



About us

Technology

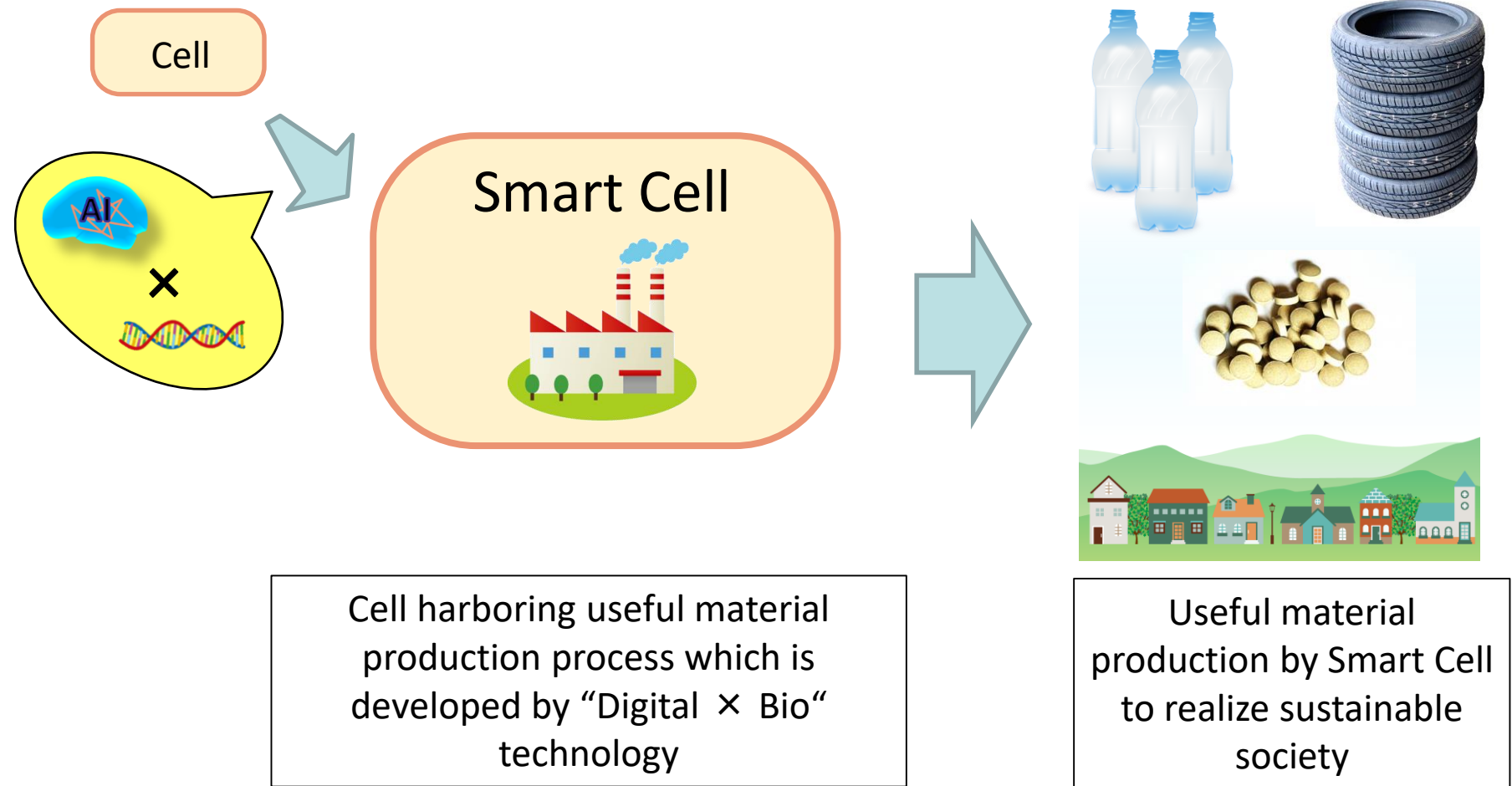
Precision, long-chain DNA.
Remarkable efficiency. No compromises.



<https://www.synplogen.com/>

Smart Cell Industry

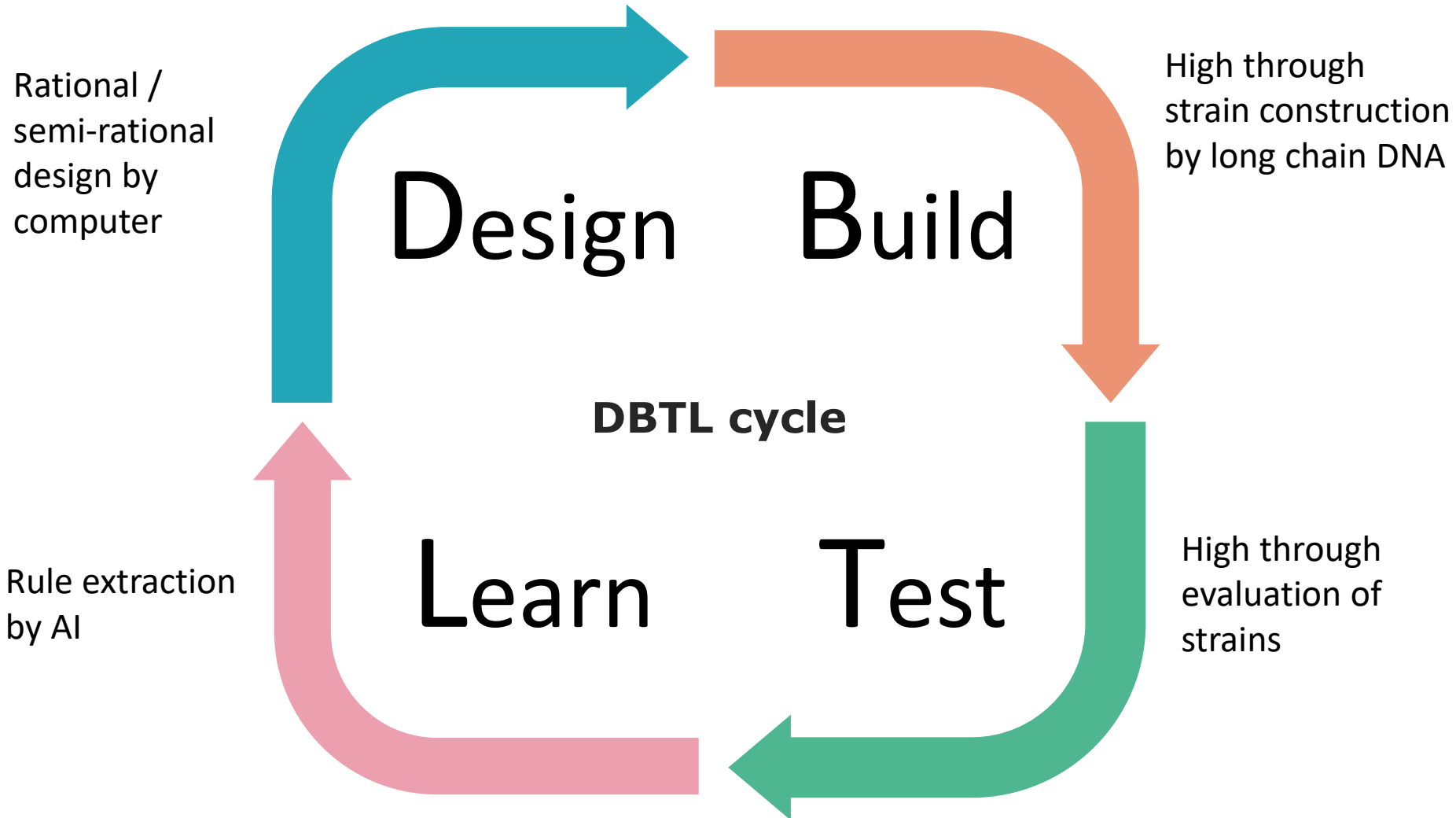
Smart Cell Industry



This figure is modified from original figure in NEDO booklet (focus NEDO, No.70).

DBTL cycle –Platform for Synthetic Biology

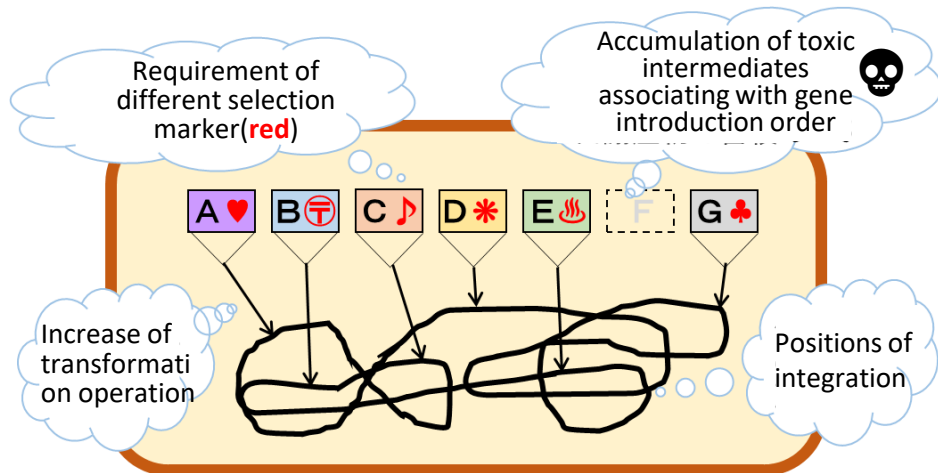
Fine tuning of multiple parameters of gene circuit in single step is almost impossible!!
Repetitive cycling of DBTL is necessary to achieve higher production



Why we need long DNA?

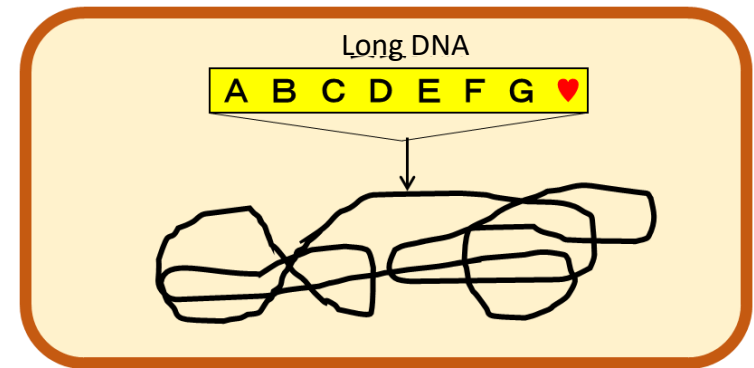
Long DNA contributes to reduce both money cost and time cost for designer organism construction

Conventional genetic manipulation



takes much money and time

Long DNA strategy



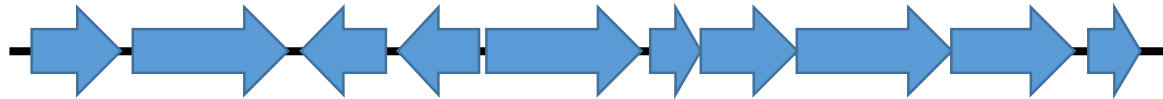
finishes within one-step

What is the size of “Long DNA”

We focus on size range of 10 to 50 Kb at initial business
But we are developing method to construct up to Mega-size DNA

How many genes in 10 kb DNA ?

Prokaryote



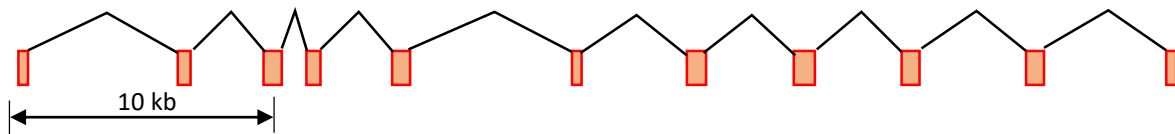
Averagely 10 genes / 10 Kb

Lower Eukaryote



Averagely 5 genes / 10 Kb

Human



Averagely 1 gene / 53 Kb

We use *Bacillus subtilis* for long DNA construction

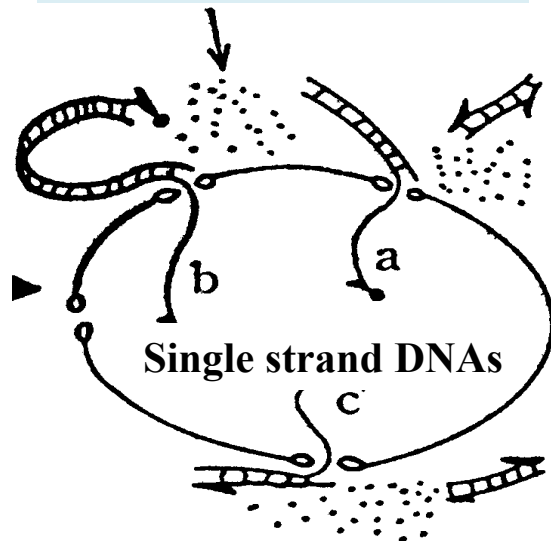
B. subtilis is good at large DNA manipulation

Natto (Japanese traditional food)



Safe bacteria

Natural transformation

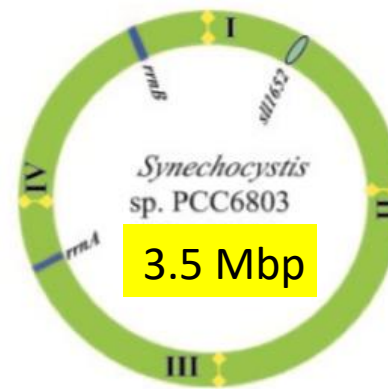


Canosi, U., et al. MGG 181, 434-440 (1981)

First instance of hybrid genome of 2 bacteria

Itaya, M., Tsuge, K., Koizumi, M., Fujita, K. *PNAS*, 102, 15971-15976 (2005)

Mbp-cloning feasible



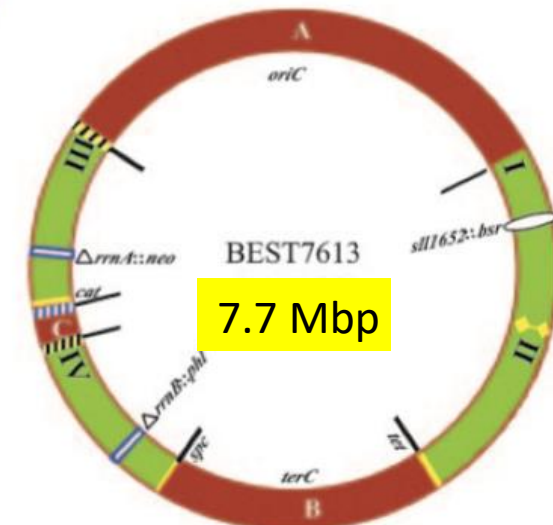
3.5 Mbp



Bottle ship construction manner



4.2 Mbp



7.7 Mbp

Key technology: OGAB method (1st Gen.)

Ordered Gene Assembly in *Bacillus subtilis*

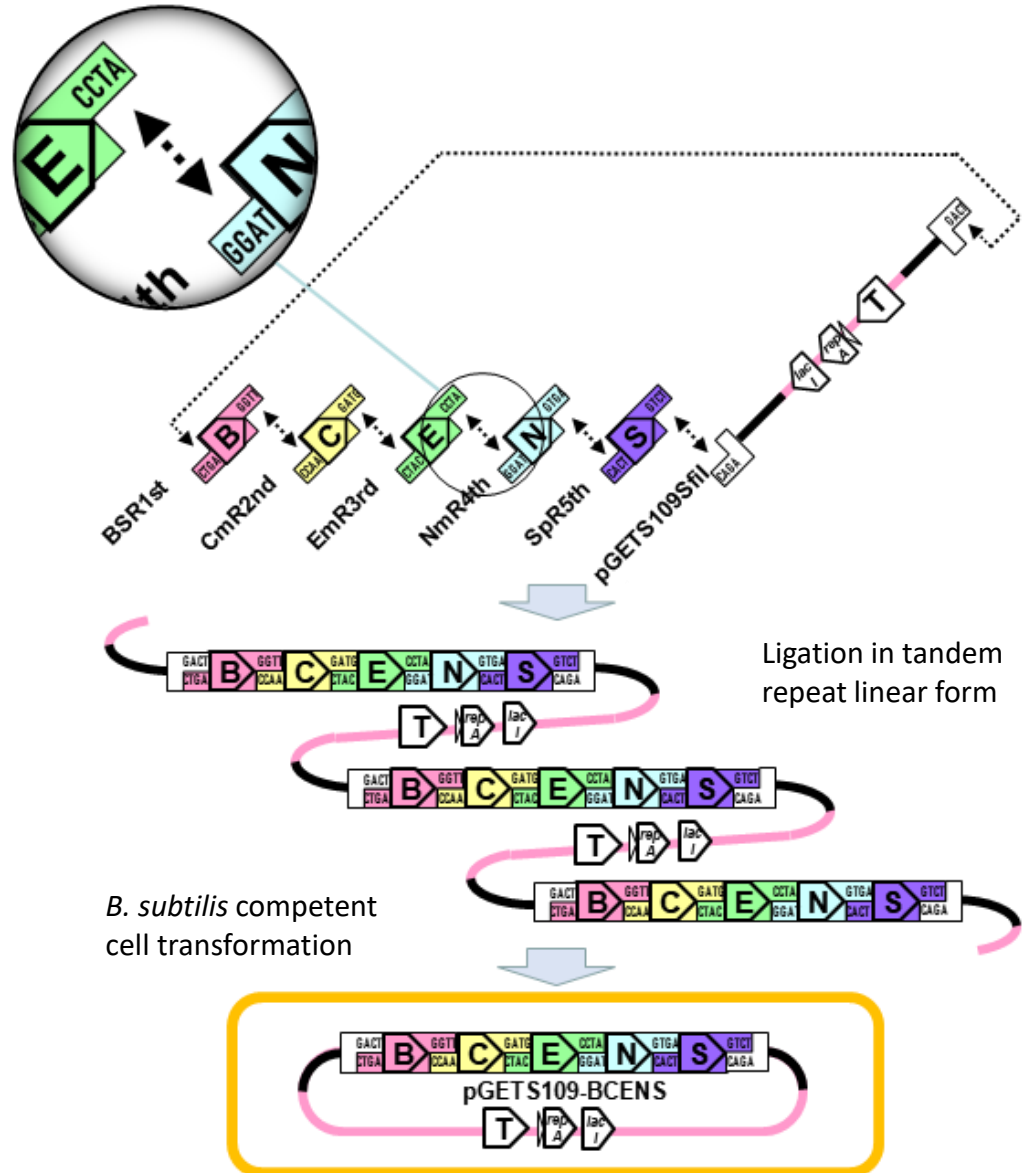
Tsuge, K., Matsui, K., Itaya, M.

Nucleic Acids Res. 31,e133 (2003)

Patent No. 4479199 (Japan) Synplogen

Assembly method using *B. subtilis* plasmid transformation system

- Assembly completes in one-step
- High efficiency and fidelity
- No limit on fragment size
- Feasible for up to **15-fragment** assembly

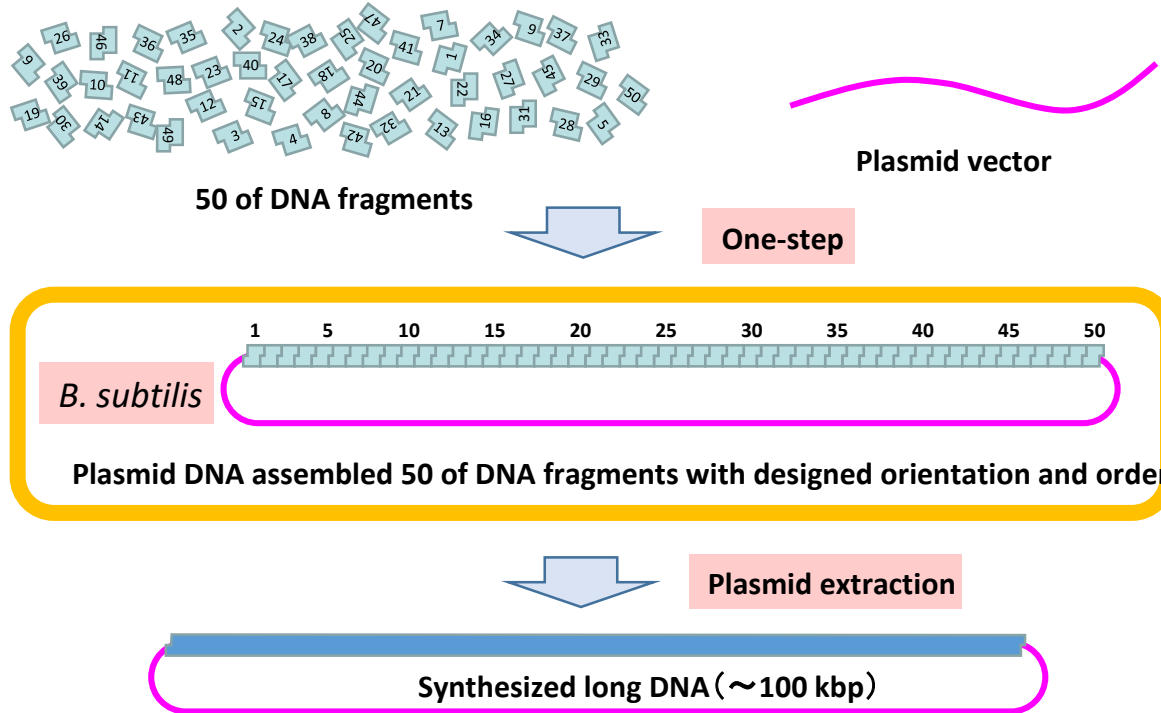


Automation system for 2nd Gen. OGAB

By adjusting size of material DNA fragments strictly, gene assembly of more than 50 fragments is feasible

Tsuge, K. et al. *Sci. Rep.*, **5**, 10655 (2015)

WO 2015/111248 Synprogen



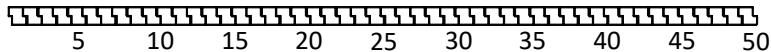
- Feasible to construct ~100 kb DNA with designed sequence
- Very precise DNA compare to other method
- Construction finishes with in one-step
- Automation friendly

Detail of 2nd Gen. OGAB method

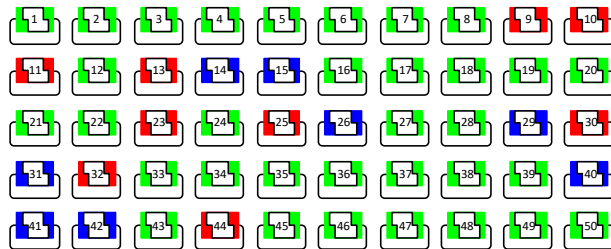
Rationally simplified with elevated reliability

Tsuge, K. et al. *Sci. Rep.*, **5**, 10655 (2015).

Designed sequence



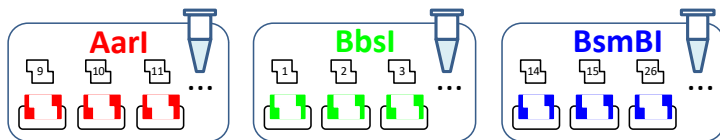
50 of OGAB block plasmids
(All plasmids are designed to be equal length (CV = 0.2%))



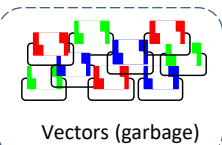
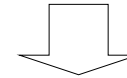
Equimolar adjustment
with microspectrometer
(CV = 2%)



Dispense equimolar DNA to each tube according to restriction enzyme use / TypellS restriction enzyme digestion

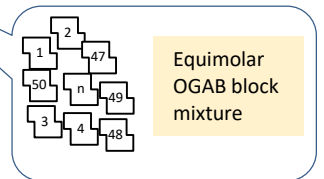


Gel electrophoresis
separation of OGAB
blocks

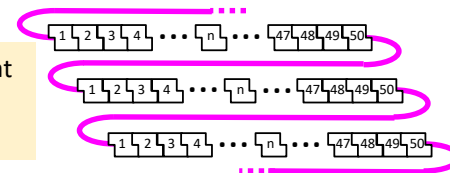


Plasmid vector for
OGAB assembly

+

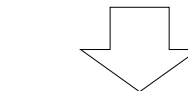


Tandem repeat
form ligation
products



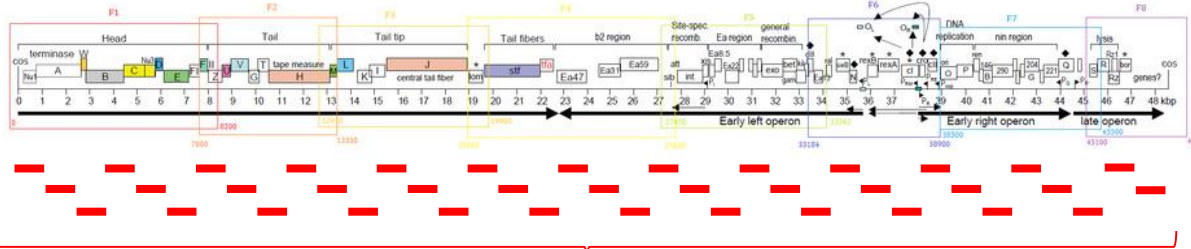
B. subtilis

Assembled plasmid



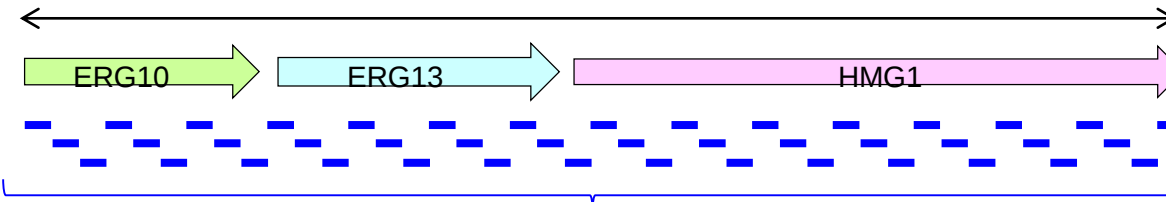
Examples of long DNA by 2nd Gen. OGAB

Lambda phage (48.5 kb)



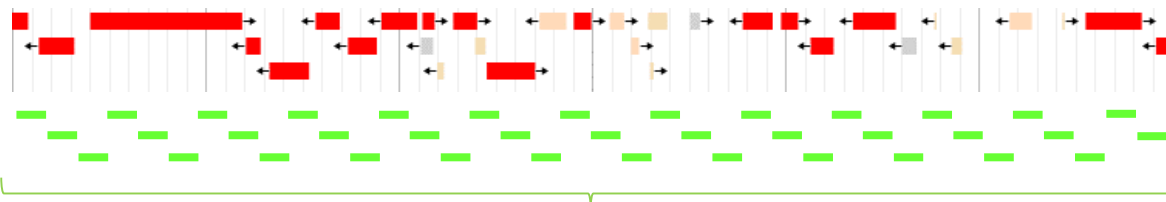
Ave. 970 bp × 50 OGAB blocks

Artificial mevalonate operon (6 kb)



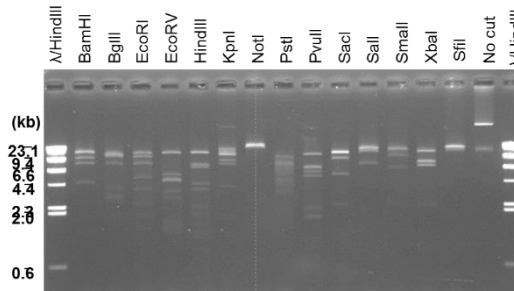
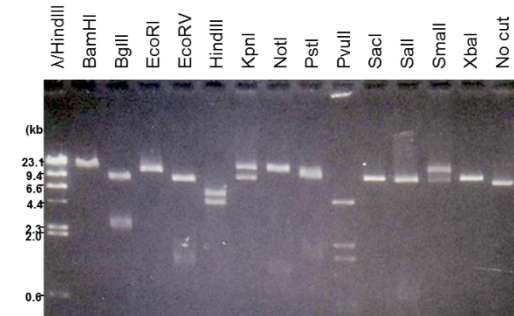
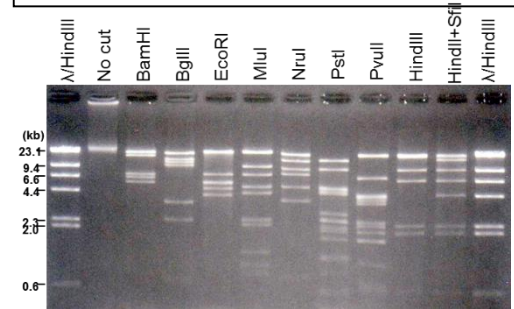
Ave. 110 bp × 55 OGAB blocks

Yeast artificial genome (30 kb)

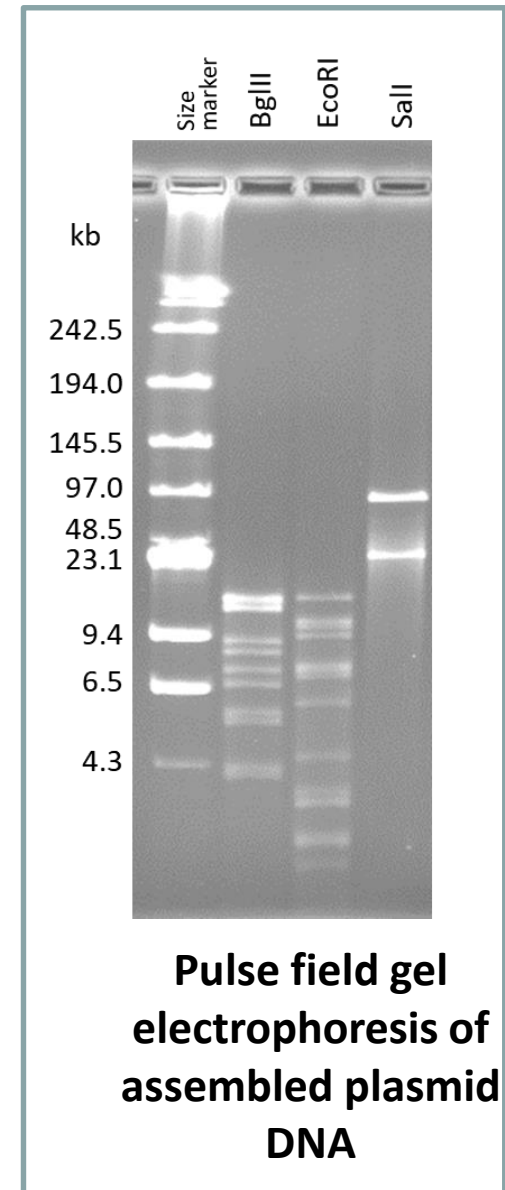
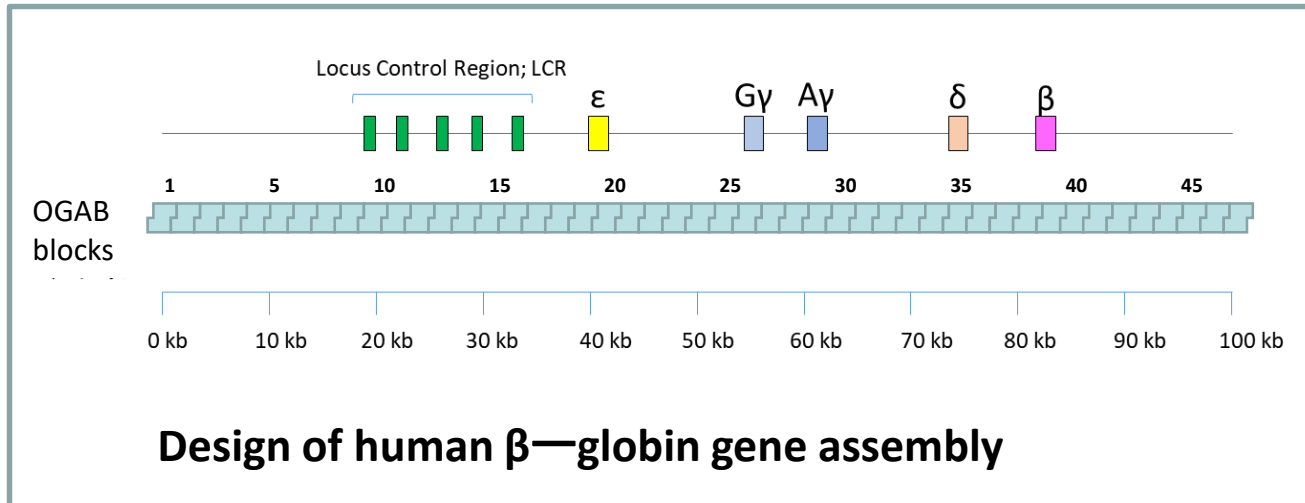


Ave. 620 bp × 48 OGAB blocks

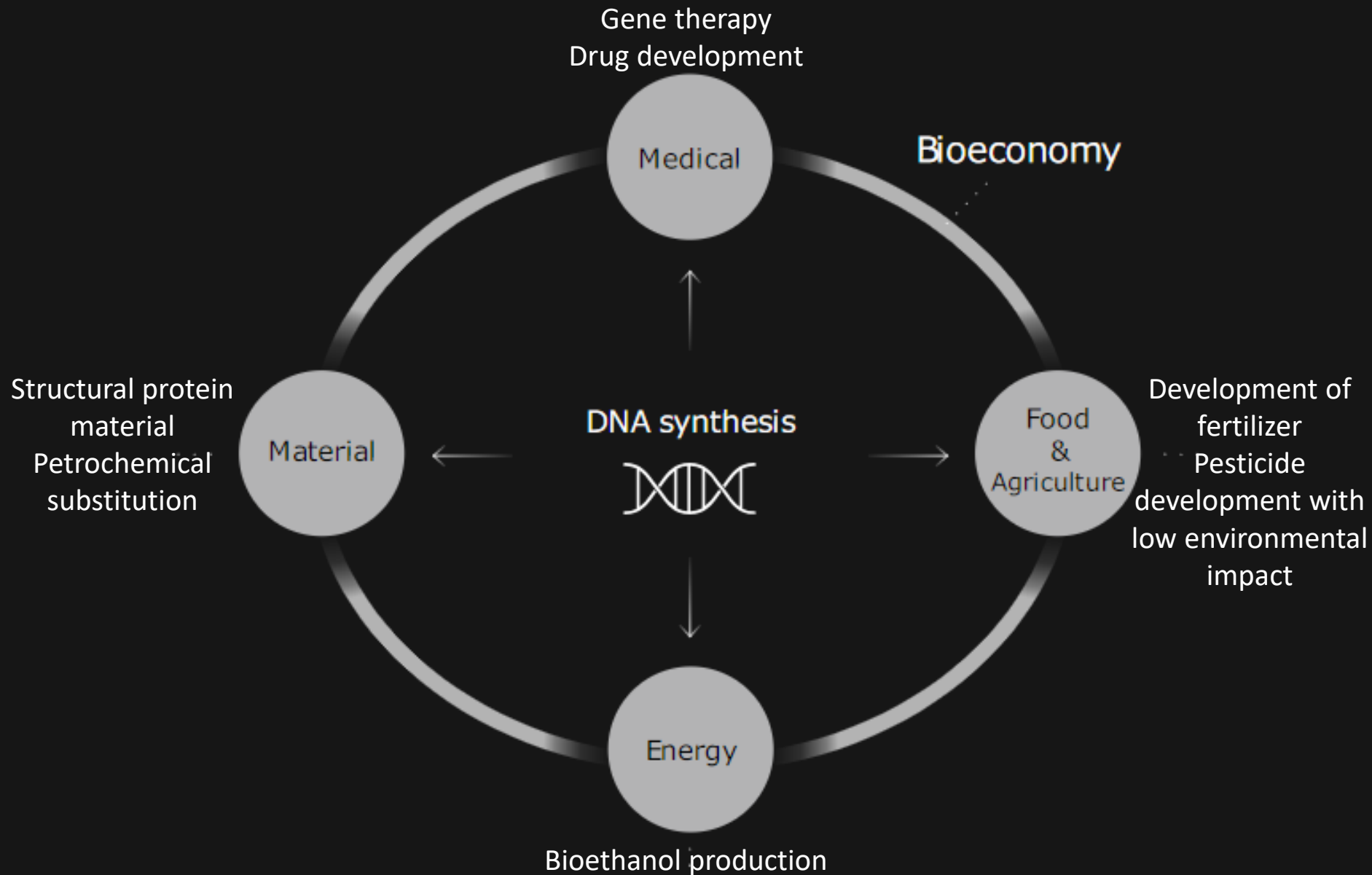
Restriction enzyme patterns



Successful re-construction of 100 kb human gene



Business field of Synplogen



Services

Long-chain DNA synthesis

Synthesis of long-chain DNA (up to 100 kbp)

Combinatorial DNA library
synthesis

Synthesis of DNA libraries using combinatorial design

Support for developing
production-use microbes

Assistance with the design and synthesis of gene
clusters for the development of production-use
microorganisms

About Us

Company Name	Synplogen Co., Ltd.	
Address	HQ: 1-1 Rokkodaimachi, Nada-ku, Kobe-shi, Hyoto, 657-0013, Japan Branch office: 234-1 Mizukami, Kakuganji, Tsuruoka-shi, Yamagata, 997-0052, Japan	
Founded	February 21, 2017	
Paid-in Capital	¥118 million (incl. capital reserves)	
Board of Directors	+ CEO	Junichi Sugahara, PhD
	+ Director	Akihiko Kondo, PhD
	+ Director	Kenji Tsuge, PhD
	+ Director	Kazuhiro Yamamoto
	+ Director	Yoshihiro Oshima
	+ Director	Daniel Meyer
E-mail	info@synplogen.com	