

CSIR Synthetic Biology and Precision Medicine Centre Bio-foundry Program

Deepak B. Thimiri Govinda Raj^{1*}

¹Synthetic Nanobiotechnology and Biomachines, Synthetic Biology and Precision Medicine Centre, Council for Scientific and Industrial Research (CSIR), Pretoria, South Africa.

*-Corresponding author: dgovindaraj@csir.co.za

Background: Global bio-foundry Alliance (GBA) has been established between countries including the UK, US, Japan, Singapore, China, Australia, Denmark, and Canada through 16 research institutions. Global bio-foundry Alliance plays the key role in the synthetic biology driven towards a new global bioeconomy that is accelerated by advanced technology innovation. Establishment of Biofoundry program in South Africa and in Africa will plan key scientific and the strategic role in promoting synthetic biology and precision medicine program in Africa. This would further enable bioeconomy and industrial development towards SME program. At our CSIR Synthetic Biology and Precision medicine Centre, we are currently establishing biofoundrylab that will implement various synthetic biology and precision medicine projects in South Africa.

Methods: We are currently establishing two research components in the CSIR Synthetic Biology and Precision Medicine Centre Bio-foundry program which includes industrial synthetic biology and functional precision medicine program. We implement Biofoundry biodesign and biological engineering Design-Build-Test-Learn (DBTL) cycle into our industrial synthetic biology and functional precision medicine program. In our Industrial synthetic biology program, we are working on a) ValitaCHO: Development of superior CHO cell line system for hyper-burst protein expression system using directed evolution and synthetic biology approaches; b) Lactochassis: Designer microbes for industrial synthetic biology platform applications; In our Cancer Precision Medicine program: we are working on drug repurposing based drug sensitivity screening platform for B-cell malignancies and ovarian cancer treatment for South African patient cohort.

Results: We are currently at the Design phase of the Design-Build-Test-Learn (DBTL) cycle in our industrial synthetic biology and functional precision medicine program. We have so far have progressed in generation of the preliminary data on ValitaCHO cell-line chemstress fingerprinting profiling. We are currently designing the directed evolution approach for generation superior CHO cell line. In the Lactochassis project, we are currently designing the computational biology based genome mapping for Lactochassis. In our precision medicine platform, we are currently progressing in design and build phase of platform where we have currently procured 770 cancer drugs for drug repurposing platform which can be applied for blood and ovarian cancer cohort.

Conclusion: Using Bio-design DBTL cycle, we aim to implement our industrial synthetic biology and cancer precision medicine platform at CSIR Synthetic Biology and Precision Medicine Centre. These platforms will enable establishment of one of the first Biofoundry labs in Africa.

Funding: This work is funded by ICGEB Early Career Grant, NRF unrated Grant and CSIR Parliamentary Grant for Synthetic Biology and Precision Medicine Centre, CSIR Pretoria, South Africa.

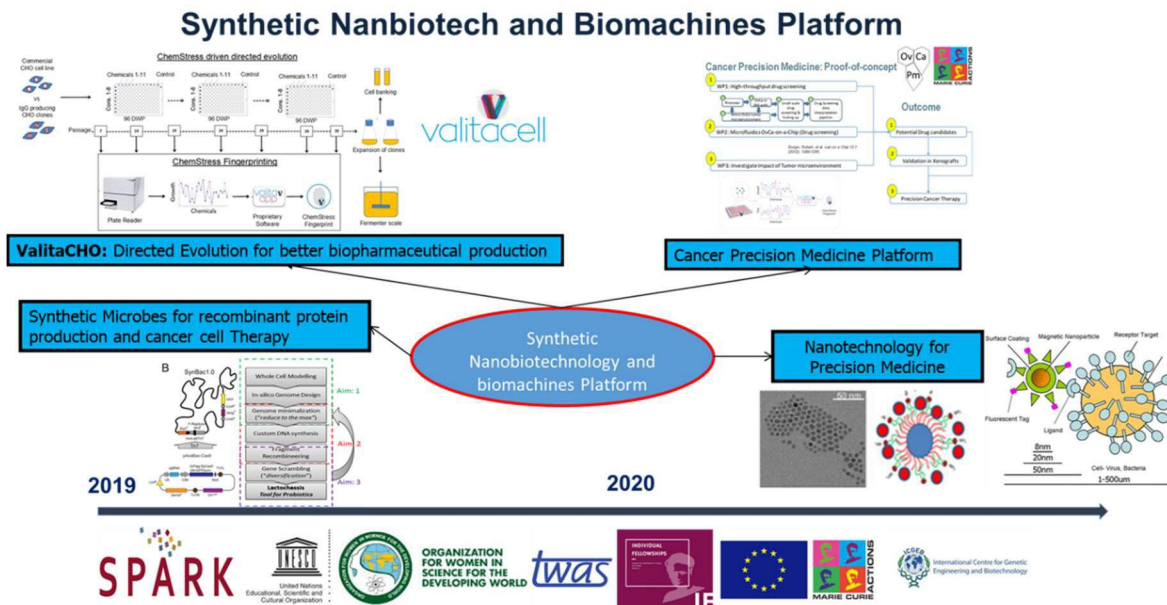


Figure 1: CSIR Biofoundry Program: Synthetic Nanobiotechnology and Biomachines Platform

References

1. Crocker H, Gorda B, Pelosse M, Thimiri Govinda Raj DB, Berger I, SynBac: Enhanced Baculovirus Genomes By Iterative Recombineering. *Methods in Mol.Biol.* 2021 in press.
2. Skånland SS, Cremaschi A, Bendiksen H, Hermansen JU, Thimiri Govinda Raj DB, MuntheLA, Tjønnfjord GE, Taskén K, An in vitro assay for biomarker discovery and dose prediction applied to ibrutinib plus venetoclax treatment of CLL. *Leukemia* in press 2019.
3. Vijayachandran LS*, Thimiri Govinda Raj DB*, Edelweiss E, Gupta K, Maier J, Gordeliy V, Fitzgerald DJ, Berger I.. Gene gymnastics: Synthetic biology for baculovirus expression vector system engineering. *Bioengineered* 2013 4(5).
4. Thimiri Govinda Raj DB, Ghesquiere B, Tharkeshwar AK, Coen K, Derua R, VanderschaegheD, Rysman E, Bagadi M, Baatsen P, De Strooper B, E.Waelkens, G.Borghs, Gevaert K, & Annaert W, A novel strategy for the comprehensive analysis of the biomolecular composition of isolated plasma membranes. *Molecular Systems Biology* (2011); 7:541.
5. Thimiri Govinda Raj DB, Cremaschi A, Skånland SS, Gade A, Schjesvold F, Tjønnfjord G, Munthe LA, and Tasken K., In-Vitro Drug Sensitivity Screening in Chronic Lymphocytic Leukemia (CLL) Primary Patient Samples Identifies Drug Candidates for Precision Cancer Therapy. *Blood* 2018 132.
6. Thimiri Govinda Raj DB, Giliberto M, Cremaschi A, Skånland SS, Gade A, Tjønnfjord G, Schjesvold F, Munthe LA, Tasken K., Drug Sensitivity Screening on MM for Precision Cancer Therapy *Blood* 2018 132 1: 4677.
7. Thimiri Govinda Raj DB, Berger I., Improved baculoviral expression systems and methods of producing the same: EP14708951, US 61/ 762. 607 and WO2014122629
8. Thimiri Govinda Raj DB, Lagae L, Annaert W, Borghs G, Plasma membrane isolation. 2015US8936935, EP2388589.