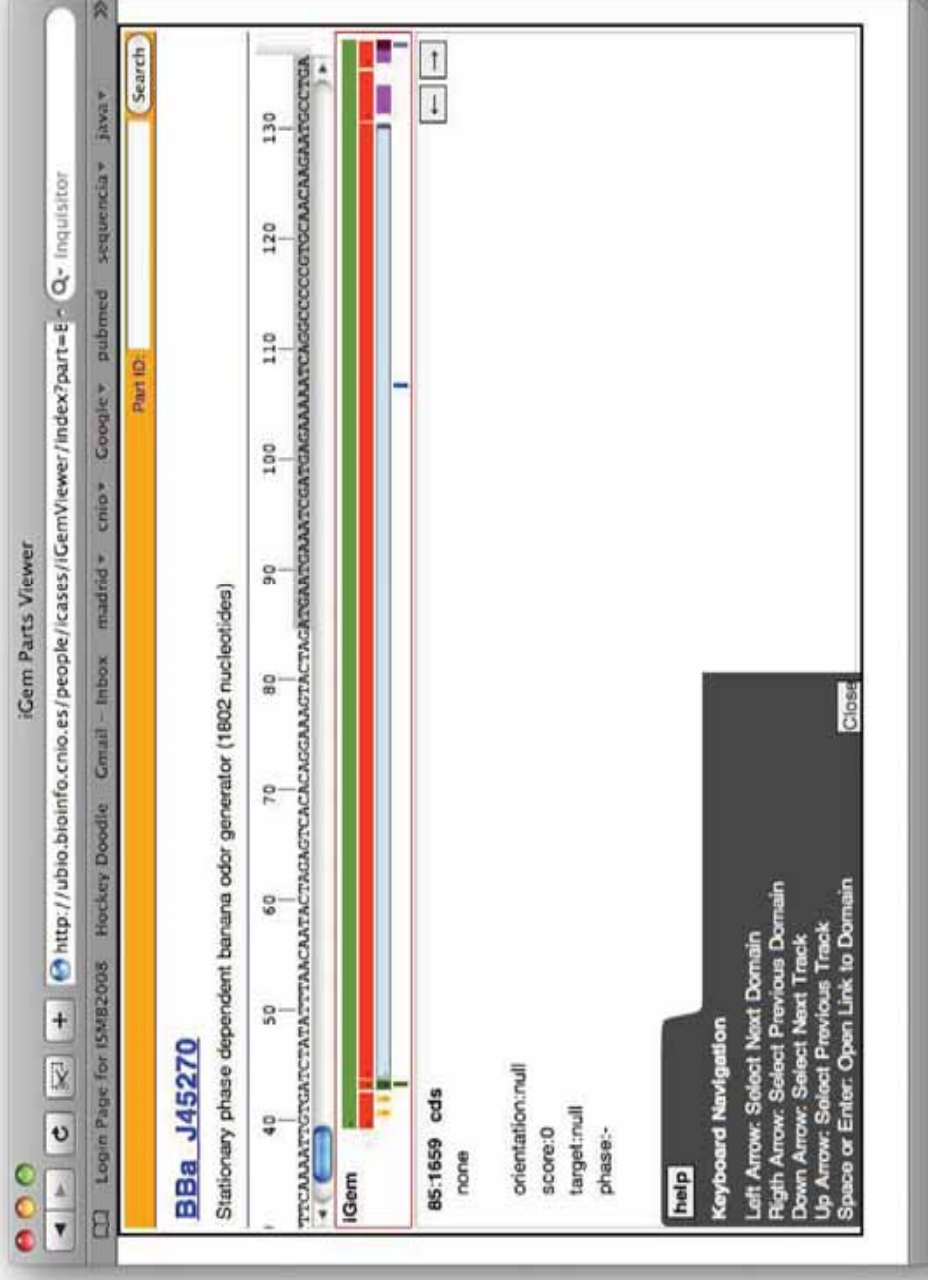


(<http://cargo2.bioinfo.cnio.es/>)

CARGO (Cases et al., NAR 2007) extended to handle information from the three prototypes DAS installed on the MIT site.

3rd step: realization of a pilot application.

IGEM Part Viewer: can visualize part content and return a possible link to Uniprot.



by courtesy of I. Cases

Circuit simulation

(ETHZ – M.A. Marchisio, J. Stelling)

Main activity: building of a “*drag & drop*” tool for circuit design

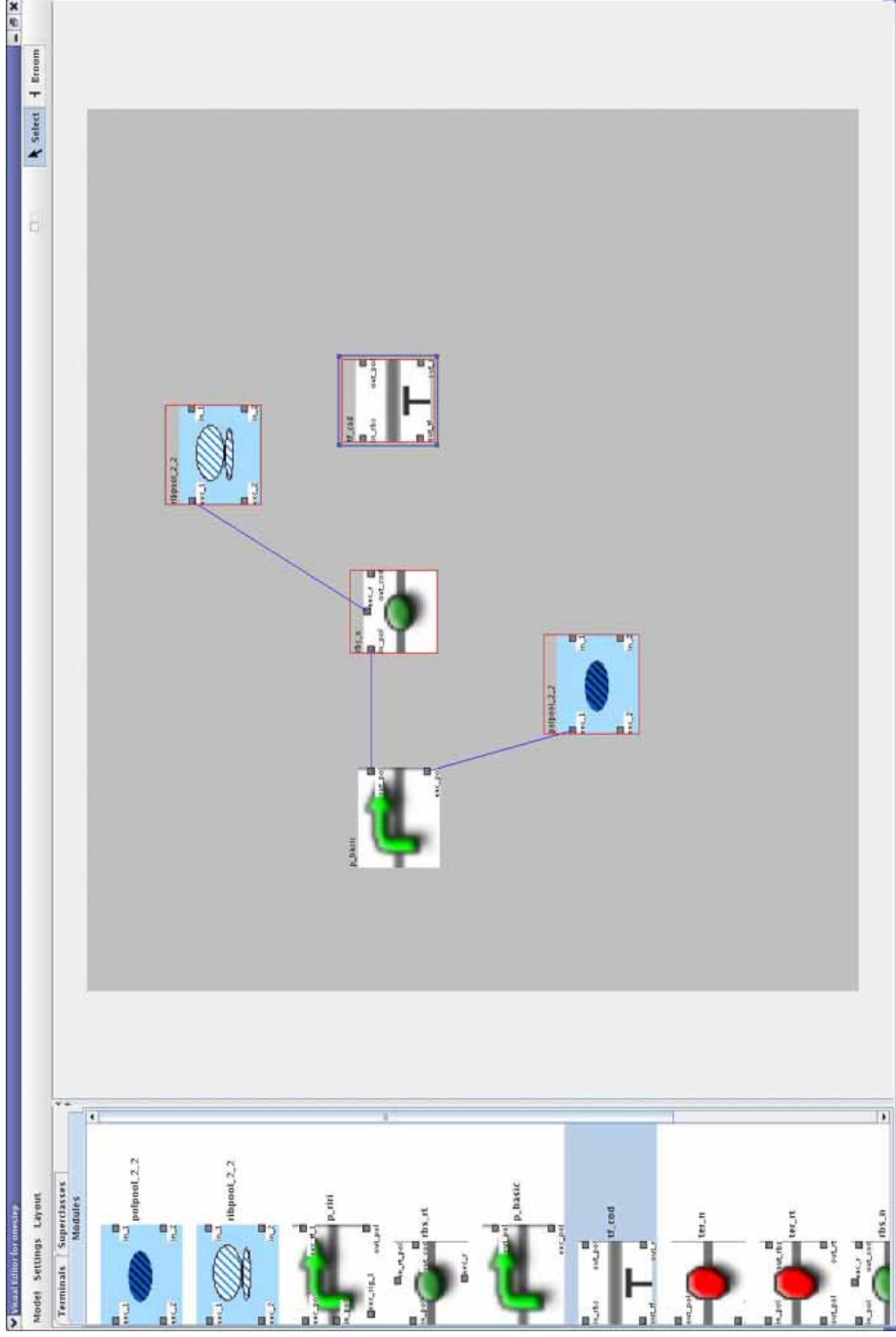
Main **problem**: composability of Registry parts

(common signal carriers definition)



Canvas provided by **ProMoT**

(<http://www.mpi-magdeburg.mpg.de/projects/promot/>)



Model/tool features

ODE based model for every basic Registry parts.

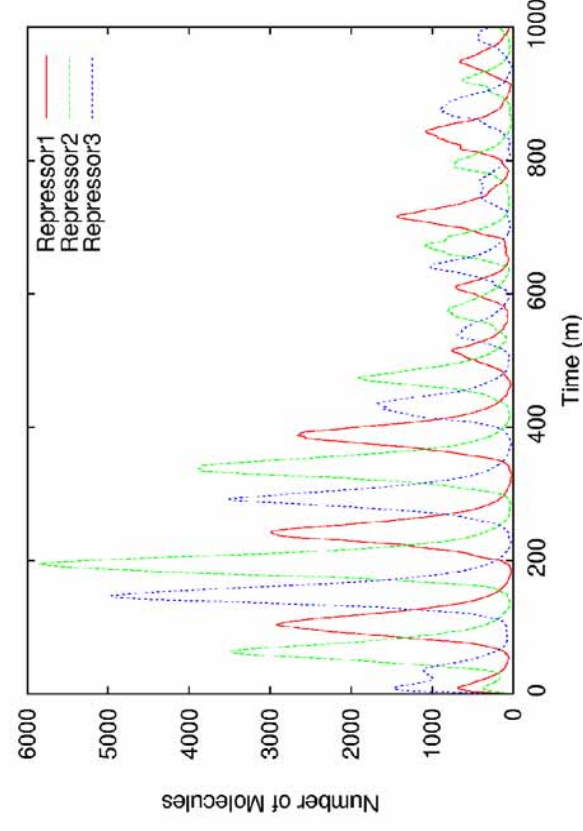
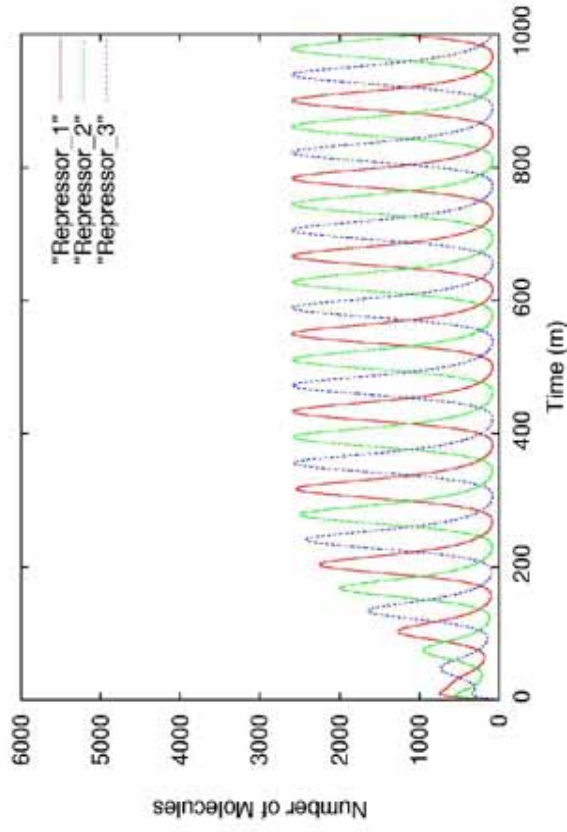
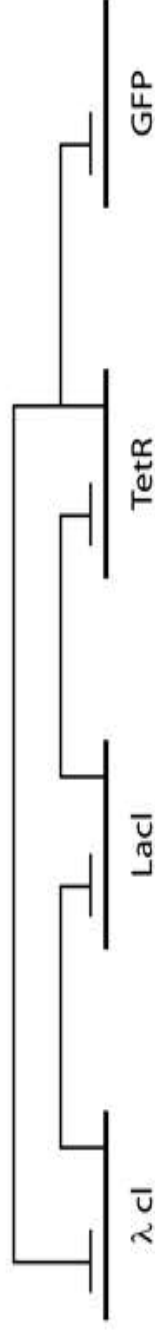
Set of Perl scripts to generalize parts' construction.

Devices (complex Registry part) realized by assembling basic parts.

Circuit files exportable into Matlab or SBML (level 1 or 2) format.

Possibility of running both deterministic and stochastic simulations.

Example: the Repressilator*



(*) Elowitz, M. B. and Leibler, S. (2000) *Nature* **403**, 335-338.

How to go on?

Definition of the European Registry mirror basic components.

They might be:

- 1) the part browser (CARGO + IGEM Part Viewer);
- 2) the access to different databases (CARGO + DAS servers);
- 3) the canvas for circuit design (ProMoT + part models).

Problems:

- necessity of a “source” of parameter values;
- integration of part generation and circuit design;
- putting together the information and the simulation side.

Biobrick Standardization



Synthetic Biology Concepts



- **Standardization** ... transferable & predictable **modules**, conditions, methods
- **Decoupling** ... separate complex design into simpler problems tackled independently
- **Abstraction** ... hide detail behind **standard interfaces** and **signal carriers**
- **Open exchange** ... organize specifications and methods

[D. Endy (2005) Foundations for engineering biology. Nature 438]

Content

- ① BBF Standardization Process
- ② from Pobol to a web of registries
- ③ BrickIt – open source biobrick management
- ④ A European web of registries project?

Standardization

Overview

http://openwetware.org/wiki/The_BioBricks_Foundation:Standards

[article](#) [talk](#) [edit](#) [history](#)

The BioBricks Foundation:Standards



[HOME](#) [OUR GOALS](#) [BOARD MEMBERS](#) [FAQ](#) [DONATIONS](#) [CONTACT](#)

Please visit:

- [The BioBricks Foundation:Standards/Technical](#) for the wiki notes from the **BBF Technical Standards** mailing list.
- [The BioBricks Foundation:Legal](#) for wiki notes from the **BBF Legal Standards** mailing list.
- [The BioBricks Foundation:MailingLists](#) to subscribe to **BBF** mailing lists, or browse the mailing list archives.

Standardization Process

1. two parties from different locations
2. demonstrate it's working
3. write it up
4. request RFC # on mailing list
5. others comment / revise
6. BBF enacts standard

Active Technical Standards Projects

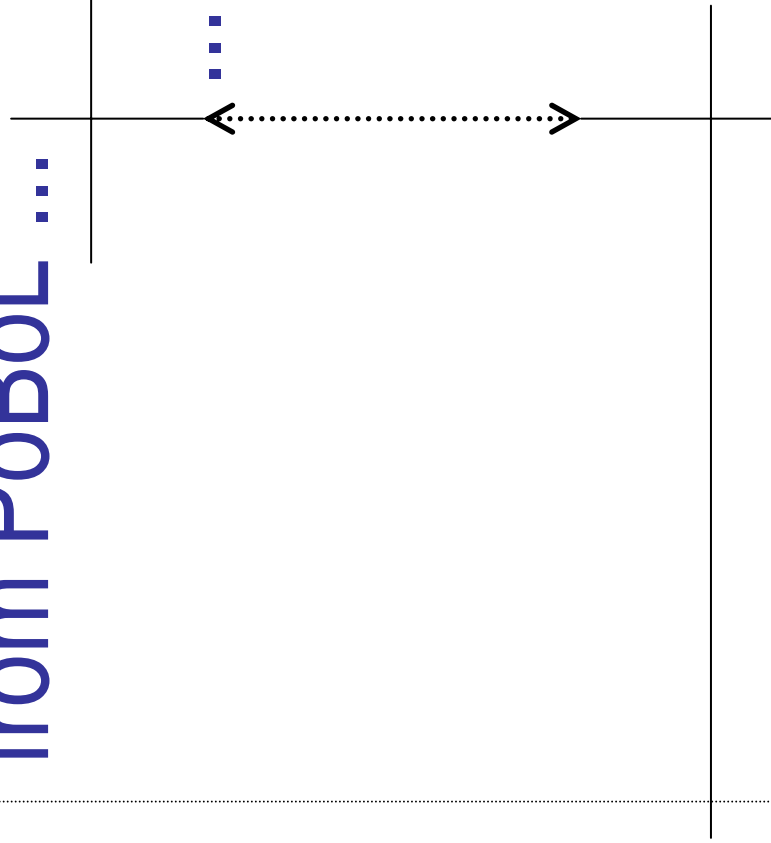
- Biobrick Formats (aka Physical Composition)
- Measurement standards
- Data exchange standards
- Technical Resources
- E.coli promoter standard

BBF Legal Working Group

- BBF Standards Mailing List
 - BBF Standards Wiki
-

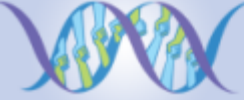
from PoBoL ...

... to a web of registries



Data Exchange Wiki

http://openwetware.org/wiki/The_BioBricks_Foundation:Standards/Technical/Exchange



OpenWetWare
Share your science.

navigation

- [Main Page](#)
- [Recent changes](#)
- [Help](#)
- [contact](#)
- [Chat](#)

research

- [Materials](#)
- [Protocols](#)
- [Resources](#)

search

toolbox

- [What links here](#)
- [Related changes](#)
- [Upload file](#)
- [Special pages](#)
- [Printable version](#)
- [Permanent link](#)
- [Cite this article](#)

[article](#) [talk](#) [edit](#) [history](#) [Log in / create account](#)

The BioBricks Foundation:Standards/Technical/Exchange

< [The BioBricks Foundation:Standards](#) | [Technical](#)

Biobrick Data Exchange Standards: This working group aims to define formats / technologies for the description of biobricks and the exchange (or networking) of biobrick-related data. This document is part of the ongoing discussion on the technical standards mailing list. The main questions to tackle are:

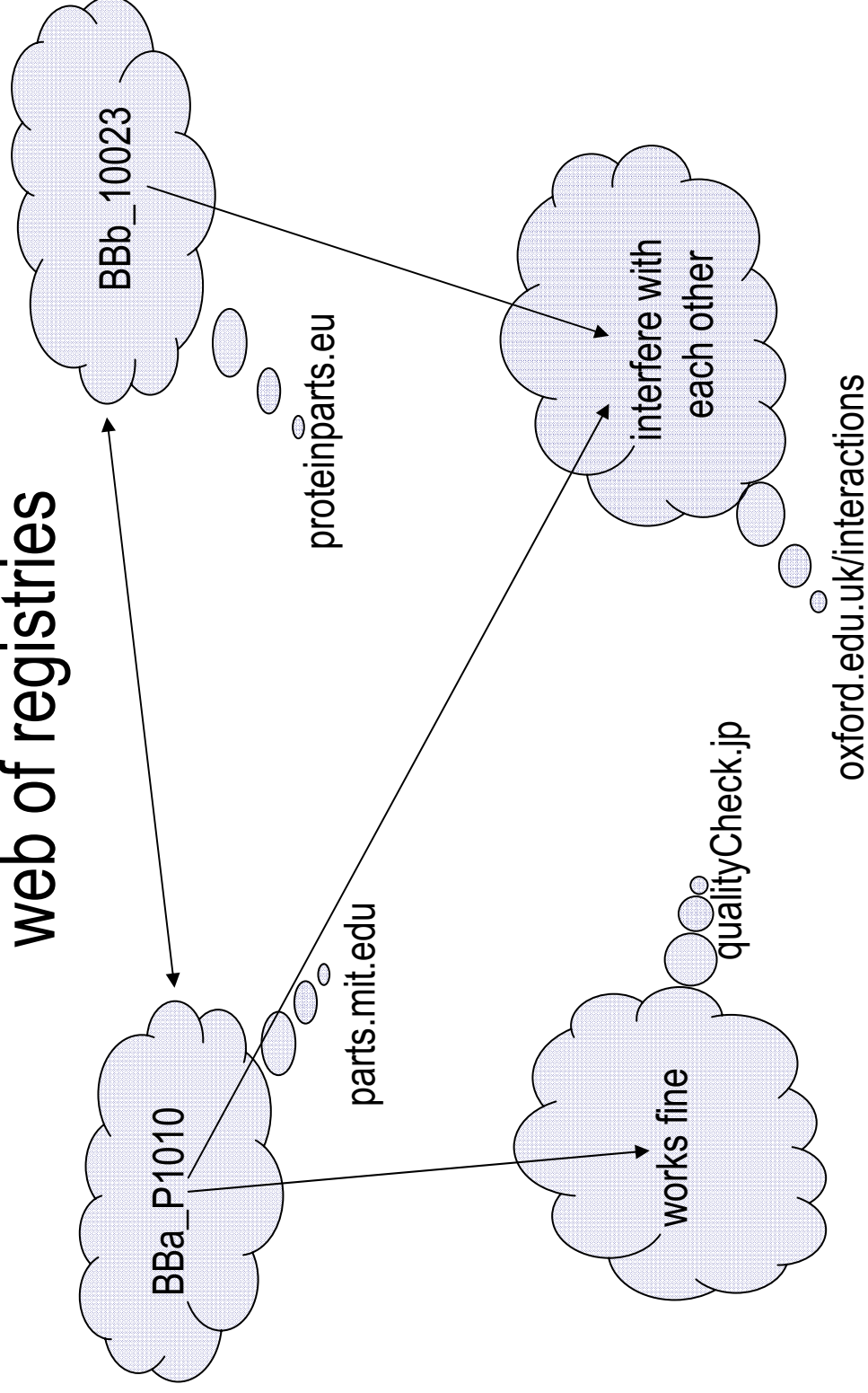
- Aim -- goal and application scenarios for this standard
- Biobrick definition -- What is a Biobrick?
- Data model -- What is the data model needed to describe a biobrick?
- Technology -- What is the best format / technology for exchange?

Contents [hide]

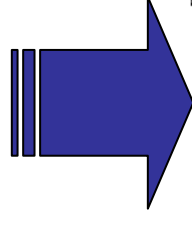
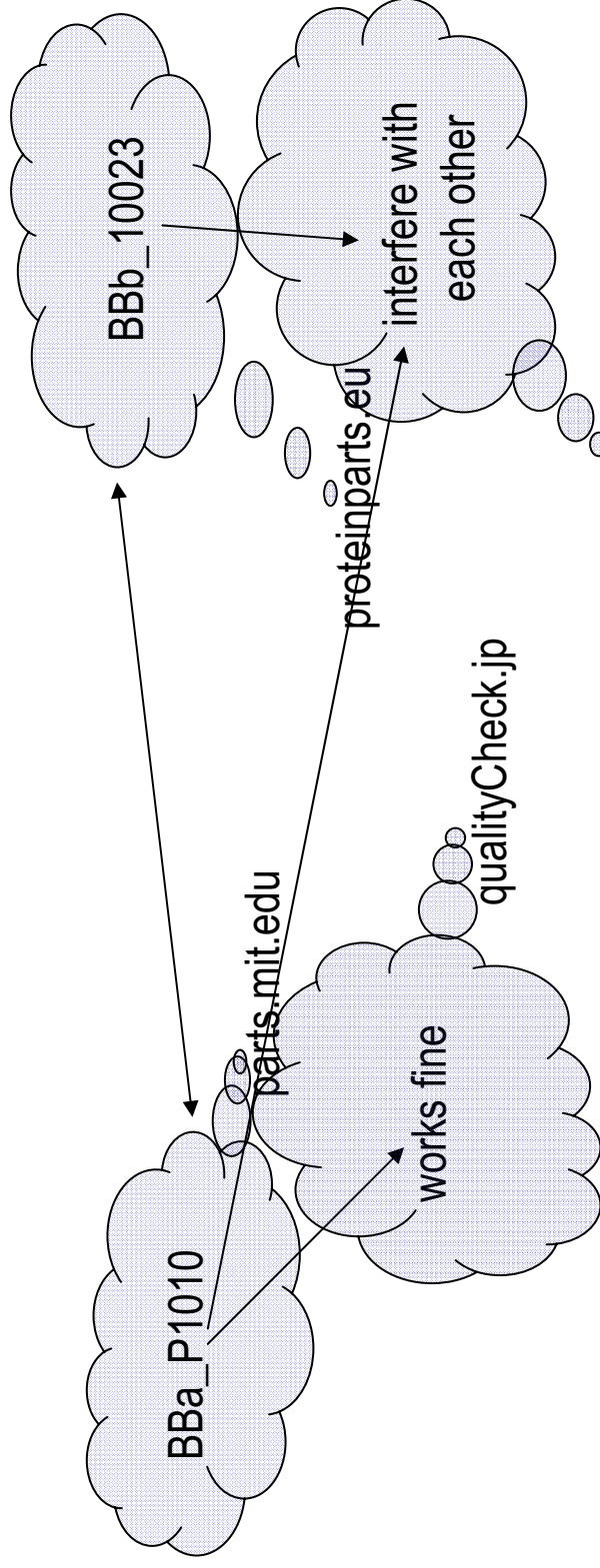
- 1 Aim / Application scenarios for this standard
 - 1.1 Concise aims
 - 1.2 Application scenarios [please discuss]
- 2 What is a Biobrick?
 - 2.1 Biobrick Definition
 - 2.2 Background
 - 2.3 Issue: BioBrick formats
 - 2.4 Device definition
 - 2.5 Biobrick & Device families
- 3 What is the data model needed to describe a biobrick?
 - 3.1 minimal Biobrick information
 - 3.1.1 some extended fields [useful but less obvious]
 - 3.2 Biobrick classification
 - 3.2.1 suggestion: Biobrick class / family

Data Exchange Architecture

web of registries



Data Exchange Architecture



Web Images Maps News Shopping Gmail more ▼

raik.gruenberg@gmail.com | Classic Home | My Account | Sign out

Google Search | I'm Feeling Lucky

Advanced Search
Search Preferences
Language Tools

Home Add a tab

To-Do List

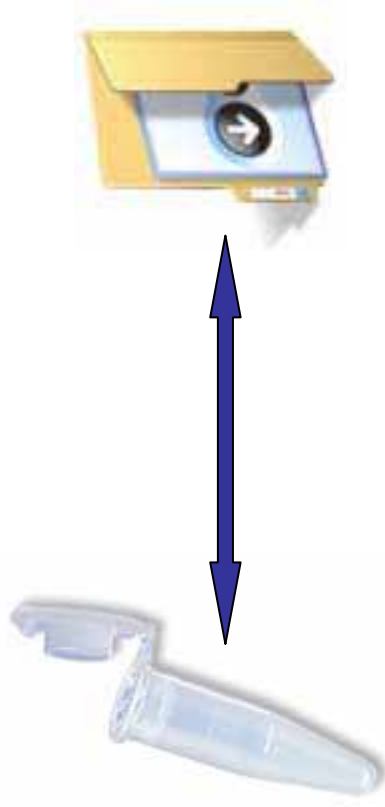
Live Weather

Gmail

New! Select theme | Add stuff »

Biobrick Definition

- A BioBrick is a standardized, continuous DNA encoding a basic biological function.
- A BioBrick has a unique DNA sequence.
- Basic Biobricks are defined by this DNA sequence.
- Composite Biobricks are defined as "sequence" of Basic BioBricks, along with intervening "scar" sequences.
- A BioBrick has a defined & standardized Format



PoBoL

Provisional Biobrick Ontology Language

minimal

extendable

RDF / OWL / XML

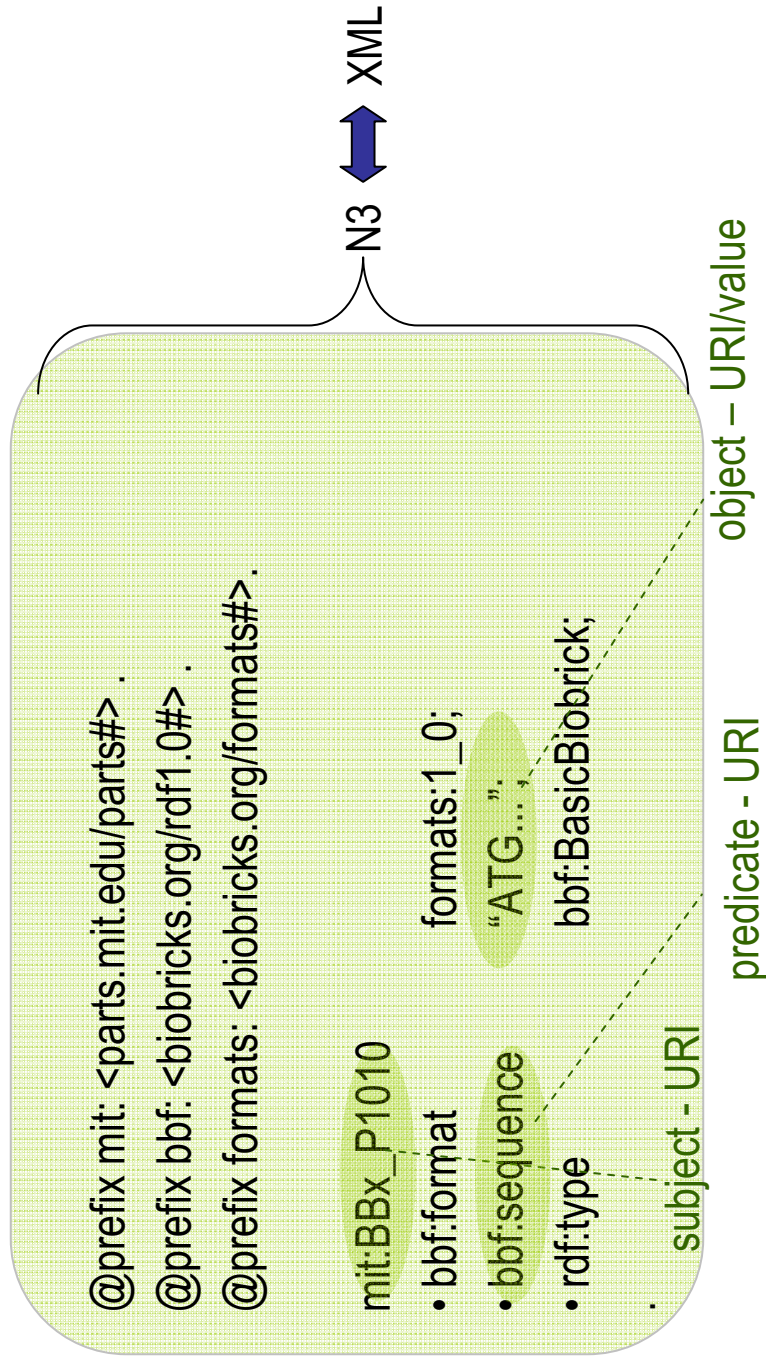
- unique ID
- DNA sequence
- optional: sequence of basic building blocks + scars
- format [specifying: prefix, suffix, self_scar, name, description]
- short_description for humans
- long_description for humans
- author(s)
- reference(s) (web / literature) [pubmed ID?, isbn?, web-address?, doi? + comment?]

PoBOL – additional concepts

- BiobrickFamily
- BiobrickDevice
- BiobrickFormat
- Vector (Biobrick)
- CompositeBiobrick (Biobrick)
- BasicBiobrick (Biobrick)



PoBOL format: OWL / RDF / XML



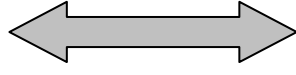
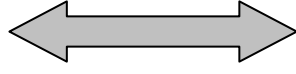
BrickIt

open source Biobrick management

Registry of Standard Biological Parts



<http://parts.mit.edu>

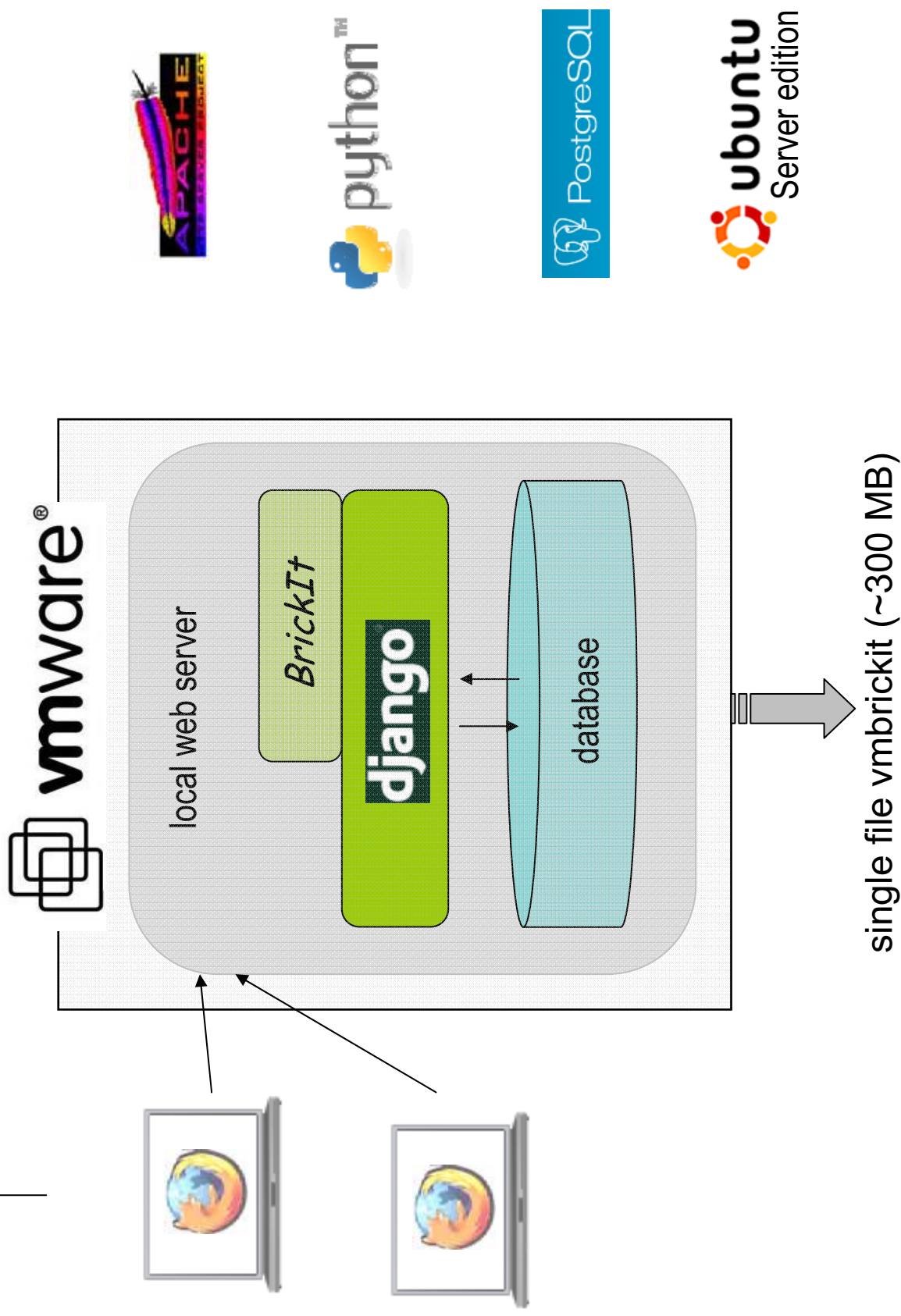


Planning —→ Construction —→ Verification



- no tracking of your local samples
- no local modifications
- all openness is nice but...
- closed development

BrickIt architecture



single file vmbrickit (~300 MB)

Brickit Registry

Site administration

| | |
|--------------------|---------------|
| Auth | |
| Groups | + Add Change |
| Users | + Add Change |
| Repository | |
| Biobricks | + Add Change |
| Brick chains | + Add Change |
| Brick formats | + Add Change |
| BrickCategories | + Add Change |
| BrickTypes | + Add Change |
| Physical DNAs | + Add Change |
| Samples | + Add Change |
| Selective markers | + Add Change |
| Storage containers | + Add Change |
| Vectors | + Add Change |

Recent Actions

My Actions

- rg0101 Biobrick
- rg0101 Biobrick
- rg0100 Biobrick
- rg0100 Biobrick
- 2.0_biofusion Brick-format
- rg0100 Biobrick
- rg0100 Biobrick
- rg0100 Biobrick
- rg0100 Biobrick
- rg0100 Biobrick

Select biobrick to change

[Add biobrick](#) +

Go

| Name | Implementation status | Experience | BioBrick format | BioBrick type | Short description |
|--------|-----------------------|------------|-----------------|---------------|---|
| rg0100 | available | works | 1.0 | B:basic part | cell death gene... construction tool |
| rg0101 | available | works | (None) | B:basic part | test |

2 bioBricks

Filter

By bioBrick format

- All
- 1.1_coding
- 1.0
- 2.0_biofusion

By implementation status

- All
- available
- planning
- ordered
- under construction
- submitted to MIT

By experience

- All
- works
- doesn't work
- none

By bioBrick type

- All
- G:primer
- M:tag
- C:protein coding
- S:intermediate
- V:cell strain
- P:protein generator
- N:general_non-coding
- B:basic part
- R:rna_coding

By brick category

- All
- measurement
- reporter
- rbis
- promoter
- rna
- rna_binding
- terminator

BioBrick rg0100 / BBa_P1010

BBa_P1010 History Edit

Description

cell death gene, construction tool

The CcdB protein, constitutively expressed by P1010, is lethal to most of the BioBrick cell strains, only DB3.1 is resistant.

Use

P1010 is used when putting BioBrick parts into BioBrick plasmids. The part to be inserted and the plasmid are cut with BioBrick enzymes and mixed. The mixture will include both the original uncut or religated plasmid and the desired structure. However, because of CcdB, all of the cells containing the original plasmid die and the surviving colonies are the desired result.

Background

Most BioBrick plasmids are delivered with the P1010 insert see P1010 Physical DNA for the current list of plasmids that are available in this form. This particular BioBrick with its Plasmid backbone is considered a Construction Plasmid.

Comments

this entry is for testing purposes only

Sources & References

Source:

Source gene:

References

Sequence

Sequence without Prefix and Suffix

```
ACTGGCTGTAT AAGGAGCCTGACATTTATATCCCGAAGACATCAGGTTAATGGCGTTTTTGATGTCAATTTTCGGG
TGCTGACATCAGCCACTTTCCCGGATAAGGGACGGCACACTGGCCATATCGTGGTCATCGCCAGCTTTC
ATCCCGATATGCACCACCGGTAAGTTACGGGAGACTTTATCTGACAGCAGCTGCACCTGCCAGGGGATCACCA
TCGGTCGCCGGCGGTGCAAT AATATCACTGTACATCCAAACAGAGATAACGGCTCTCTCTTTATAGGTGTAA
ACCTTAAACTGCATTTACCAAGCCCTGTTCTGTGAGAAAGAGCCGTTCAATTC AATAACGGGGCAGCTCAGCCA
TCCCTTCCGTGATTTCCGCTTTCCAGCGTTCCGACCGAGACGGGTTCAATTCGCATGGTTGTGCTTACCGAGCGG
GAGATATTGACATCATATGCGCTTGAGCAACTGATAGCTGTGGCTGCAACTGCACTGTAATACGCTGCTTCAATAGCA
TACCTCTTTTGGACATCTTCGGGATACATATCAGTATATATTTCTATACCGCAAAAATCAGCGCGCAAAATACGCATAC
TGTTATCTGGCTTTAGTAAGCGGATCCAGCGGT
```

Availability

This BioBrick can be found in the following Vectors and Samples:

| Physical DNA | in Vector | in Container | Sample |
|--------------|-----------|--------------------|------------------------|
| rg0100_pBS | pBS | 001: TestContainer | D_RG01 / rg01/01/08-1a |
| rg0100_pBS | pBS | 001: TestContainer | D_RG01 / rg01/01/08-1b |

Specs

Format:

1.0

Status:

available

Experience:

works

Type:

B: basic part

Users:

admin

Actions



Construct

Verify

Change biobrick

History

Name:

rg0100

start with your initials, e.g. rg0001 or rg_0001 or(for MIT bioBricks) BBa_P1010

Mit id:

BBa_P1010

Short description:

cell death gene, construction tool

Implementation status:

available

Experience:

works

Details

BioBrick format:

1.0

BioBrick type:

B:basic part

Categories:

measurement
reporter
rbs
promoter
rna
rna_binding
terminator

Hold down "Control", or "Command" on a Mac, to select more than one.

Detailed description:

The **CcdB** protein, constitutively expressed by P1010, is lethal to most of the BioBrick cell strains, only DB3.1 is resistant.

Use

=====

P1010 is used when putting BioBrick parts into BioBrick plasmids. The part to be inserted and the plasmid are cut with BioBrick enzymes and mixed. The mixture will include both the original uncut or religated plasmid and the desired structure. However, because of CcdB, all of the cells containing the original plasmid die and the surviving colonies are the desired result.

BioBrick rg0100 / BBa_P1010

Description

cell death gene, construction tool

The CcdB protein, constitutively expressed by P1010, is lethal to most of the BioBrick cell strains, only DB3.1 is resistant.

Use

P1010 is used when putting BioBrick parts into BioBrick plasmids. The part to be inserted and the plasmid are cut with BioBrick enzymes and mixed. The mixture will include both the original uncut or religated plasmid and the desired structure. However, because of CcdB, all of the cells containing the original plasmid die and the surviving colonies are the desired result.

Background

Most BioBrick plasmids are delivered with the P1010 insert see P1010 Physical DNA for the current list of plasmids that are available in this form. This particular BioBrick with its Plasmid backbone is considered a Construction Plasmid.

Comments

this entry is for testing purposes only

Sources & References

Source:

Source gene:

References

Sequence

Sequence without Prefix and Suffix

```
ACTGGCTGTAT AAGGGAGCCTGACATTTATATTTCCCAAGACATCAGGTTAATGGCGTTTTTGATGTCATTTTCGGGG
TGCGTGAGATCAGCCACTTTCCCGGATAAGGGAGACCTGGCCATATCGGTGGTCATATGGCCAGCTTTC
ATCCCGGATATGCACCACCGGGTAAAGTTCAGGGAGACTTATCTGACAGCAGACGTGGCACTGGCCAGGGGATCACCA
TCCGTCCCGGGCGGTGTCAATAATATCACTCTGTACATCCACAAGACAGATAACGGCTCTCTTTTTATAGGTGTAA
ACCTTAAACTGCATTTCAACAGCCCTGTTCTCGTCAGAAAAGCGCTTCAATCAATAAACCCGGGACCTCAGCCA
TCCCTTCTGATTTCCGGTTCCAGGTTCCGACGACAGACGGGCTTCACTGTCATGGTTGTGCTTACCGAGCCG
GAGATATTGACATATATGCCTTGAGCAACTGATAGCTGTCCGCTGTCAACTGTCACTGTAATACCGTCTCATAGCA
TACCTCTTTTGGACATACCTTCGGGTATACATATCAGTATATATTTCTTATACCGCAAAAATCAGGGCGCAAAATACGCATAC
TGTTATCTGGCTTTAGTAAGCCGGATCCAGCGT
```

Availability

This BioBrick can be found in the following Vectors and Samples:

| Physical DNA | in Vector | in Container | Sample |
|--------------|-----------|--------------------|------------------------|
| rg0100_pBS | pBS | 001: TestContainer | D_RG01 / rg01/01/08-1a |
| rg0100_pBS | pBS | 001: TestContainer | D_RG01 / rg01/01/08-1b |

Specs

Format:

1.0

Status:

available

Experience:

works

Type:

B:basic part

Users:

admin

Actions

Edit

Construct

Verify

File Edit View History Bookmarks Tools Help

http://parts.mit.edu/registry/index.php/Part:BBa_P1010

Create an account or log in

DNA Available Experience Works Entered: 2004-07-28

article discussion edit history

Part:BBa_P1010

Designed by Leon Chan

cell death gene, construction tool

The CcdB protein, constitutively expressed by P1010, is lethal to most of the BioBrick cell strains, only DB3.1 is resistant.

P1010 is used when putting BioBrick parts into BioBrick plasmids. The part to be inserted and the plasmid are cut with BioBrick enzymes and mixed. The mixture will include both the original uncut or religated plasmid and the desired structure. However, because of CcdB, all of the cells containing the original plasmid die and the surviving colonies are the desired result.

BioBrick Construction Plasmids Containing This Part

[edit]

Most BioBrick plasmids are delivered with the P1010 insert see P1010 Physical DNA for the current list of plasmids that are available in this form. This particular BioBrick with its Plasmid backbone is considered a Construction Plasmid.

Usage and Biology

[edit]

For more information on how to use this brick, visit its Featured Parts:Cell Death page

Jump to part: BBa_

BBa_P1010

- Part Main Page
- Part Design
- Experience
- Hard Information
- Physical DNA

navigation

- Main Page
- Browse Part Types
- iGEM 2007 Wiki
- Community portal
- Recent changes
- Recent part changes

resources

- User Accounts
- Add a Part
- Part Searches
- DNA Repositories
- Sequence Analysis
- Assembly Tool
- Help

search

Go Search

toolbox

- What links here
- Related changes
- Upload file
- Special pages
- Printable version

Powered By

BioBrick rg0100 / BBa_P1010

BBa_P1010 History Edit

Description

cell death gene, construction tool

The CcdB protein, constitutively expressed by P1010, is lethal to most of the BioBrick cell strains, only DB3.1 is resistant.

Use

P1010 is used when putting BioBrick parts into BioBrick plasmids. The part to be inserted and the plasmid are cut with BioBrick enzymes and mixed. The mixture will include both the original uncut or religated plasmid and the desired structure. However, because of CcdB, all of the cells containing the original plasmid die and the surviving colonies are the desired result.

Background

Most BioBrick plasmids are delivered with the P1010 insert see P1010 Physical DNA for the current list of plasmids that are available in this form. This particular BioBrick with its Plasmid backbone is considered a Construction Plasmid.

Comments

this entry is for testing purposes only

Sources & References

Source:

Source gene:

References

Sequence

Sequence without Prefix and Suffix

```
ACTGGCTGTAT AAGGAGCCTGACATTTATATCCCGAAGACATCAGGT AATGGCGTTTTTGATGTCAATTTTCGGG
TGCTGACATCAGCCACTTTCCCGGATAAGGGACCGGCACACTGGCCATATCGTGGTCATCGCCAGCTTTC
ATCCCGATATGCACCACCGGTAAAGTTCACGGAGACTTATCTGACAGCAGGTGCACCTGCCAGGGGATCACCA
TCGGTCCCGGGCGTGAAT AATATCACTGTACATCCAAACAGAGATAACGGCTCTCTTTTATAGGTGTAA
ACCTTAAACTGCATTCACAGCCCTGTTCTGTGAGAAAGACCGTT CATTTC AATAACGGGGCAGCTCAGCCA
TCCCTTCCGTGATTTCCGCTTTCAGCGTTCGGCACGACGAGCGGCTT CATTTC GCATGGTTTGCTTACCGAGCG
GAGATTTGACATCATATGCGCTTGAGCAACTGATAGCTGTGGTGTCAACTGCACTGTAATACGCTGCTTCA TAGCA
TACCTCTTTTGGACATCTTCGGGATACATATCAGTATATATTTCTATACCGCAAA AATCAGCGGCAAAATACGCATAC
TGTTATCTGGCTTTAGTAAGCGGATCCAGCGT
```

Availability

This BioBrick can be found in the following Vectors and Samples:

| Physical DNA | in Vector | in Container | Sample |
|--------------|-----------|--------------------|------------------------|
| rg0100_pBS | pBS | 001: TestContainer | D_RG01 / rg01/01/08-1a |
| rg0100_pBS | pBS | 001: TestContainer | D_RG01 / rg01/01/08-1b |

Specs

Format:

1.0

Status:

available

Experience:

works

Type:

B: basic part

Users:

admin

Actions



Construct

Verify

Change sample

[History](#) [View on site](#)

| | |
|------------------------------|--|
| Label: | <input type="text" value="rg01/01/08-1a"/> <small>example: rg23/12/07-1a (rg...user initials)</small> |
| Bar code: | <input type="text"/> |
| Container: | 001: TestContainer |
| Well: | <input type="text"/> |
| Type of well or tube: | tube |
| Content | |
| Physical DNA: | rg0100_pBS |
| Stored as: | DNA |
| In cell: | |
| Concentration: | <input type="text" value="100.00"/> |
| Conc. unit: | mg/l |

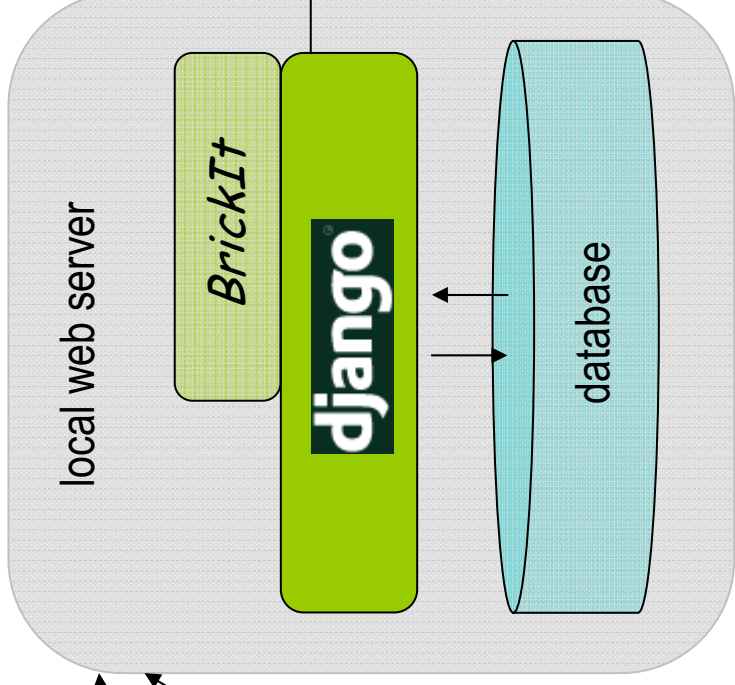
Additional information

Users:

Hold down "Control", or "Command" on a Mac, to select more than one.

Comments:

BrickIt architecture



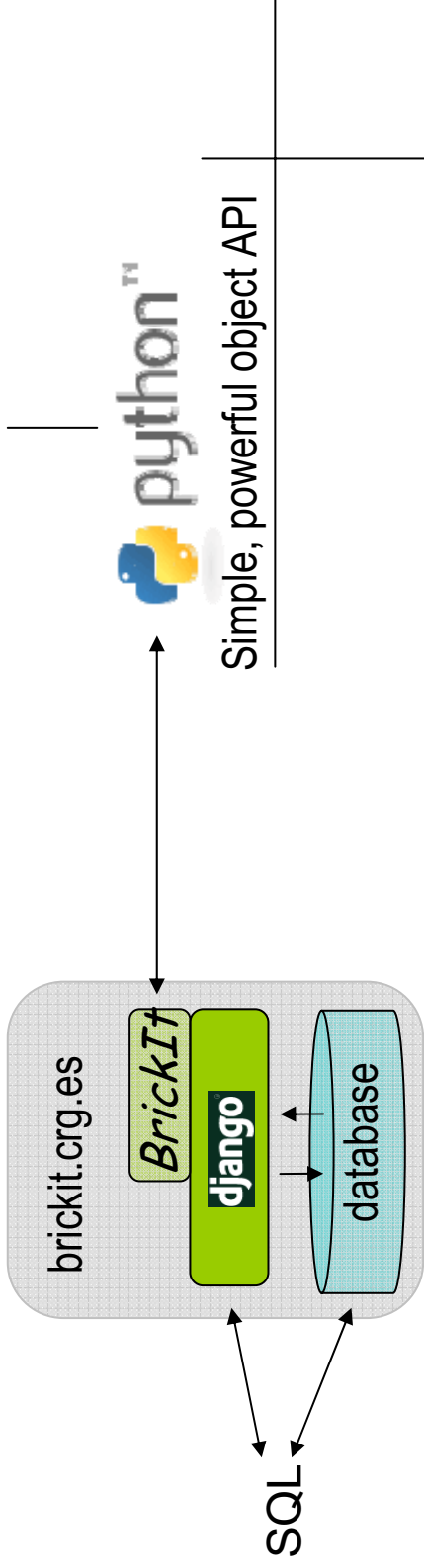
- + rapid development
- + clean yet pragmatic design
- + data models defined in Python
- + well documented & supported

www.djangoproject.com

The Web framework for perfectionists with deadlines.
Django makes it easier to build better Web apps more quickly and with less code.

Flexible

huge reservoir of Python libraries,
Biopython, numpy, SciPy...

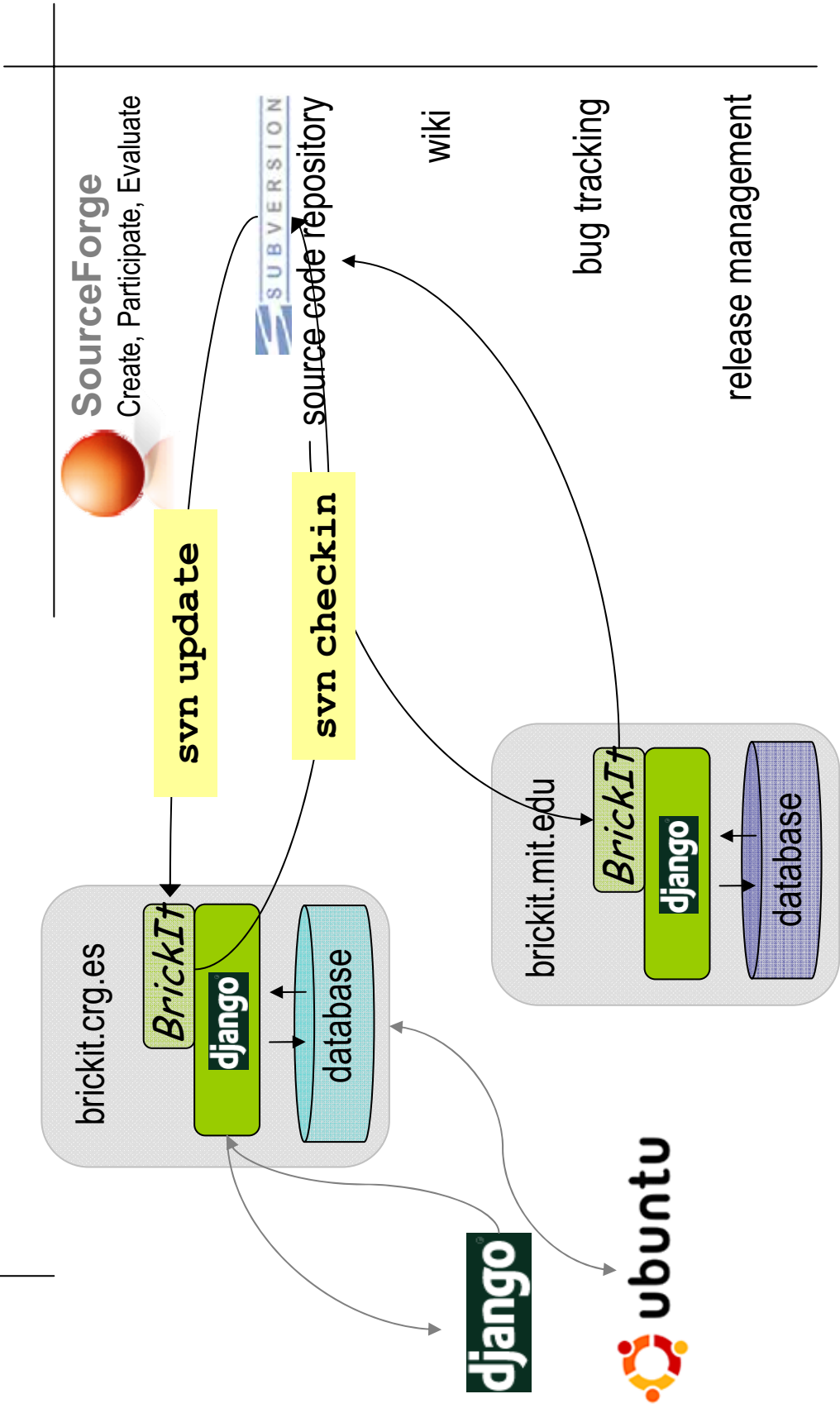


```
from djbrickit.repository.models import Vector, SelectiveMarker
```

```
amp = SelectiveMarker( name='amp', short_description='AmpR' )  
amp.save()
```

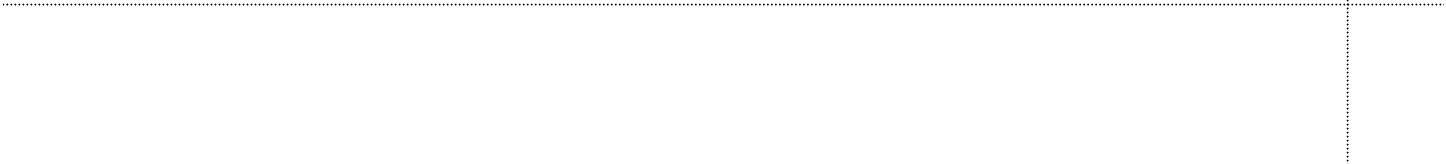
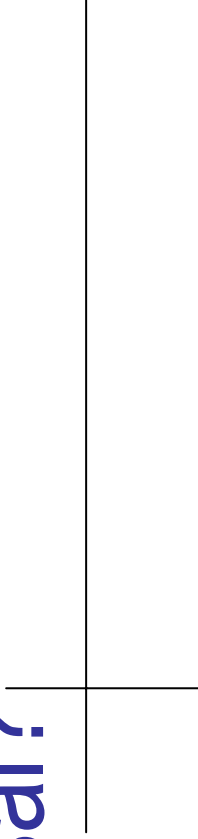
```
pBS = Vector.objects().get(name='pBS')  
pBS.marker = amp  
pBS.save()
```

Shared development



divided data but shared infrastructure & development

E.U. project proposal?



In conclusion, ...

- ...check out the BBF Standardization process!
- ...PoBOL = minimal Biobrick description language
- ...BrickIt = open source development platform for custom part registries
- ... weaving a web of registries
- ... lobby for an E.U. project?

Corner stones

- extension of standards
- open source developer community
- registry platform
- design tools
- aggregation / data mining tools
- ...

Funding for:

- programming sprints
- people
- web servers / infrastructure
- travel for outside participants
- outreach / teaching / iGem

Acknowledgements

BBF Standards
mailing list



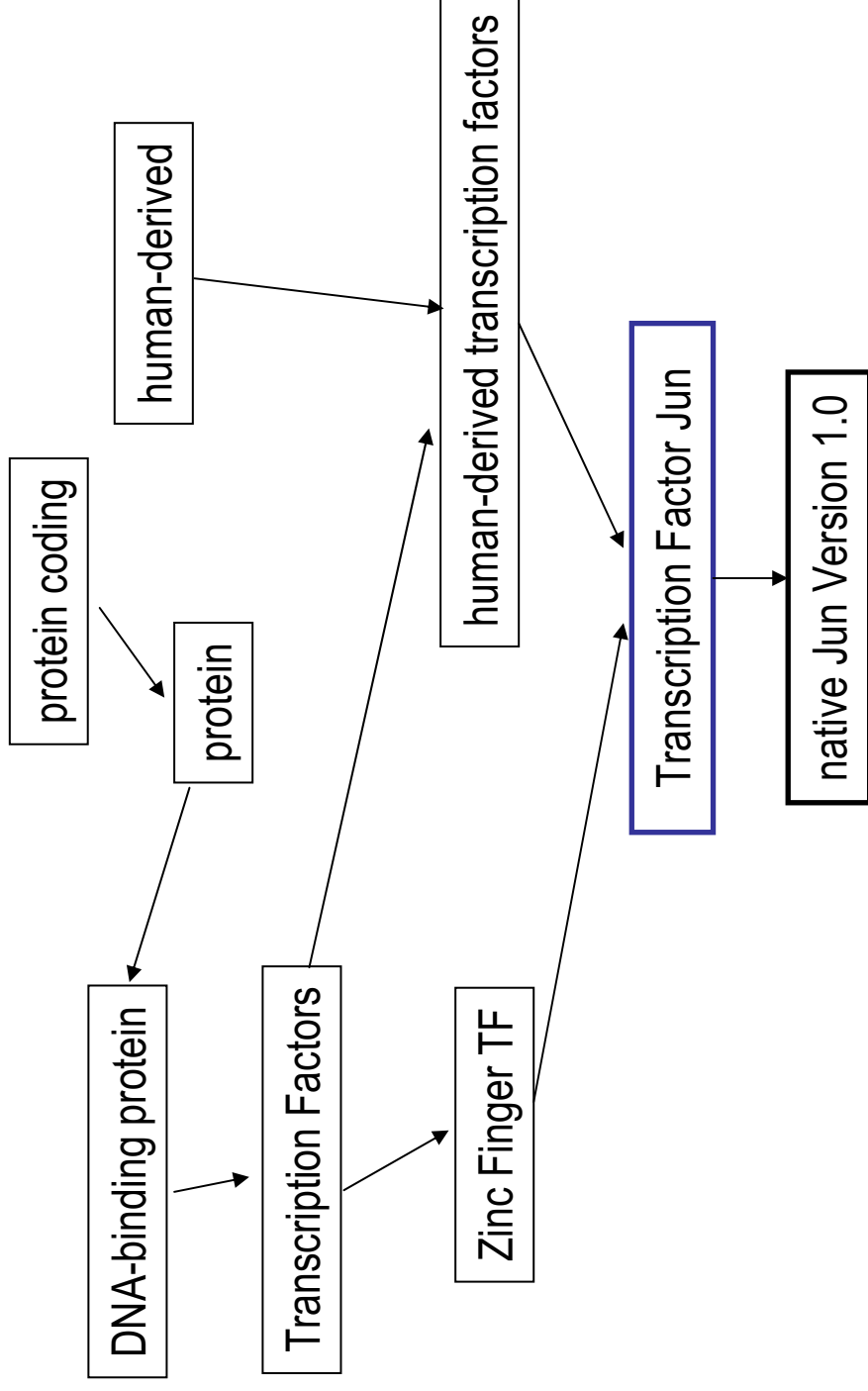
Luis Serrano



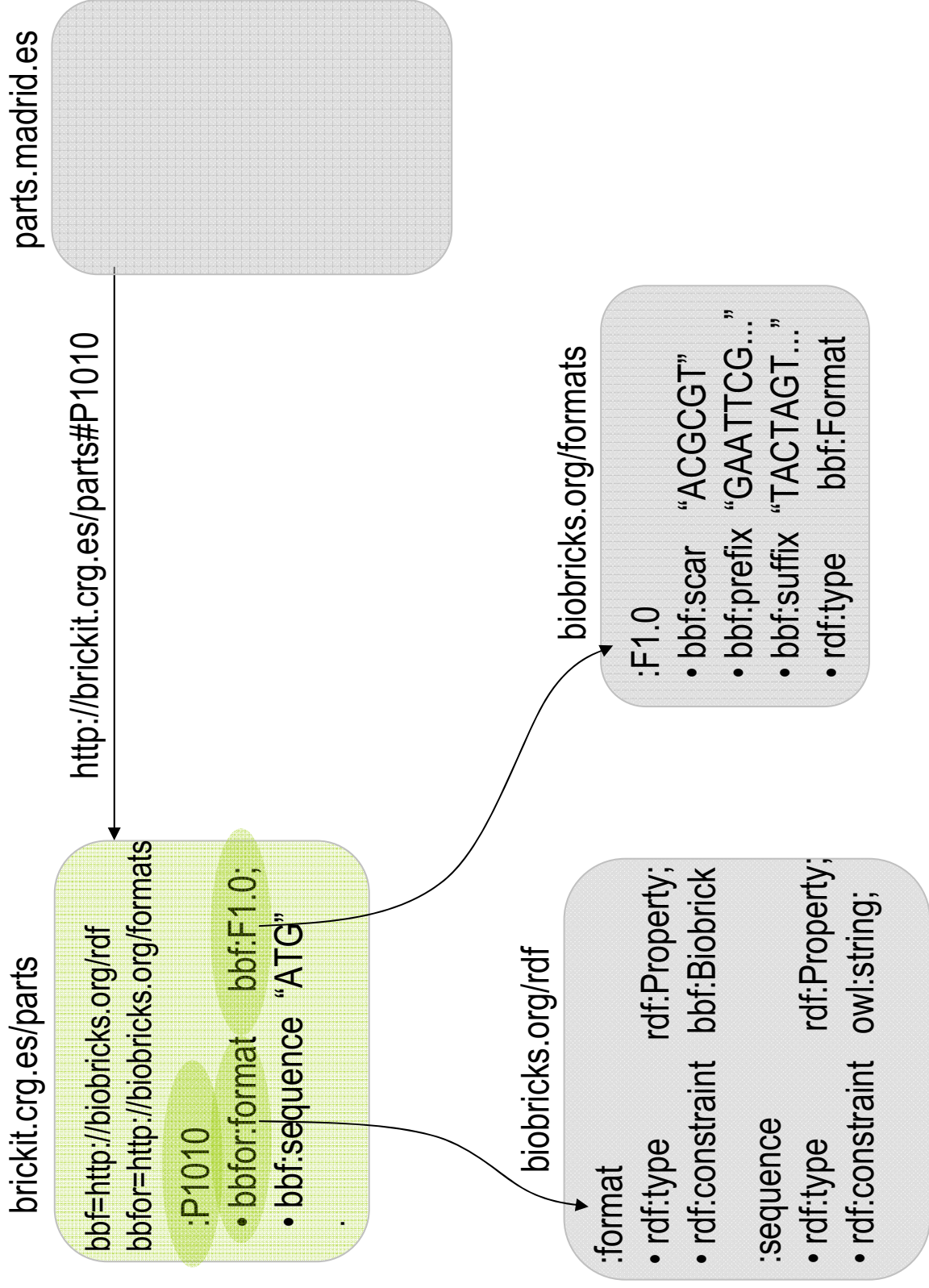
Human Frontier Science Program

Biobrick Classification

- Biobricks are grouped into Families
- Each Family can have many sub- and many parent families



Data Exchange: REST / RDF+OWL



EMERGENCE WP4

Towards a **consensus language** for SB:

Conceptual and **hermeneutical** tools for

Formatting and categorization of
Transcriptional Working States

P2 (CSIC), P4 (CRG), P7 (UCL)

The central question:

How to

- describe
- de-construct
- re-construct

Biological complexity?

Hermeneutics

The development of interpretation tools and understanding of texts and systems of meaning. The concept of "text" is here extended beyond written documents to any number of objects subject to interpretation.

The hermeneutical problem:

- All languages for describing reality are metaphoric
- Understanding complex systems relies on suitable metaphores
- No descriptive language is neutral, they all have an agenda

Metaphor #1: The radio

Building a radio with parts



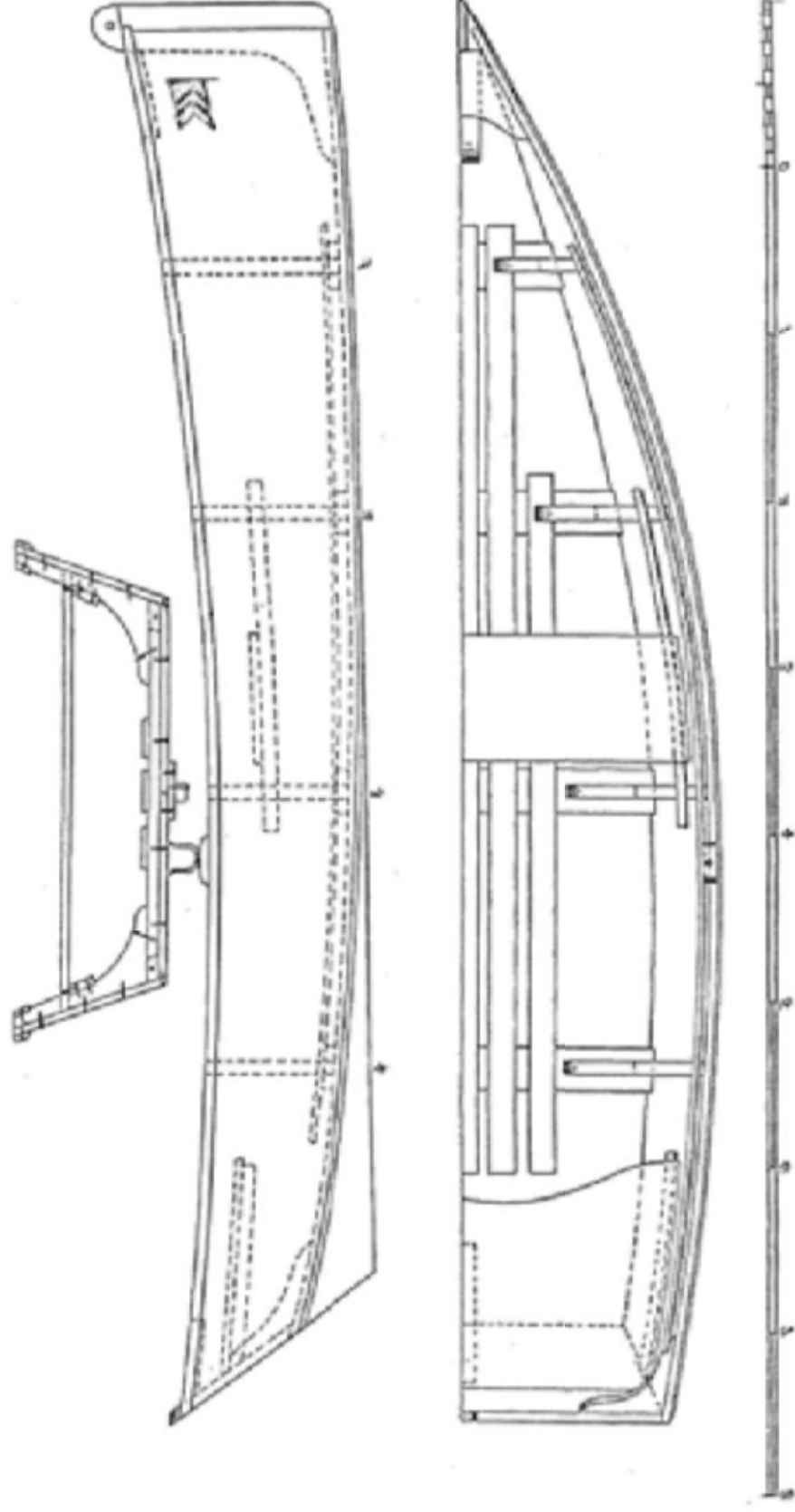
J. Keasling

Metaphor #2: The chassis (iGEM favourite!)



D. Endy

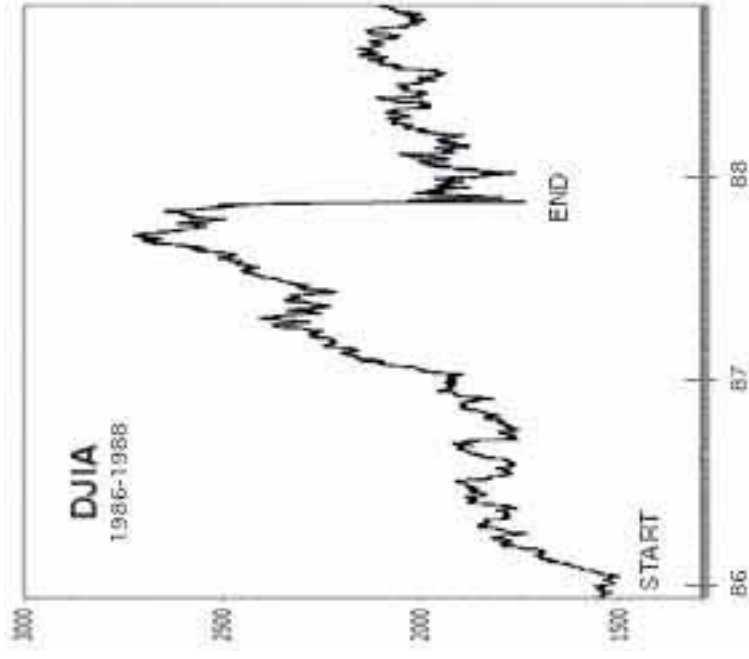
Metaphor #3: The Delphic boat



A. Danchin

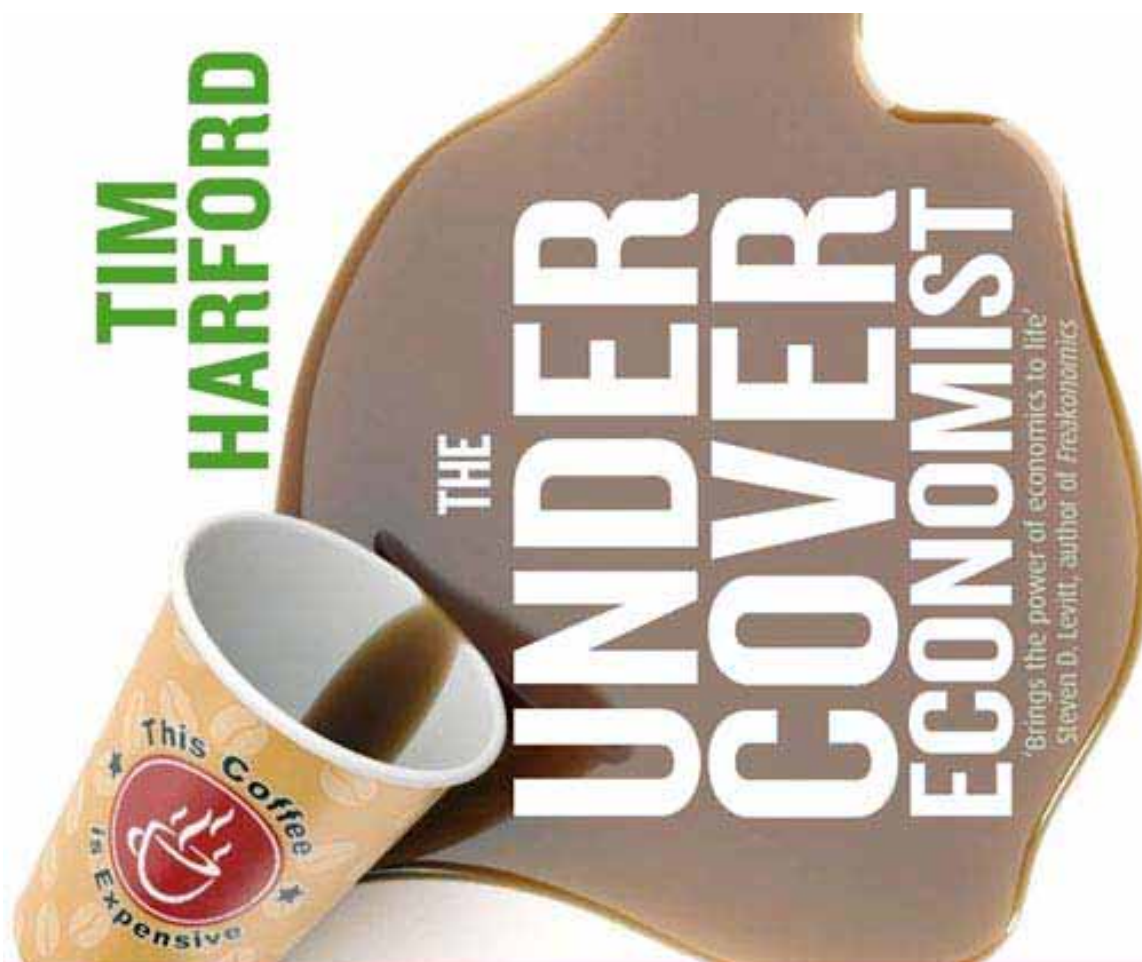
Metaphor #4: The Company

```
ARGENTINA'S PIEDRI
24450 DJ MICROSOFT CORP (MSI
24460 DJ MOTOROLA INC (MOT)
24470 DJ ANHEUSER BUSCH (BU
INDU -27.22 VOLU 289,915.00
INDP 11008.48 UVOL 112,745.80
UTIL -.99 DUOL 134,141.60
TRAN -23.70 TRIN .77
SHRS LSS CR HWP AVE
56 8% 4%
```



JA Ranea

**TIM
HARFORD**

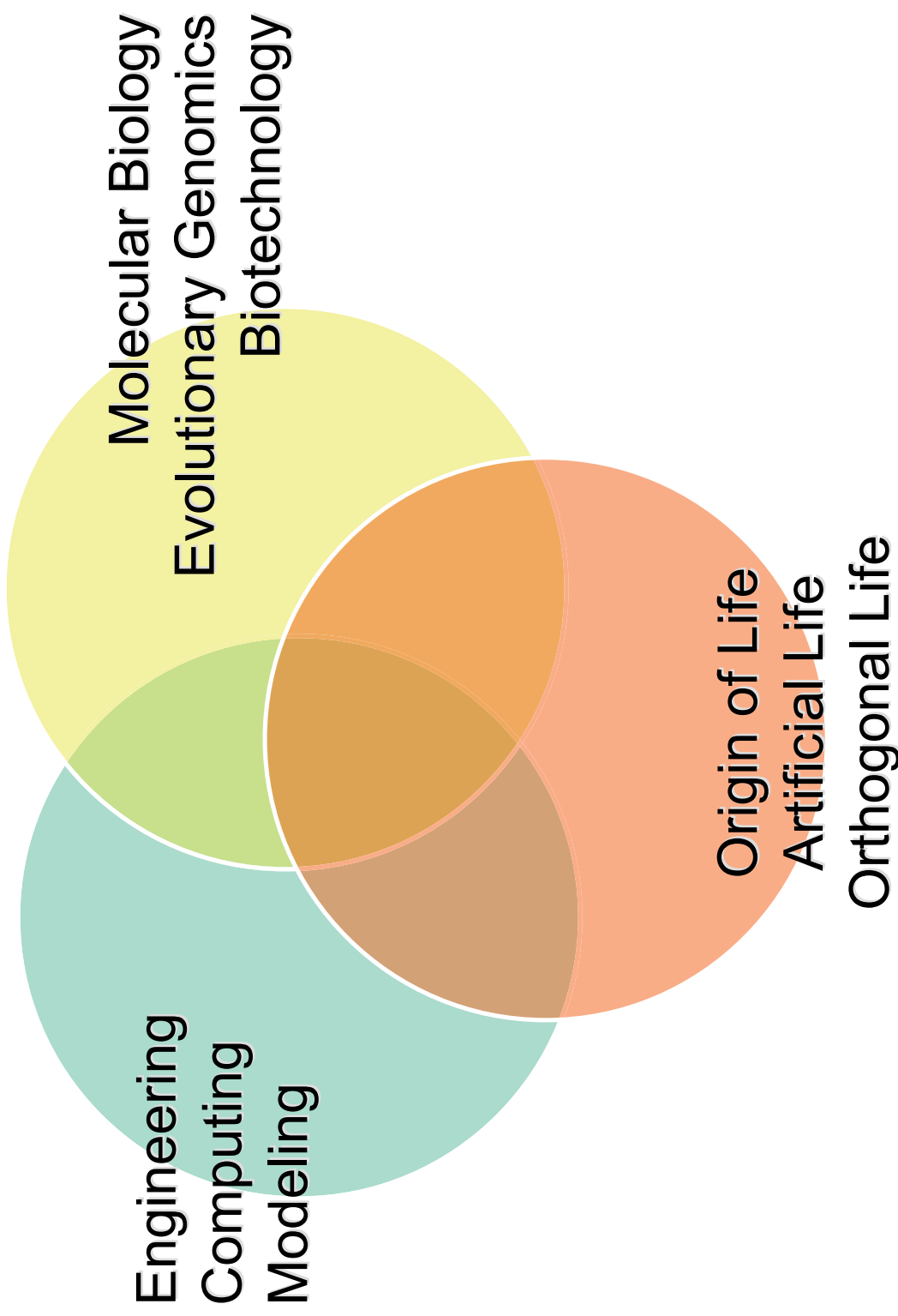


hachette | AUDIO

read by Cameron Stewart

unabridged on 7 cds

The 3 agendas/stakeholders of SB in Europe

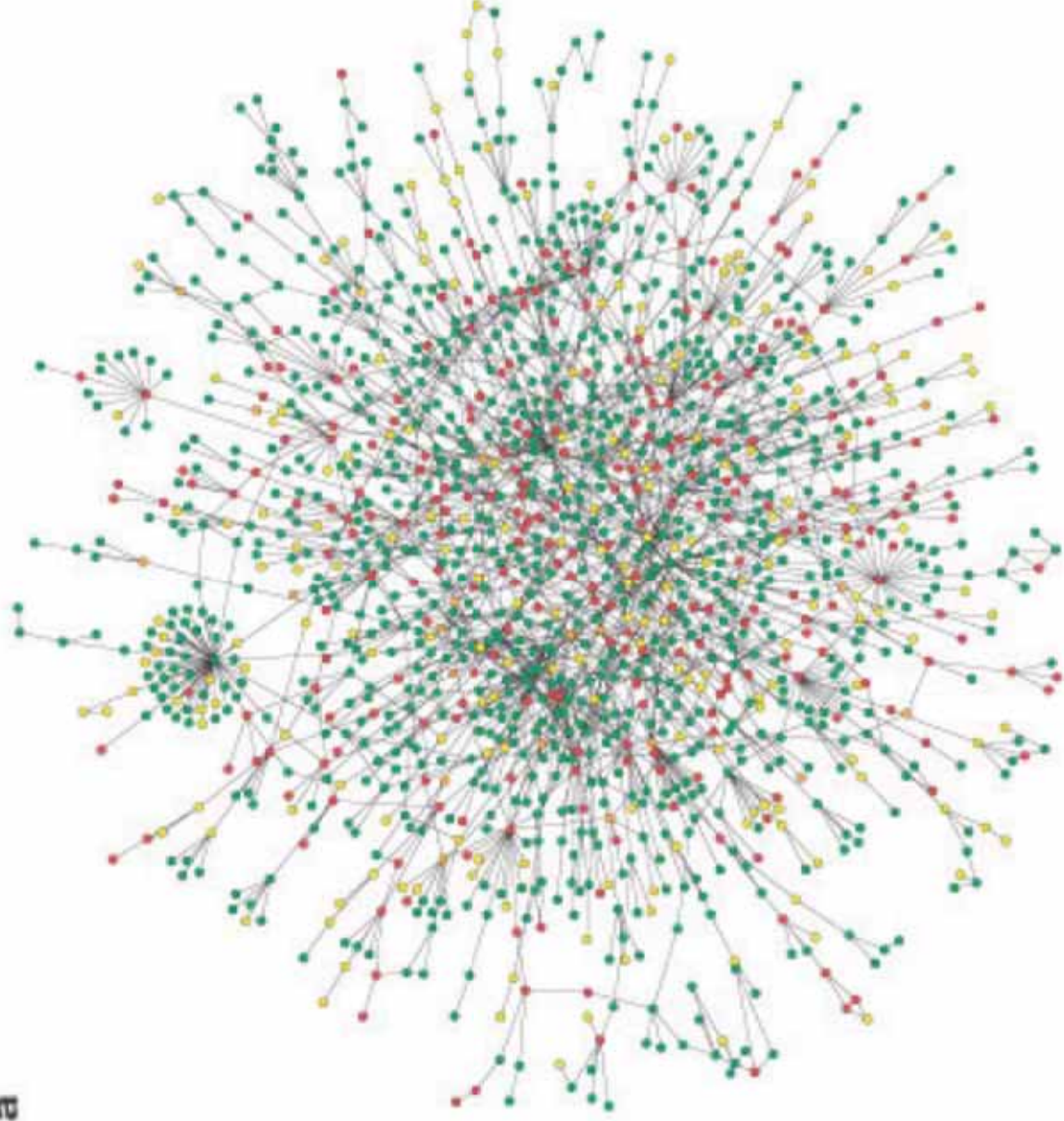


*" ... Engineers hate complexity. I hate emergent properties.
I like simplicity. I don't want the plane I take tomorrow
to have some emergent properties while it is flying..."*

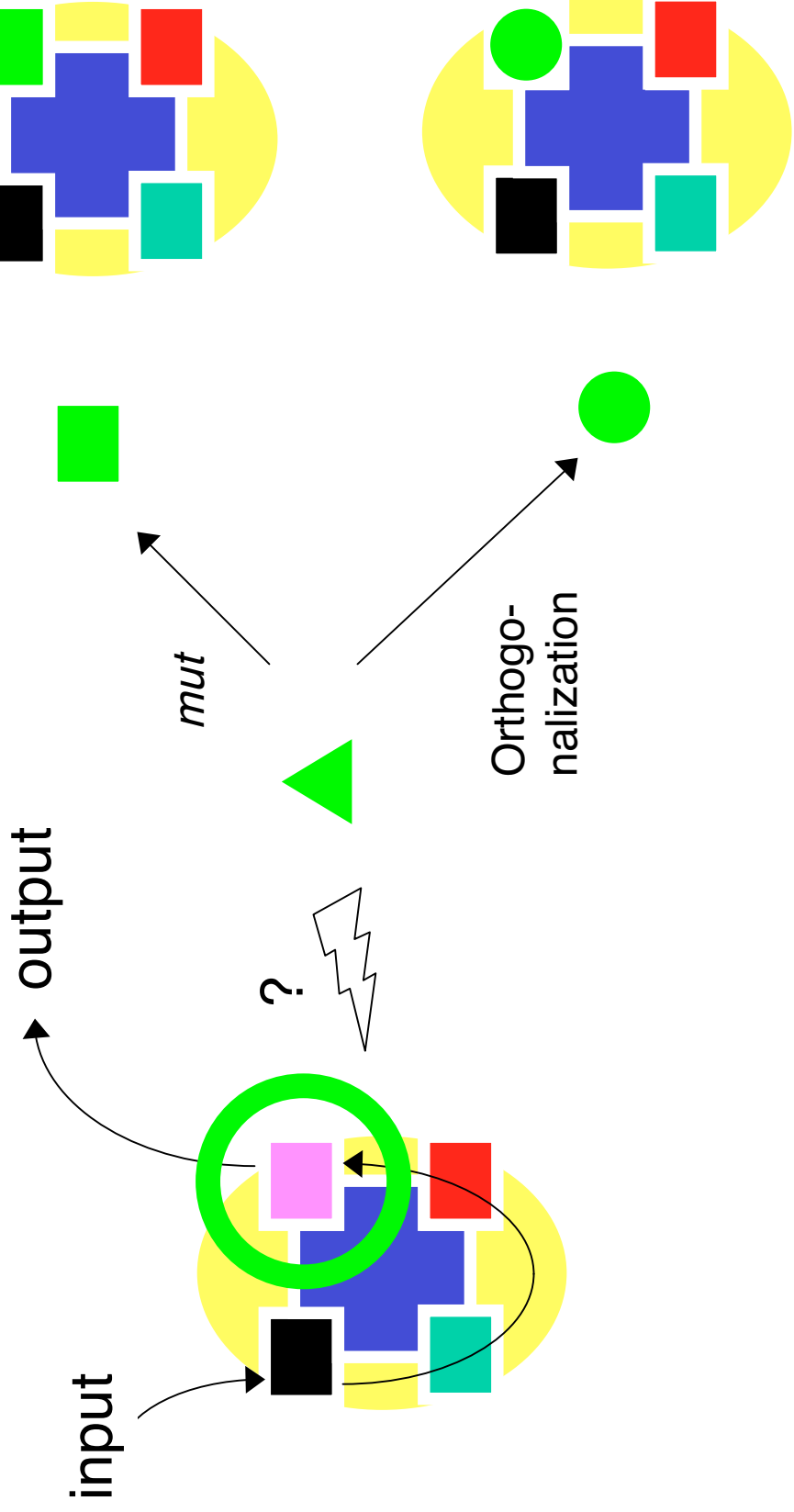
*(www.edge.org; Engineering Biology:
a talk with Drew Endy, 2008)*

Implantation of circuits in a pre-existing network?

a



Resistance to implantation of new functions in a *cellular hyperstructure*



What to do under WP4?

- Early assembly of an expert group on standardization and connectivity of minimal functions.
- Text-mining on quantitative data relevant to promoter functioning (link to WP3) & production of a database
- Modelization of 4 types of prokaryotic promoters as the standard components of choice for building complex regulatory circuits.
- In silico analysis of the data collected for developing a concept aimed at reaching a transatlantic consensus.

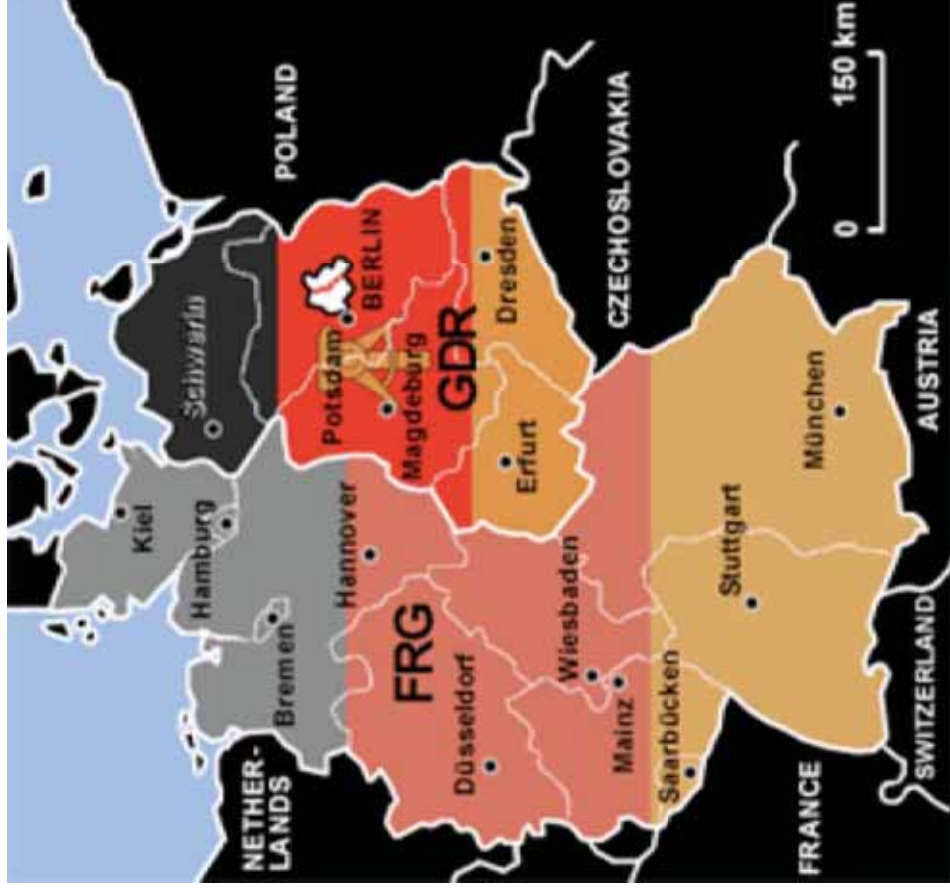
Minimal parts vs. minimal functions

- A better conceptual frame is badly needed here to grasp what

minimal engineer-able biological building blocks

are. Just calling them Biobricks™ and make them equal to singular biological components (as is the case in the MIT-run catalogue of biological parts) can give the perception that the issue is already solved!

Finding the right descriptive terms....



Minimal parts vs. minimal functions

- The cell as an *automaton processing information* in an algorithmic fashion: the program and the machine.
- The quest for a minimal set of functions for a self-maintaining system is not exclusive of Synthetic Biology...

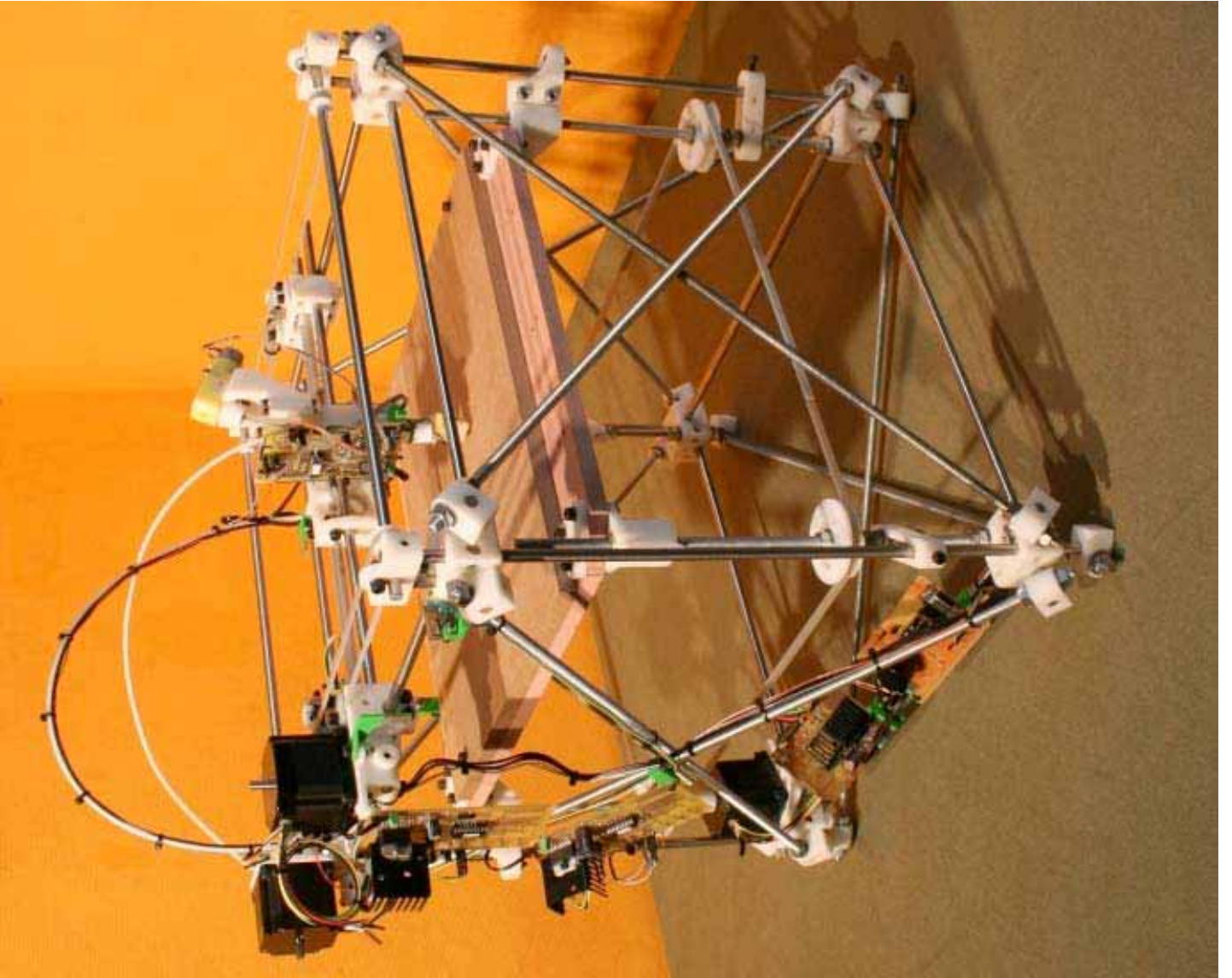
3D Printer: The RepRap Community Project

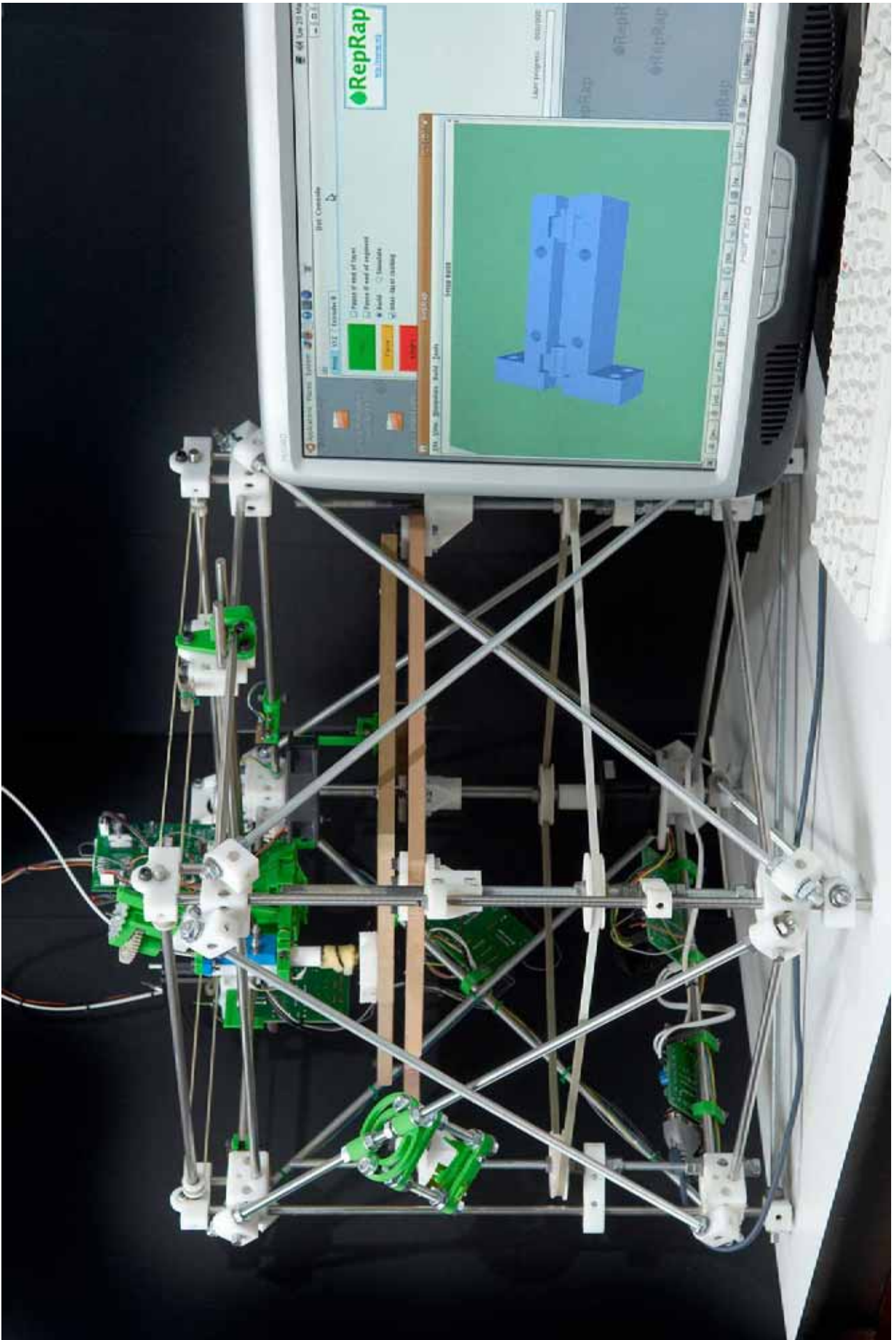
<http://www.reprap.org>

RepRap 1.0 "Darwin" is a rapid prototyping machine that is capable of making the majority of its own component parts.

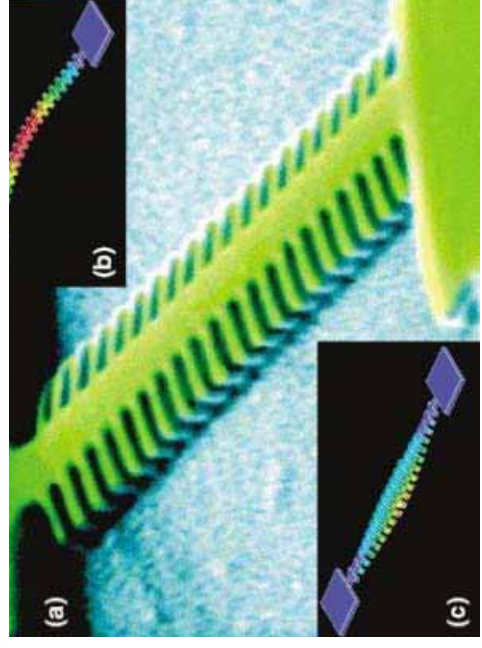
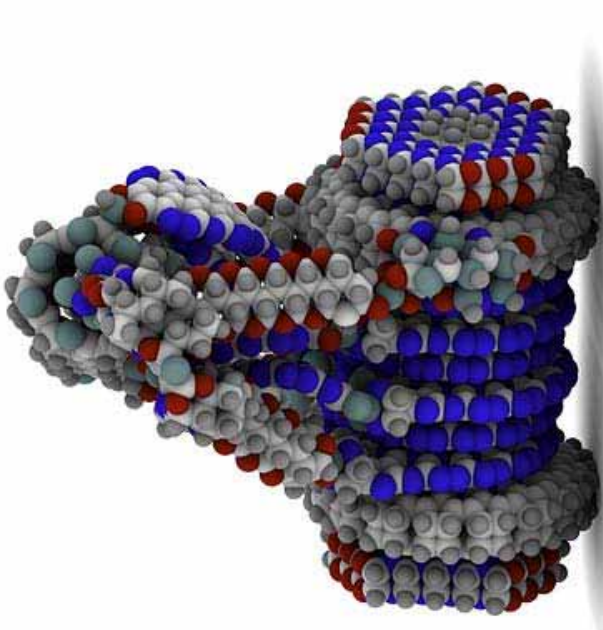
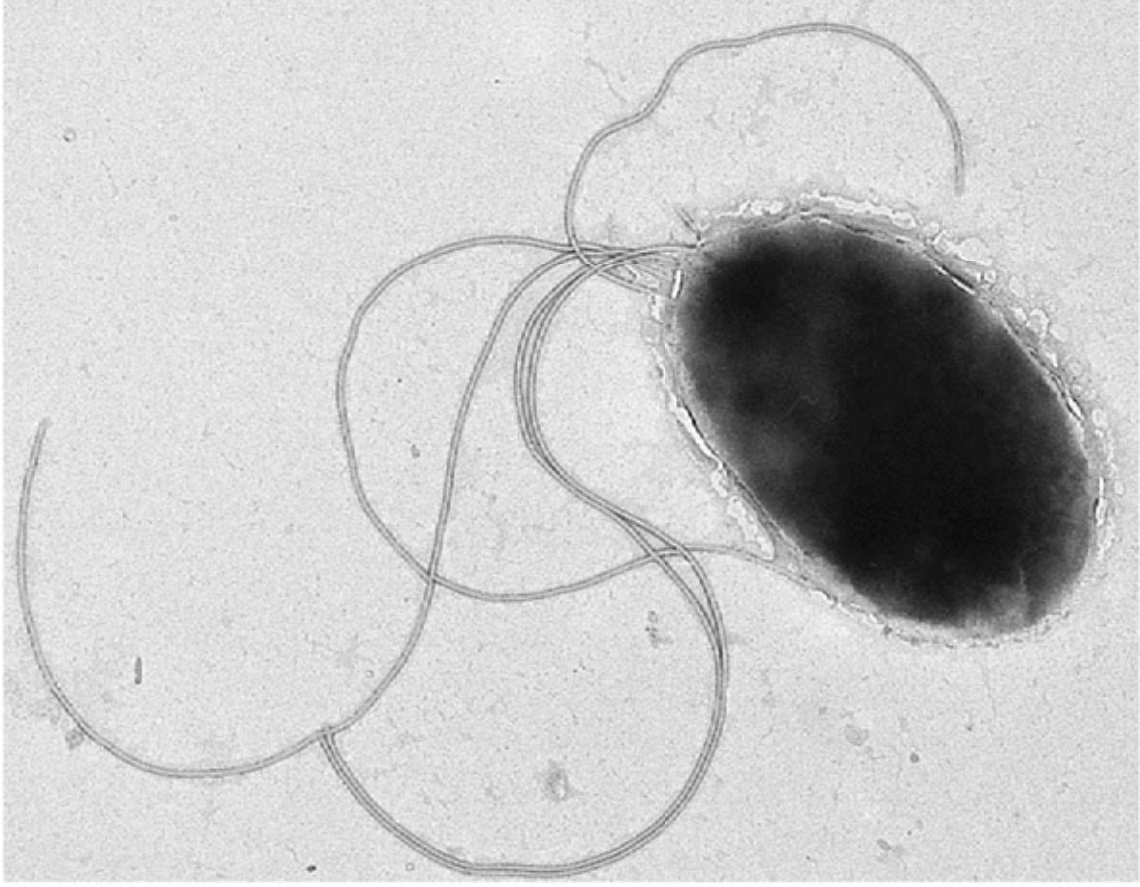
Darwin consists of a frame made from rods and printed parts. A flat build platform moves vertically in that frame, driven on screw threads by a stepper motor. At the top of the frame there are two write heads that move horizontally (driven by toothed belts and two more steppers) extruding a thin stream of molten plastic to form new layers on the build base.

The machine prints layer by layer to form a solid object. The build base then moves one increment down, the second layer is extruded, and so on. There are two heads to allow a filler material to be laid down as well as the plastic. This filler is used to support overhanging parts of the objects being built, and is removed when the process is finished.





The cell as a machine built on nanomachines



*2 remarkable features of the
Minimal self-replicating 3D printer:*

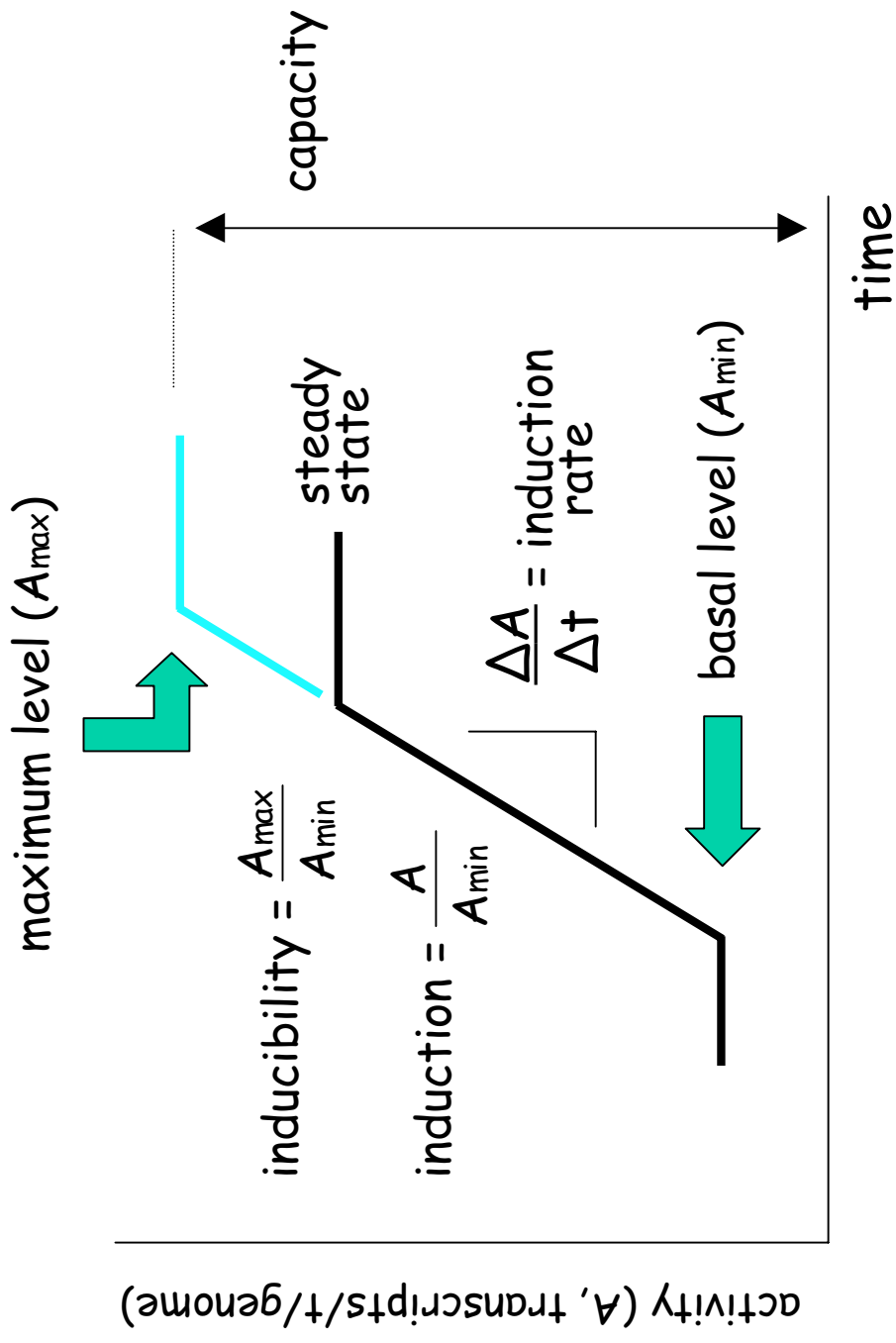
- **Very little regulation needed**
- **Lubricants are essential!**

EMERGENT ISSUE

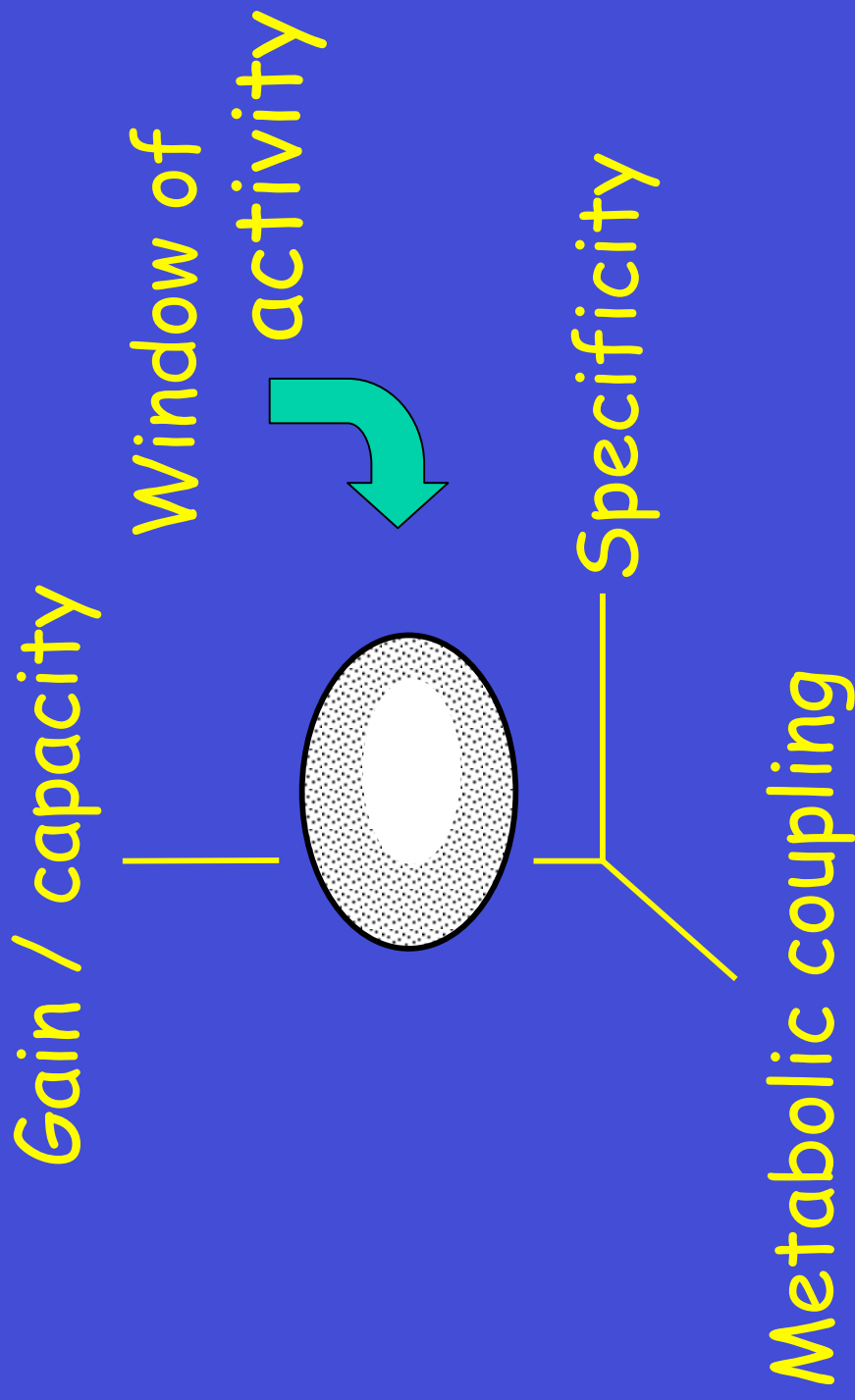
Metabolic transactions impose a permanent chemical and energetic frame to the cells, a sort of background economy.

While the link between translation of mRNA and the ribosome is straightforward, the organization of metabolism is much less so.

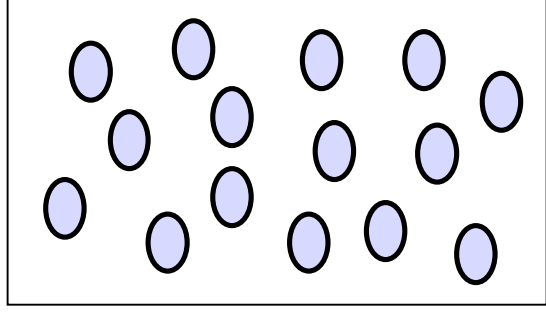
Promoter parameters



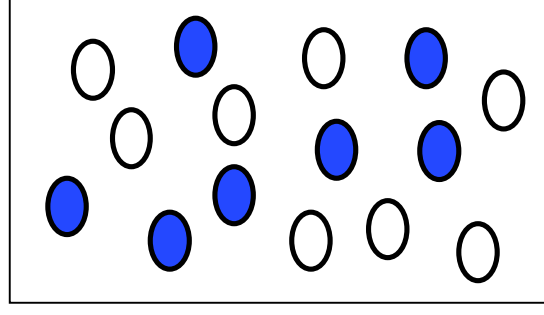
Definition of promoter activities *in vivo*



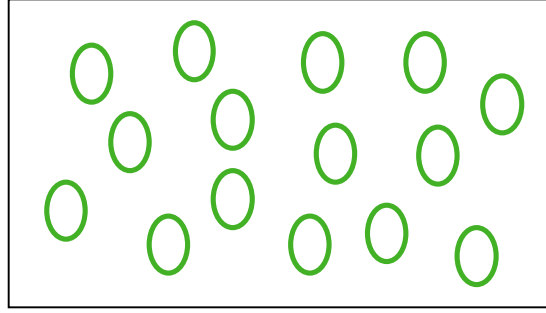
even



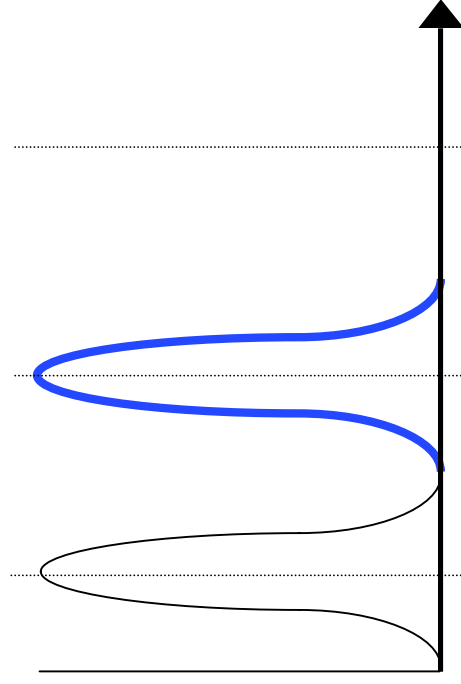
stochastic



uninduced

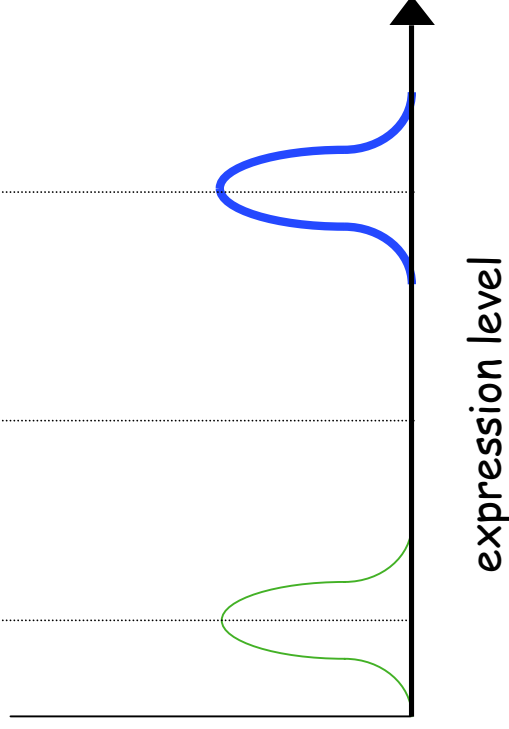


basal medium high



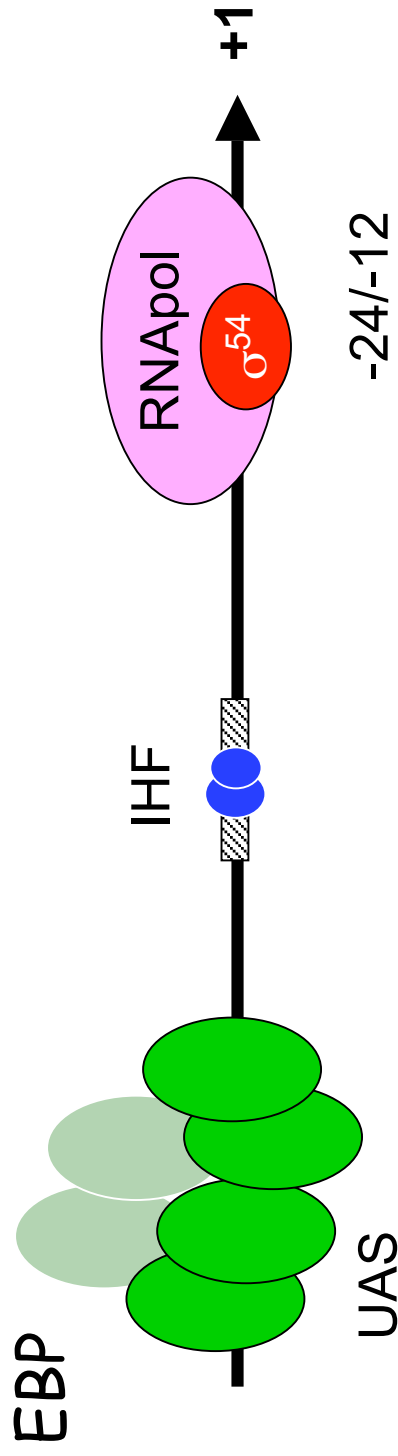
expression level

population

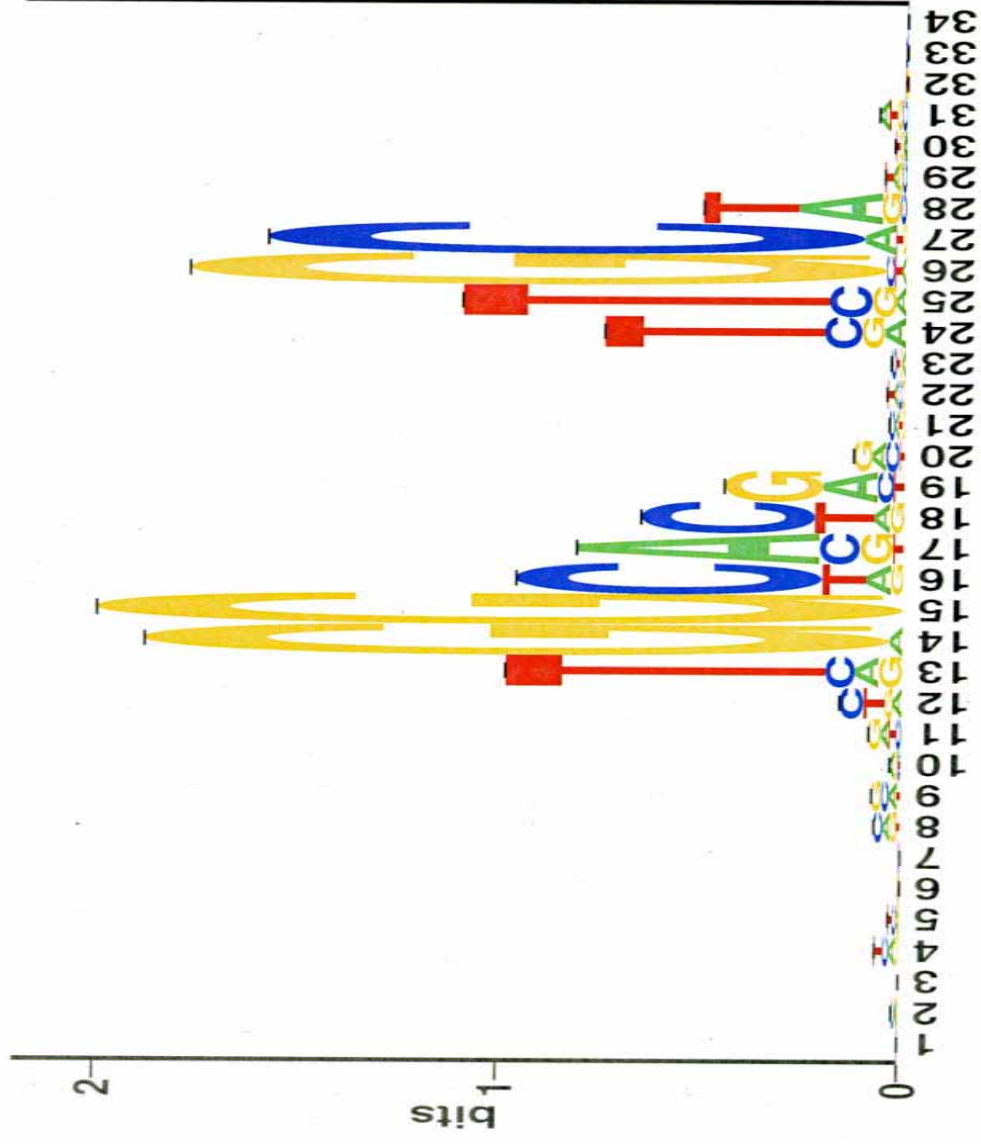


expression level

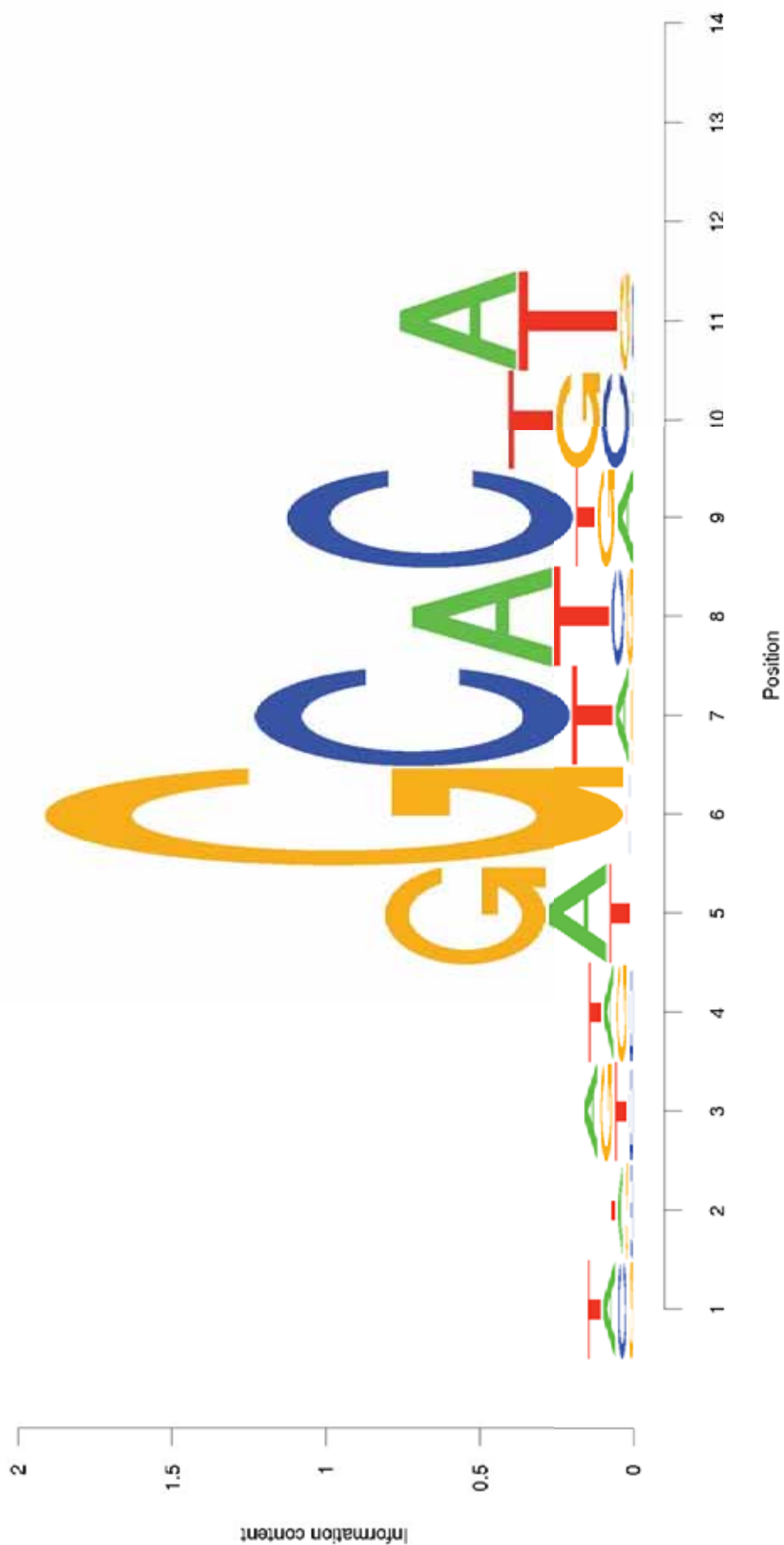
Sigma 54 (RpoN) dependent promoters



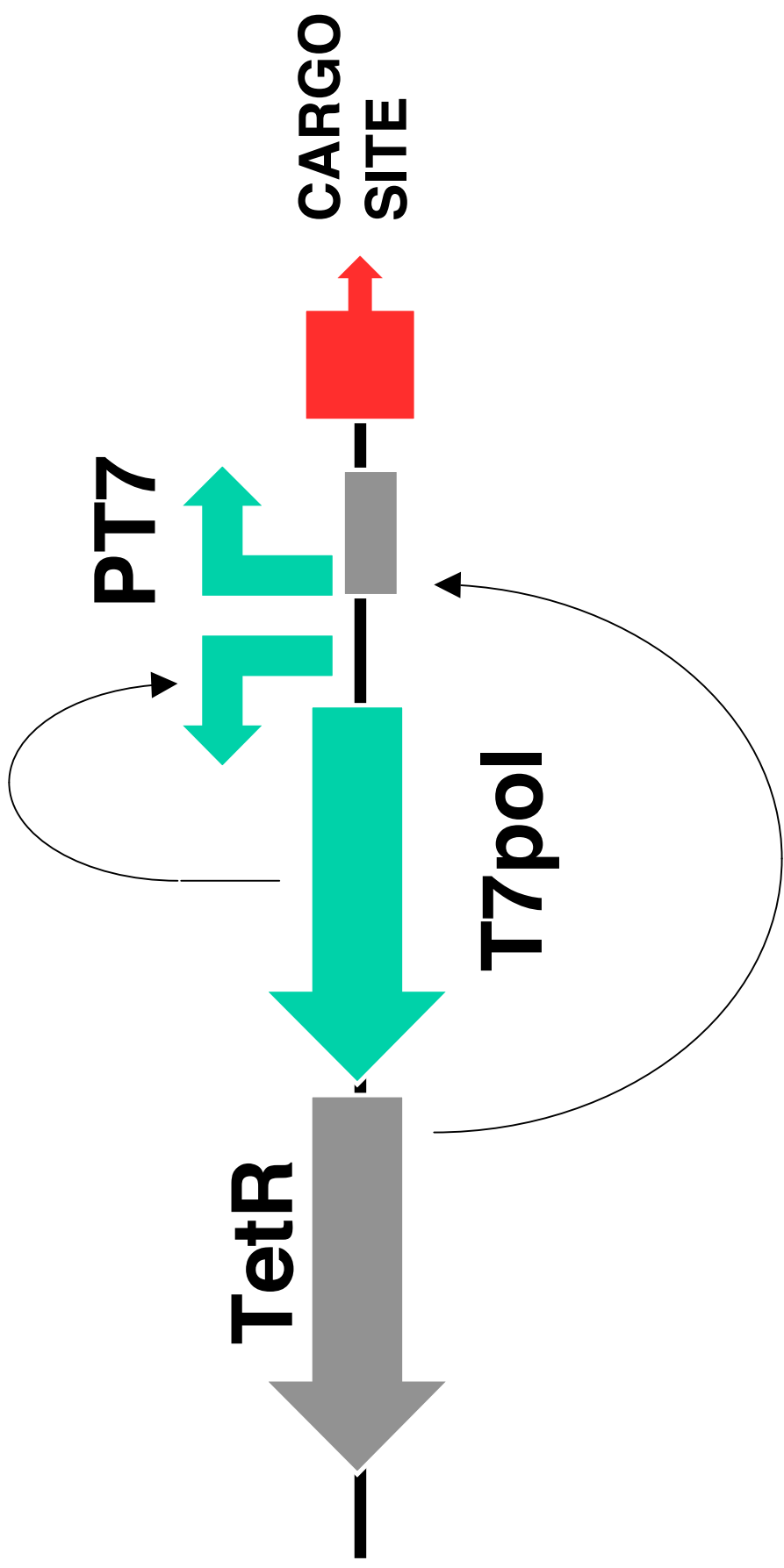
The -12/-24 site



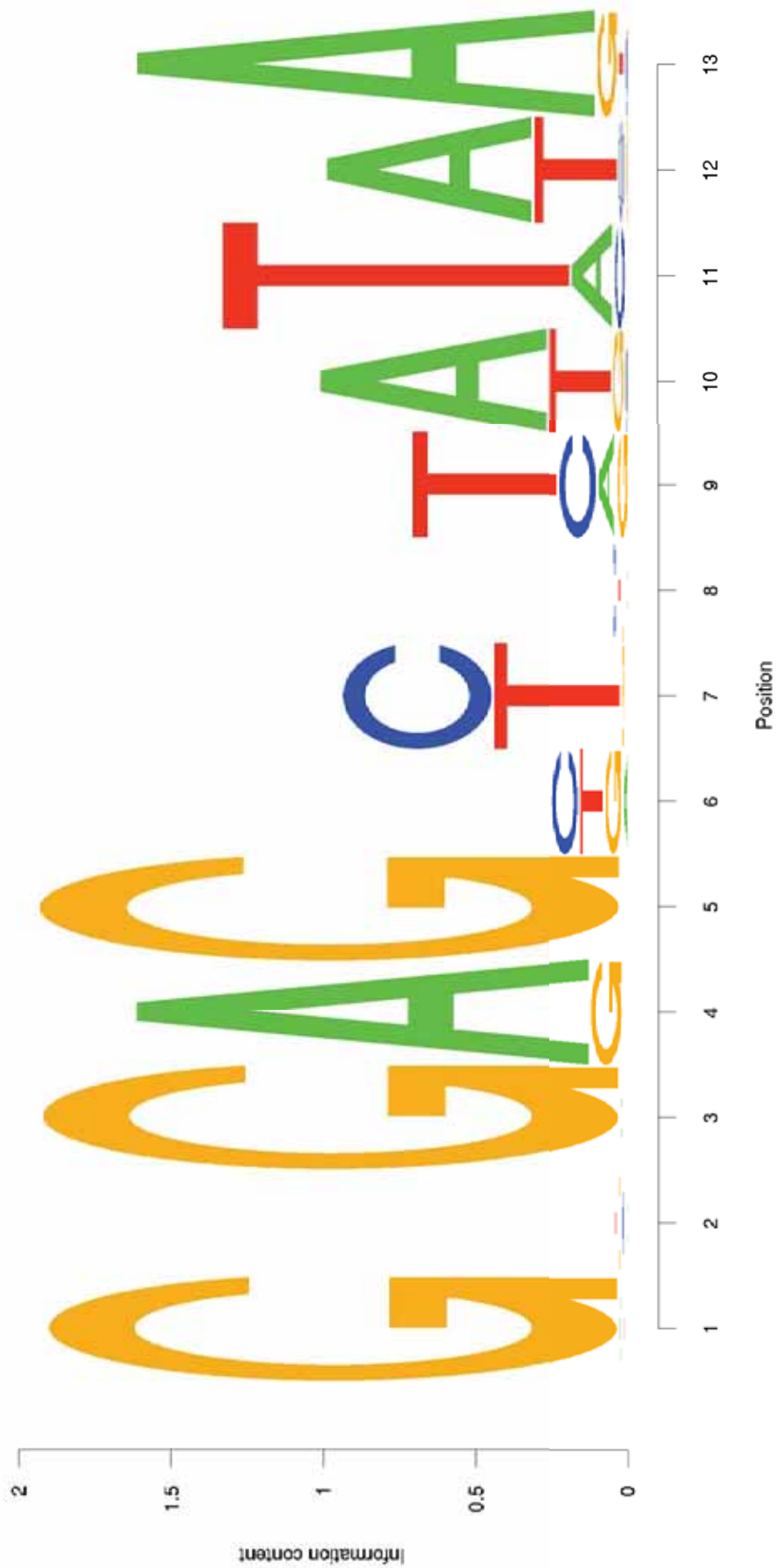
Bases relevant for RpoN-DNA recognition



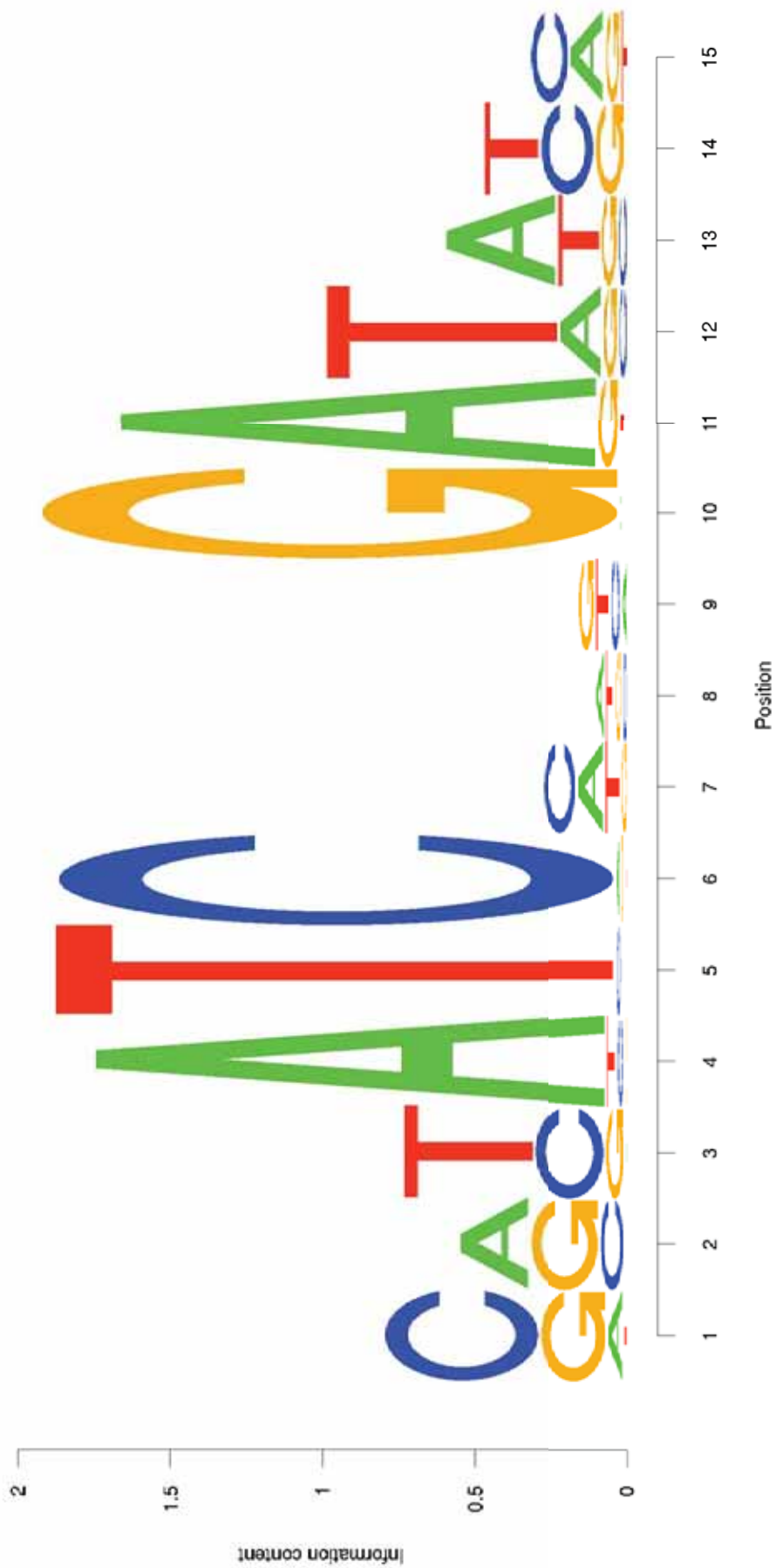
Towards a *bona fide* orthogonal regulatable promoter system



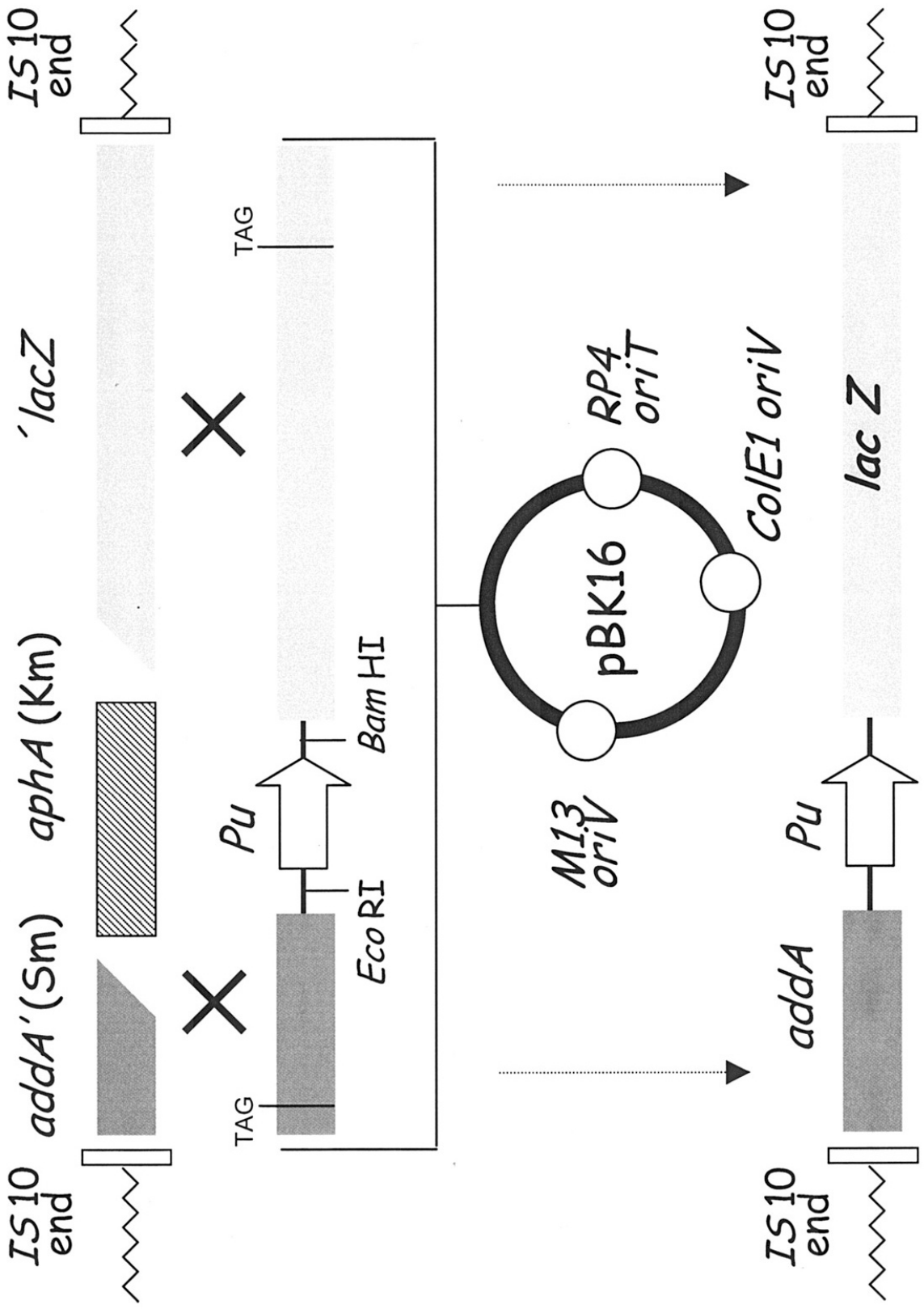
Bases relevant in T7Pol-DNA complex



Bases relevant for TerR-DNA complex



Chromosomal homology fragment



UAS

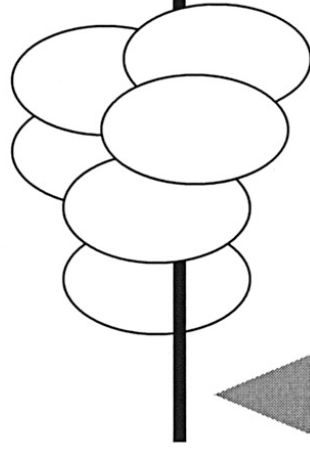
-190  -120

IHF

-24/-12 

XyIR

-79  -52

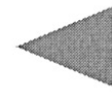


UP (-79)

UP (-104)

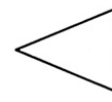
RNAP

σ 54

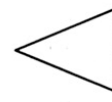


Sma I
(- 205)

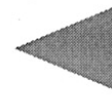
Eco RI (- 106)
pFH15



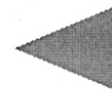
Eco RI (- 44)
pFH14



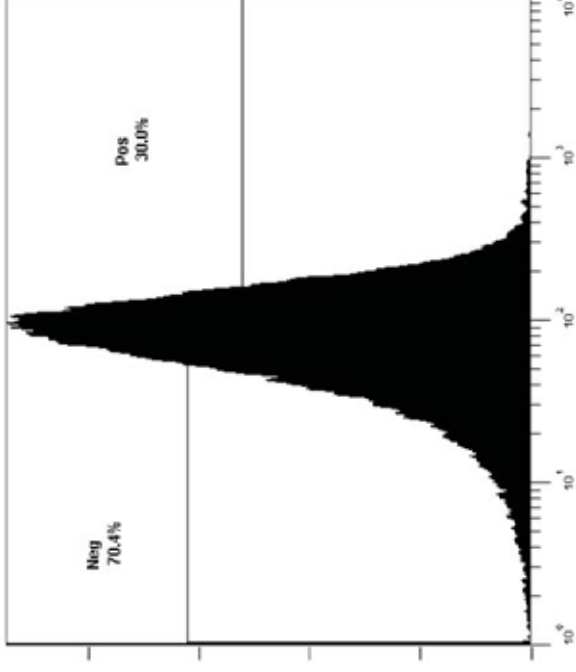
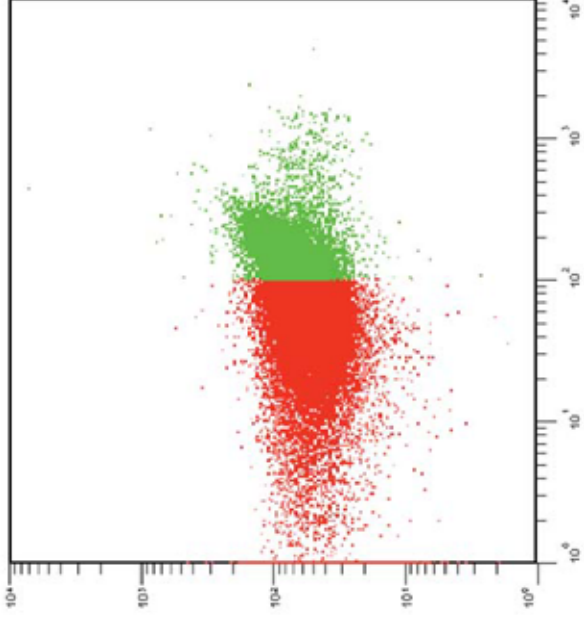
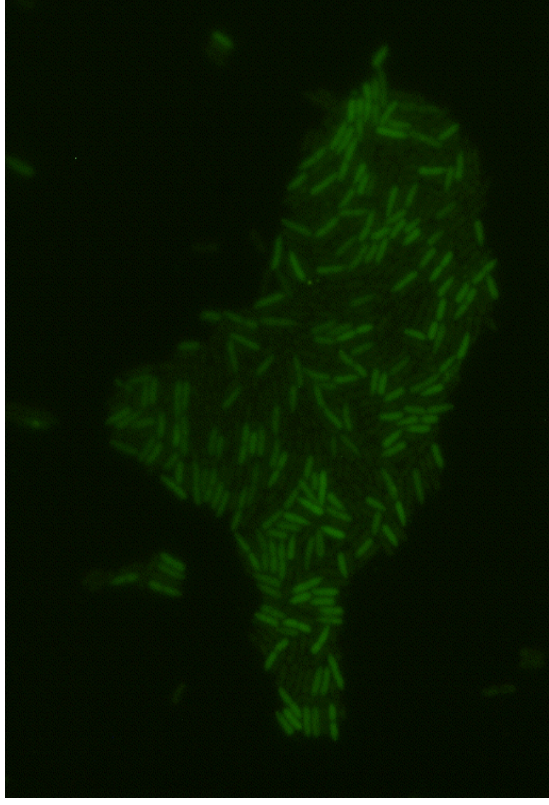
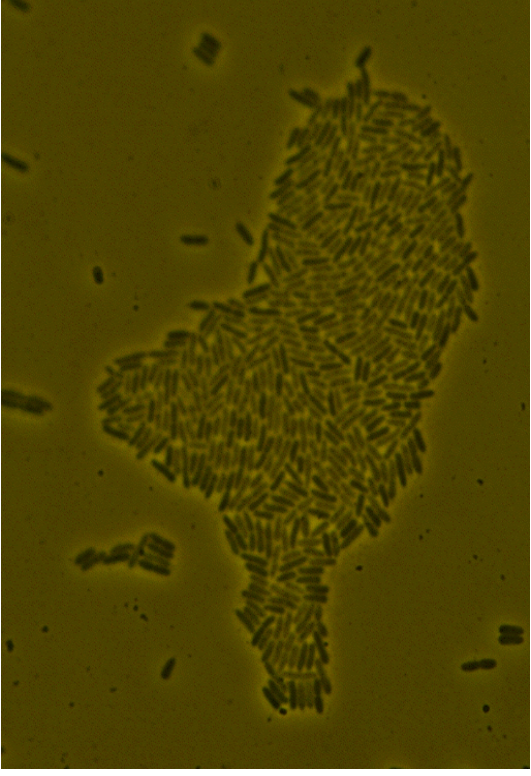
Bst EII
(+ 19)



Hae III
(+ 107)



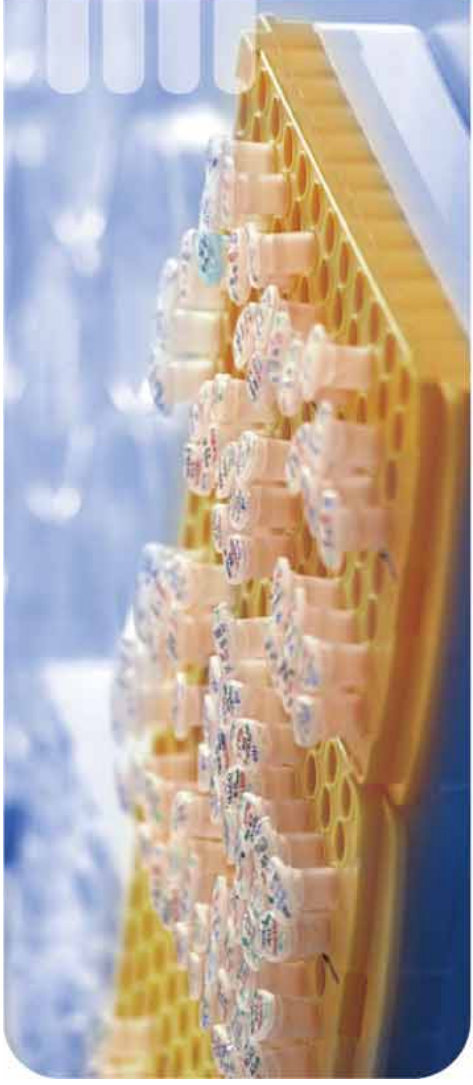
“Digital” response of XylR/Pu \rightarrow GFP to *m*-xylene exposure



Deliverables

D1.4. Report on recommendations of the intra-consortium expert group on suitable promoter standardization formats

D4.1. Database on quantitative prokaryotic promoter performance



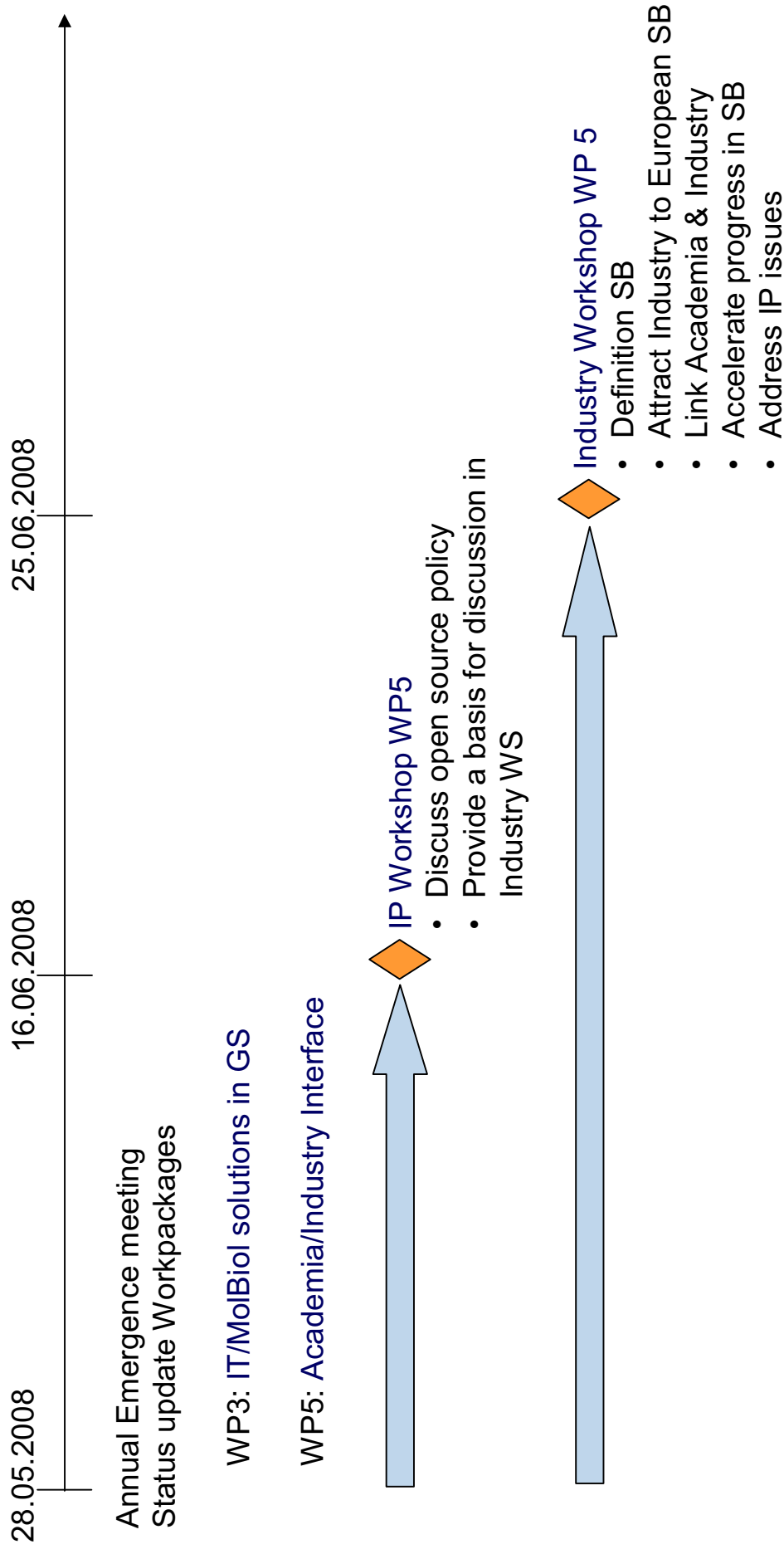
EMERGENCE Meeting Zürich May 2008

Workpackage 5: Building the Academia-Industry interface

Frank Notka, Ralf Wagner, May 2008

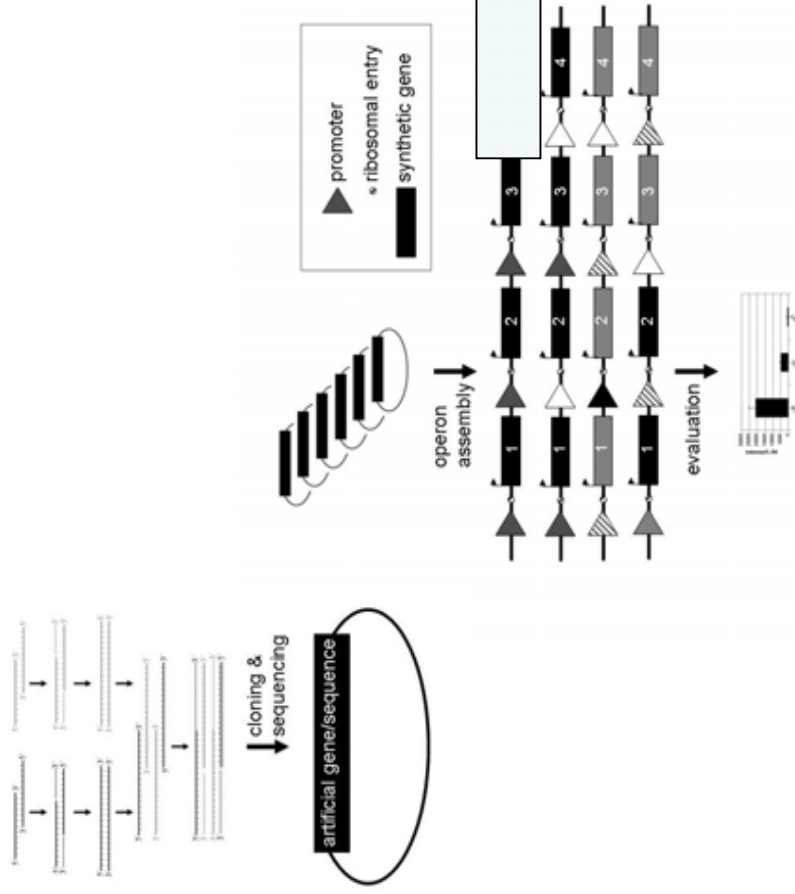
Status & Time lines

Contribution to different Workpackages



Strategies and tools for gene synthesis and assembly:

Gene sized segments can be assembled from synthetic oligonucleotides allowing maximum freedom for rational operon design. Differently designed elements in operons can be evaluated and optimized.:



- I) Design
 - adaptation to host
 - interchangeable parts
 - high flexibility
- II) Synthesis capacities
 - up to 2 mio bp/month
 - expanding automation platform
 - introduction of Laboratory Information Management System (LIMS)
- III) Variants of CDR/regulatory elements
 - epigenetic control
 - metabolic control
- IV) Functional circuit design
 - cooperative task

Workpackage 3 European IT Infrastructure for SB

Strategies and tools for gene synthesis and assembly:

Providing tools to effectively manage the complex and integrative process in genesynthesis to meet an expected increasing demand

Laboratory Information Management System (LIMS):

Operation level:

1. Order process
2. Gene optimization and design
3. Definition of oligonucleotides
4. Production and post-processing of oligos
5. *Gene assembly and cloning*
6. *Quality control*
7. *Export*

Information level:

1. Process steering
2. Process monitoring
3. Production parameters
4. Innovation management
5. Cost controlling
6. Customer support
7. Statistics

Workpackage 3 European IT Infrastructure for SB



Strategies and tools for gene synthesis and assembly:

Provide a strategy to evaluate parts in regard of biosecurity in order to avoid misuse

BioInformatics @ Geneart: Providing highest biosecurity level

Initial check of gene synthesis:

- (1) Country of customer (K-List, Embargo states)
- (2) Nature of customer (HADDEX List)
- (3) Nature of sequence (Internal data-base, blast)

Involvement of regulatory authorities/guidelines (BAFA and Australian group)

Check for associated pathogenicity/toxicity (dual-use components)

Based on these information a Go/No-Go decision is made

Workpackage 5

WS Define needs and interests of Industry

Objectives:

- To introduce Synthetic Biology and defining the industrial expectations, priorities and concerns
- To discuss topics regarding regulation, collaboration, and business challenges
- To link the actual development of Synthetic Biology in Europe with major projects, funding and network options

➔ Promote the Integration of Industry into the European SB development

Invited Participants (<http://spreadsheets.google.com/ccc?key=pMuMDXic0bDYn66S-6rJHAW&hl=en>):

Experts from leading European industries covering:

- Chemistry (Lonza, **BASF**, Novozymes, Henkel)
- Pharma (**AstraZeneca**, **Roche**, Novartis)
- Environment/Biomaterials (**Metabolix**, BASF)
- Energy (Shell, Butalco)
- Biotechnology (**Lifewizz**, Brain, Collectis) and

European academic Synthetic Biology exponents (DKFZ)

Workpackage 5

WS Define needs and interests of Industry



WS date: 25.06.2008 from 10 a.m. – 5 p.m.

Venue: Airport Munich
Hotel Kempinski

Welcome Address and Introduction

11:00 – 11:10 **Ralf Wagner**, GENEART AG

11:10 – 11:30 **Sven Panke**, ETH Zurich

11:30 – 12:00 Showcase by **Luis Serrano**, Centre for Genomic Regulation (CRG)

Topic I, SB and Chemical Manufacturing

12:00 – 13:00 Overview by **Sven Panke**, ETH Zurich & Discussion

13:00 – 14:00 Lunch

Topic II, SB and Pharma Research

14:00 – 15:00 Showcase by **Martin Fussenegger**, ETH Zurich & Discussion

Topic III, SB and IP Landscape

15:00 – 16:00 Open discussion

16:00 Final discussion & Closing remarks

Workpackage 5 WS on IP issues



WS date: 16.06.2008 from 10 a.m. – 2 p.m.

Venue: Munich
to be announced

Participants: Sven, Joachim Henkel, Berthold Rutz, DSM and Geneart

Main Goal: To develop a satisfactory and sustainable concept to regulate patent issues between participants, e.g. while dealing within a Registry network

Specific Objectives:

- Start an European IP discussion (provide a basis for the Industry WS)
- Is there a specific European aspect that we can contribute to the IP issue?
- Link the different perceptions of the academic and the industrial R&D processes (accelerated development vs. exploration of IP rights)

Proposed Agenda:

- Sven Panke (ETH): MIT Registry and IP
- Jo Henkel (TUM): SynBio and open source?
- Berthold Rutz (EPO): Scenarios for Synthetic Biology IP solutions
- DSM/Geneart: The industrial perspective
- Discussion: What is a realistic goal for a registry-related IP strategy?
Will a (European?) registry drown in IP problems

Workpackages 3 and 5: Deliverables

| Deliverable | Month | Progress |
|--|---------------------|---|
| 3.4 Document describing the proof-of-concept study exploiting the integrated workflow for genetic circuit design | 12/09 | In progress |
| 5.1 Reports on two industry workshops <ul style="list-style-type: none"> ■ to define the priorities of the European industry in the field of synthetic biology, and ■ to evaluate the fit of the European synthetic biology projects with the industry needs | 06/07 & 06/08 | Delay (involvement in 10/07) 1. WS fixed, registration i.p. |
| 5.2 Reports on two workshops (associated to industry-relevant scientific conferences) to teach the industry in synthetic biology concepts and tools | 12/07 & 12/08 | Dependent on 5.1 |
| 5.3 Position paper on the priorities of the European industry in the field of synthetic biology, evaluation of fit with current EU synthetic biology projects, and decision on how to address the potential gaps | 12/08 | In progress |
| 5.4 Intermediate and final report on status of discussion regarding IP strategy in the field of synthetic biology, originating from company internal assessments and summarizing the ideas on IP-management (same workshops as in D5.1) | 12/07 & 12/09 | Delay (involvement in 10/07) 1. WS fixed |