

SYNTHACE ■■

**SOFTWARE TO INCREASE THE USE OF
AUTOMATION IN
LIFE SCIENCES**

Spring 2019 Allotrope™ Connect Meeting

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INTRODUCTION



People: 55



Location: Central London & USA (Q2 2019)



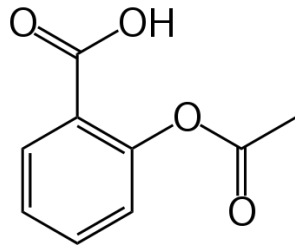
History:



Funding: \$44m raised to date

CHALLENGE: EVER INCREASING BIOLOGICAL COMPLEXITY

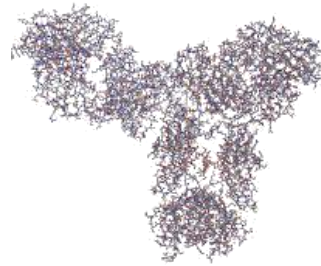
SMALL MOLECULES



Aspirin

21 atoms

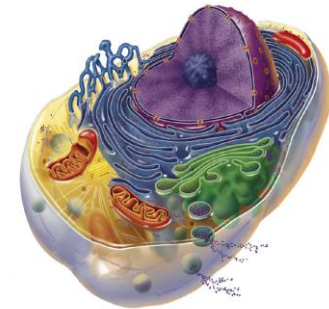
BIOLOGICS



Humira

20,067 atoms

CELL & GENE THERAPY



Human cell

10^{14} atoms

CHALLENGE: EVER INCREASING BIOLOGICAL COMPLEXITY

English
Representation
of Biological
Processes

+

Manual
Execution

+

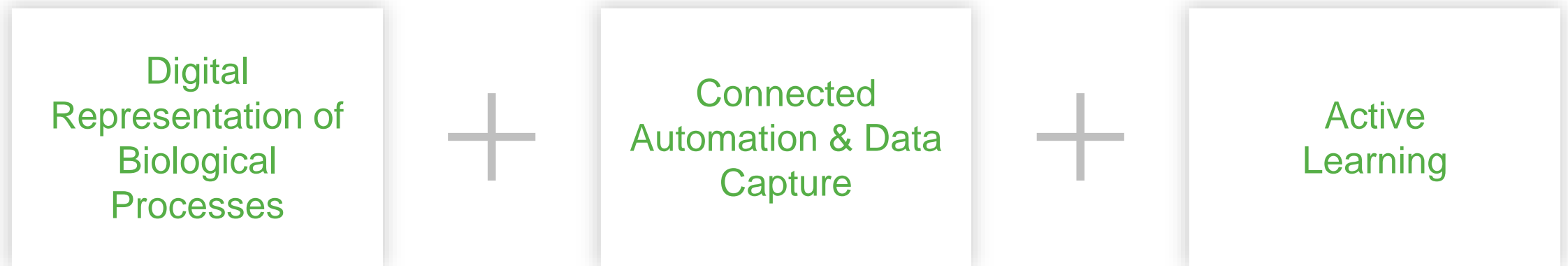
Artisanal
Insight

NOT FIT FOR PURPOSE

.... Eroom's Law....

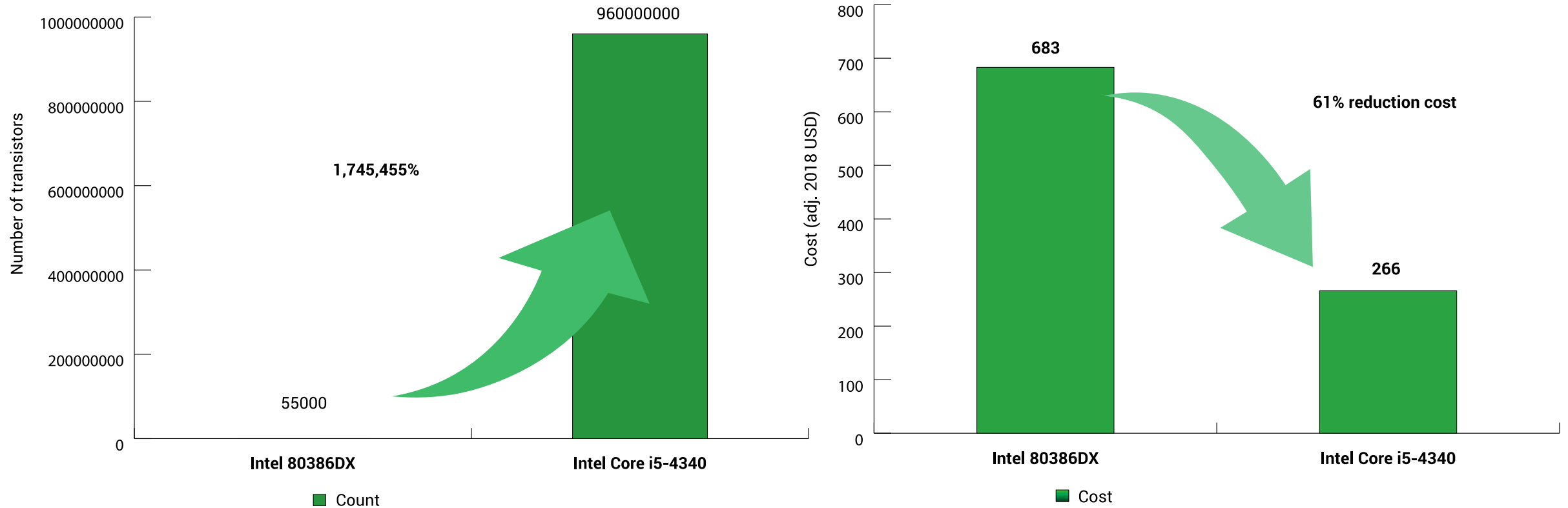
A NEW WAY OF WORKING: COMPUTER AIDED BIOLOGY

A way of working, reliant upon Automation and Analytics



BASIC CONCEPT – MANUAL TO AUTOMATED

OTHER INDUSTRIES HAVE REDUCED COST WHILST DELIVERING AN INCREASINGLY COMPLEX PRODUCT



Electronic Design Automation tools comprised of abstracted design and simulation allowed for the rapid design, testing and production of increasingly complex integrated circuits; in part leading to Moore's Law



A GROWING INTEREST IN AUTOMATING BIOLOGY

The Economist

Topics ▾

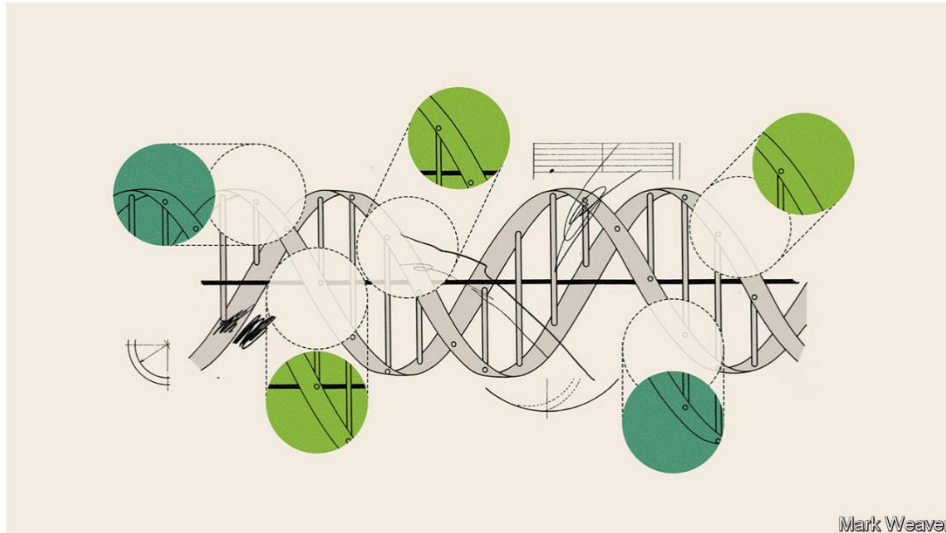
Current edition

More ▾

Automation

Remaking life means automating biology

Synthesised DNA enables doing biology on an industrial scale



Mark Weaver

Print edition | Technology Quarterly >

Apr 4th 2019



Biotech

+ Add to myFT

Microsoft moves into biological computing with Station B

Tech giant rolls out new system to analyse vast volumes of biomedical data



A scientist at work in Oxford BioMedica's main facility in the UK © Jonathan Banks for Microsoft

Clive Cookson in London MARCH 12, 2019

5

Microsoft will make a big move into [biotechnology](#) on Tuesday with the launch of a new research system that enables scientists to engineer living cells using machine learning and data analysis.

COMPUTER AIDED BIOLOGY: A NEW WAY OF WORKING



DIGITAL
Powered by AI

Models

Bio Design

Bio Simulation

Experiment Design

Experiment Simulation

Experiment Execution

Building Blocks

Analysis

Data Management



PHYSICAL
Powered by
Automation

SYNTHACE II

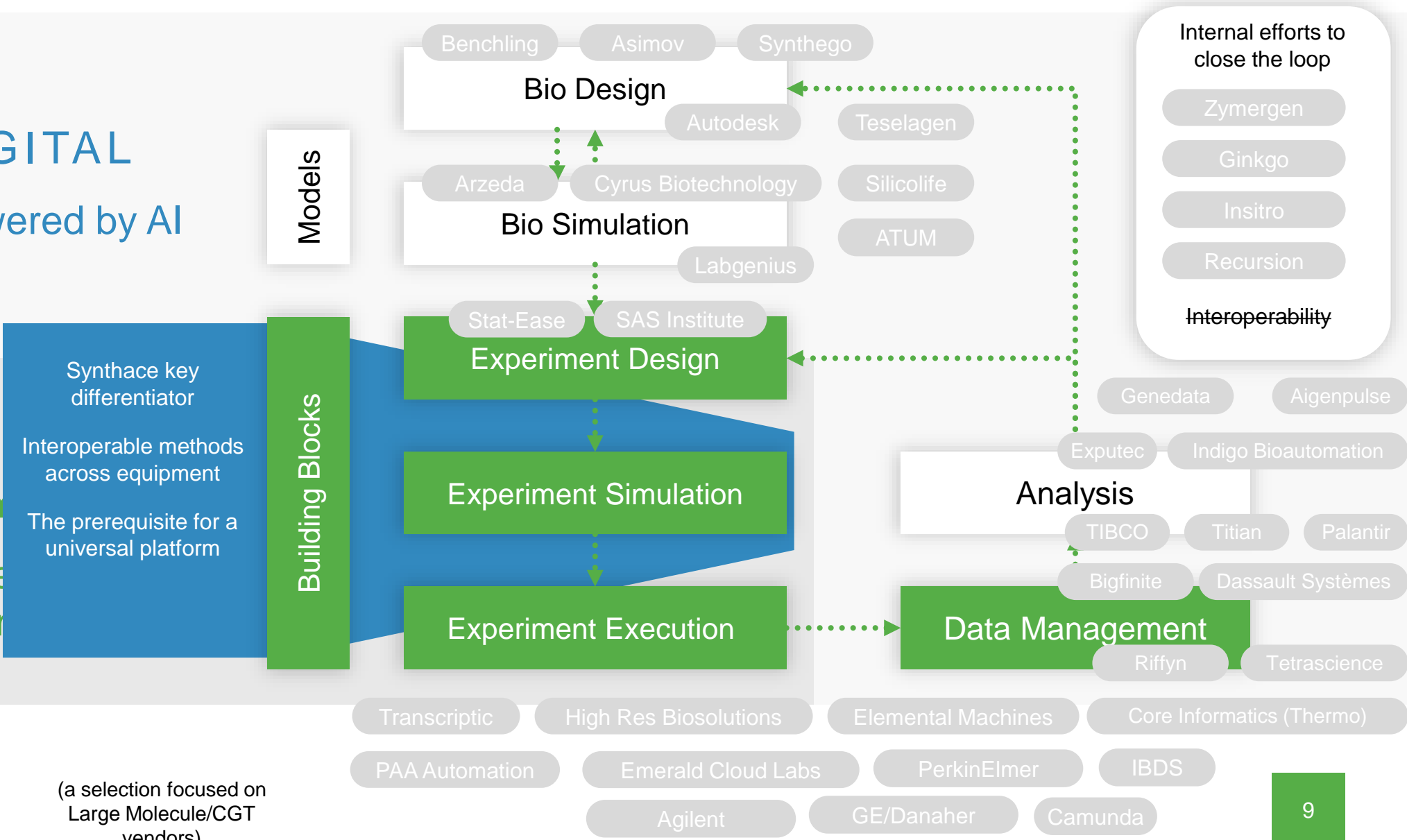
COMPUTER AIDED BIOLOGY: AN EMERGING ECOSYSTEM



DIGITAL
Powered by AI



PHY
Power
Autom



WE USE CAB TO TEST OUR OWN SOFTWARE





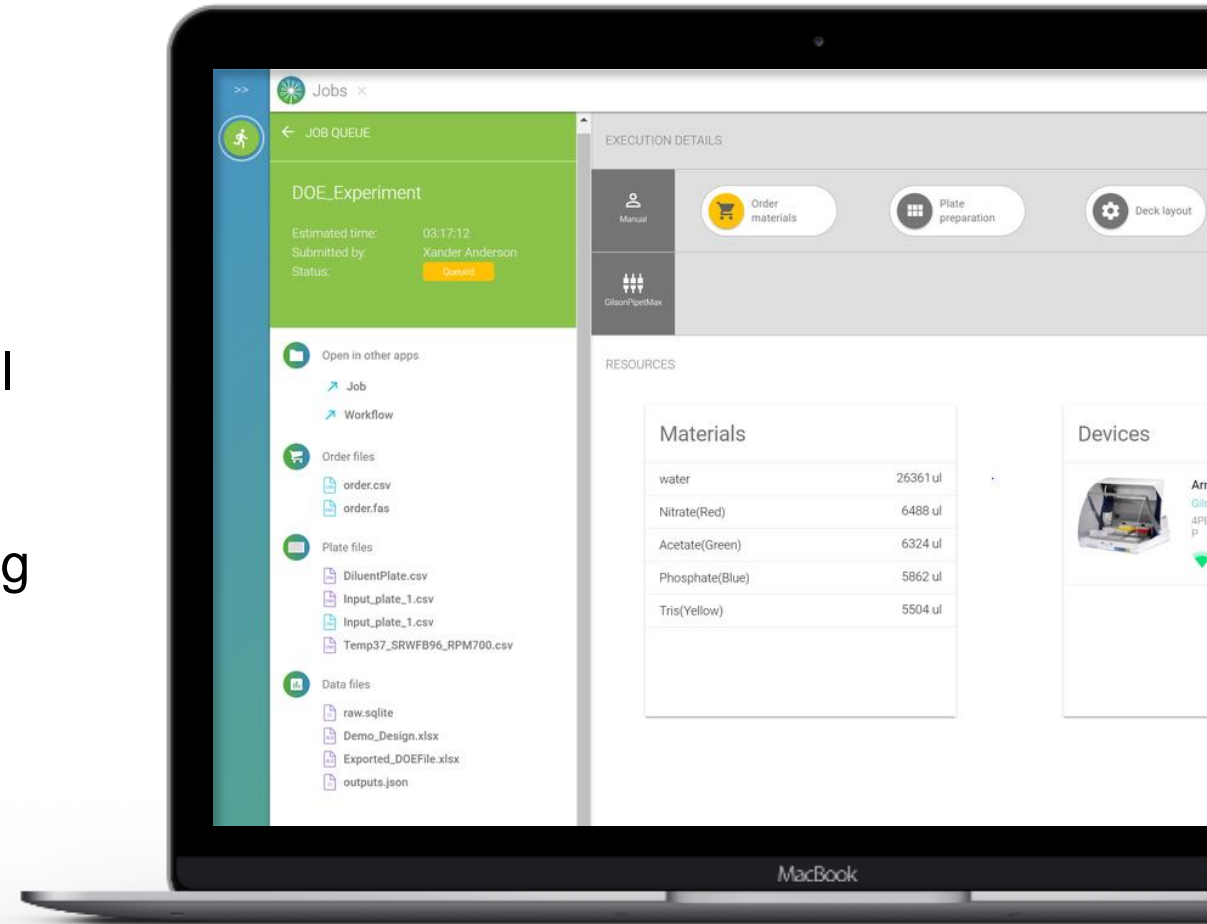
Case studies...

OUR PLATFORM



A software platform for interoperable experimental execution and data integration.

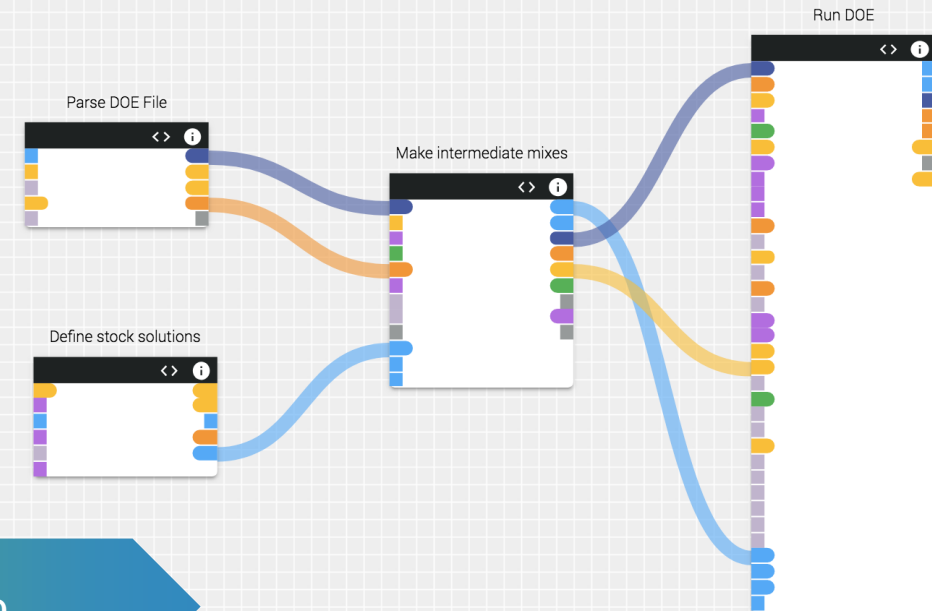
Built with a high-level language for biology, making it easy to rapidly compose and execute reproducible workflows using individually testable and reusable Antha Elements



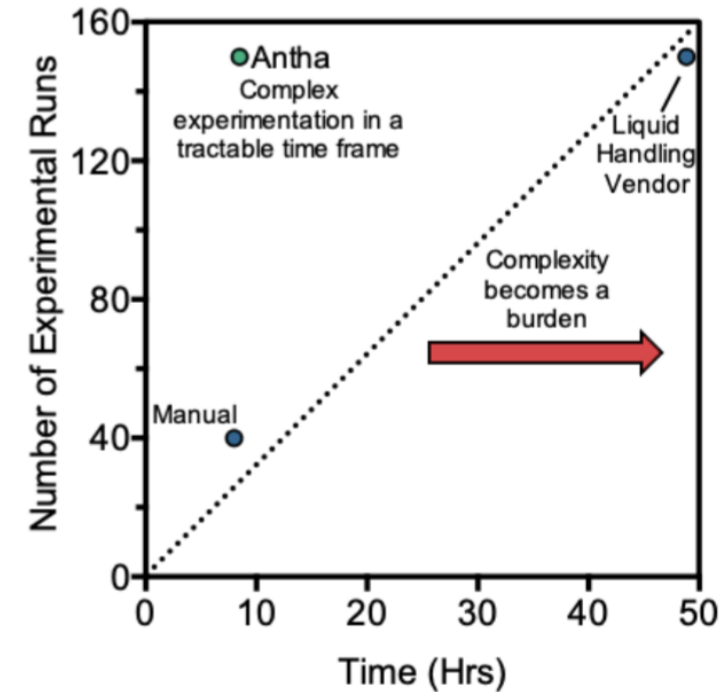
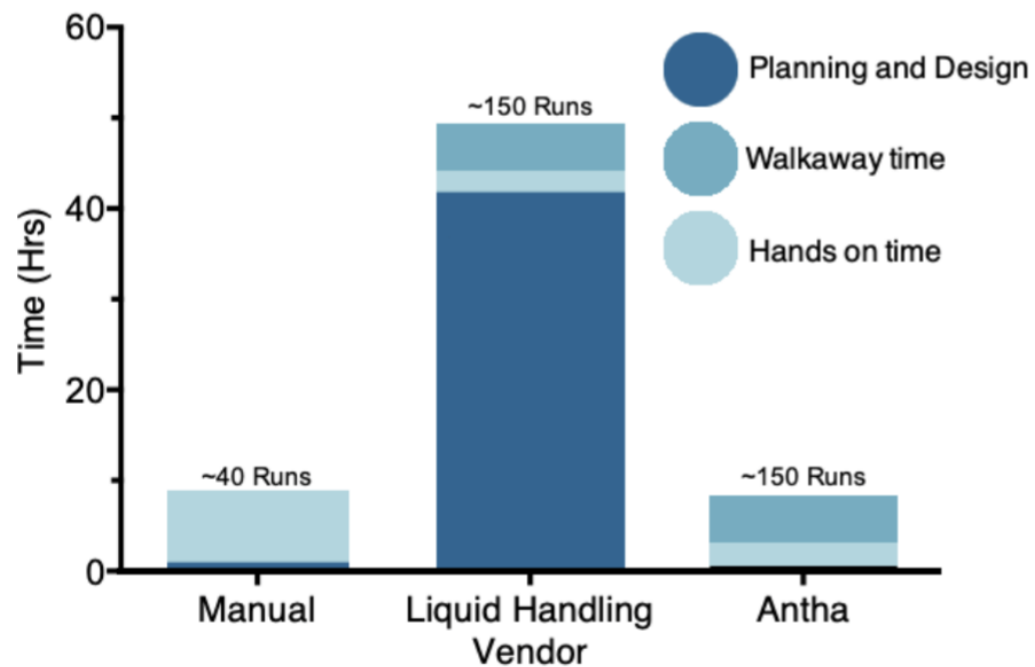
CASE STUDY: ABSTRACTED DESIGN OF EXPERIMENTS AUTOMATION

1 MOPS pH 7 (mM)	Glucose (g/L)	Magnesium Sulphate (mM)	Ferric Chloride (uM)	ACE METALS (mM)	(1e6X Timmino Acid Supp (mM)	(50X Tim Ammonium Sulphate
55	5	1	100	0.1	0.1	
55	10	1	100	0.1	1	
110	10	5	10	0.1	0.1	
55	10	5	10	1	1	
55	5	1	10	0.1	1	
55	5	1	10	1	0.1	
55	10	5	10	0.1	0.1	
55	5	5	100	1	0.1	
110	10	5	10	0.1	1	
55	10	1	100	0.1	1	
110	5	1	10	1	1	
110	10	1	100	0.1	1	
110	10	1	100	1	0.1	
110	10	5	10	1	0.1	
55	10	1	100	0.1	1	
110	5	1	100	1	0.1	
110	10	1	10	0.1	1	
110	5	1	10	0.1	1	
110	5	1	10	0.1	1	
110	10	1	10	0.1	0.1	
110	10	5	100	0.1	0.1	
55	5	1	10	1	0.1	
110	5	5	10	0.1	0.1	
110	10	5	100	1	1	
55	5	5	10	1	1	
110	5	1	10	1	1	
110	5	1	100	0.1	0.1	
55	10	1	10	0.1	0.1	
110	10	1	100	0.1	1	
55	5	1	100	0.1	1	
55	5	1	100	0.1	1	
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110	10	1	100	0.1	0.1	
55	5	1	100	0.1	1	
55	5	1	100	0.1	1	

abstraction



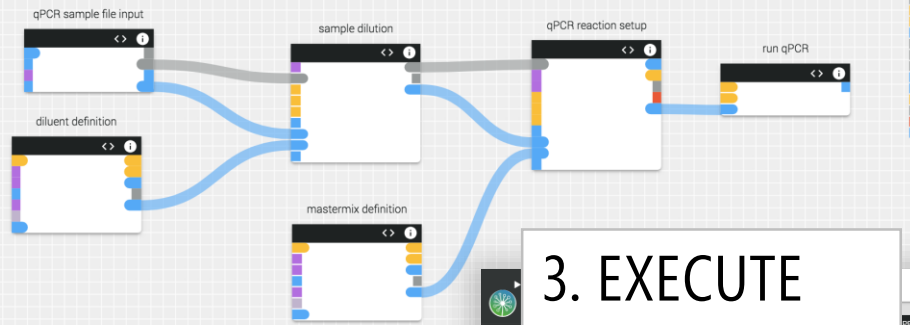
SCIENTIFIC & OPERATIONAL BENEFITS IN DOE WORKFLOWS



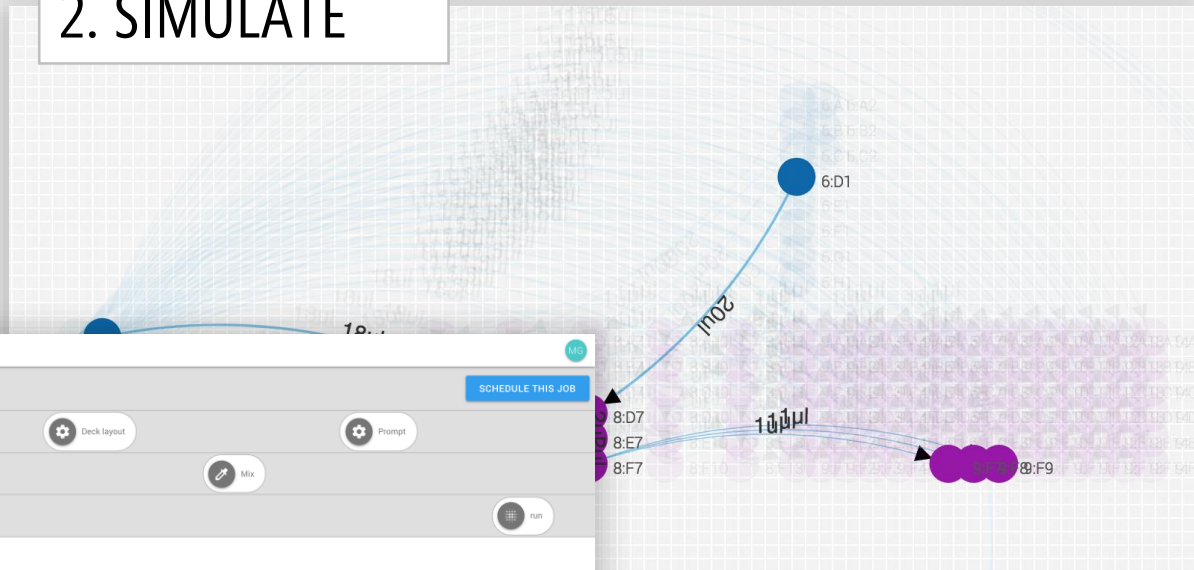
Anthra enabled the optimization of transfection conditions to give 3-10 fold increase in viral titre, whilst providing 83% time and 32% resource savings.

CASE STUDY: QPCR ANALYTICS AUTOMATION

1. DESIGN



2. SIMULATE



3. EXECUTE

QPCR workflow - Markus Gershtator

SEE DESCRIPTION

Estimated time: 01:17:58
Submitted by: Mark Gershtator
Status: **Simulation**

Open in other apps

- Job
- Workflow

Order files

- order.csv
- order.fas

Plate files

- qPCR_OutputPlate.csv.csv
- qPCR_OutputPlate.csv.csv
- auto_input_plate_1_mole.csv
- DilutionPlate1.csv
- auto_input_plate_1_mole.csv
- qPCR_plate1.csv

Data files

- raw.sqlite
- 1_X_qPCR_mastarmix_RecipeSheet.html
- CGTC_QPCR_DesignFile.csv
- qPCR_plate reaction layout.csv
- outputs.json

RESOURCES

Labware	
Plates	
96 well plate with cooler + no riser	1
384 well skirted optical PCR plate (AppliedBiosystems) + 18mm riser (Gilson)	2
96 well plate (skirted) + 18mm riser (Gilson)	1
Tips	
DL10 Tip Rack (PIPETMAX 8x20)	245 (3 boxes)
Tipwaste	
Wastebbox - gilson	1

Materials	
Reaction1	50 ul
Reaction2	50 ul
Reaction4	50 ul
Reaction5	50 ul
Reaction8	50 ul
Standard1	50 ul
NonTemplateControl	100 ul
Reaction3	50 ul
Reaction5	50 ul
Reaction7	50 ul
qPCR mastarmix	1182 ul
EASY Dilution Buffer	403.5 ul

Devices

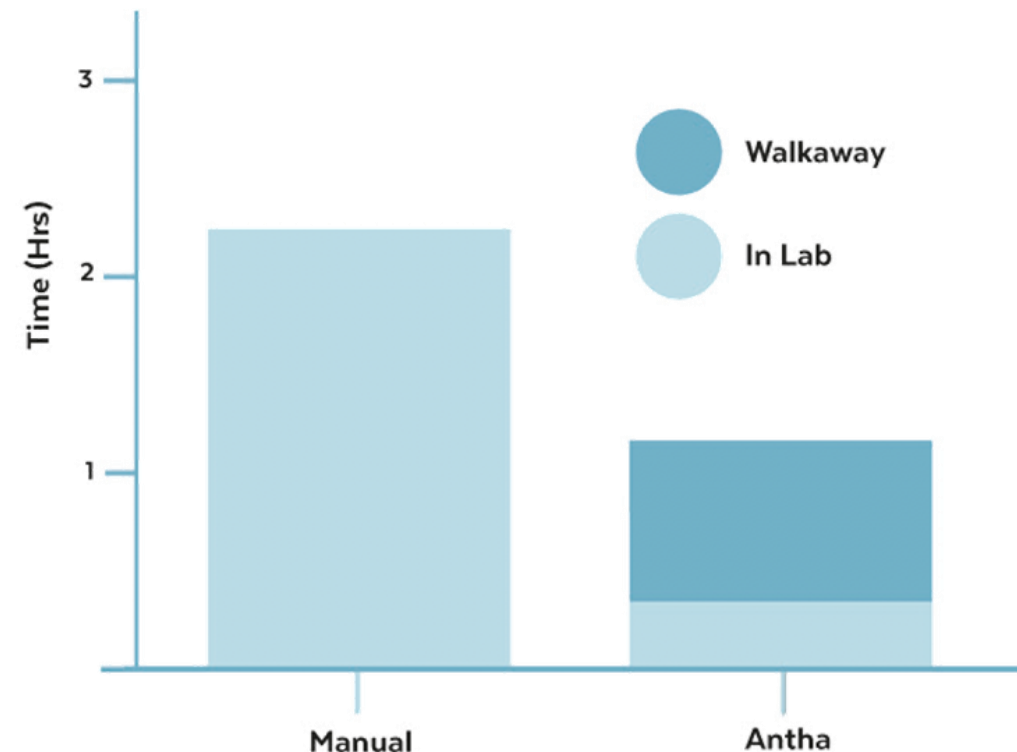
qPCR_MTG_w7vm_0
QPCRDevice
2HVBTVMK0KBAWQ137XPKWKT4
Last seen 03:47 24/05/2018

Arnold
GilsonPipetMax
8NJTOLJNGENR8S48BA67AS6W7
Last seen 09:50 20/04/2018

run

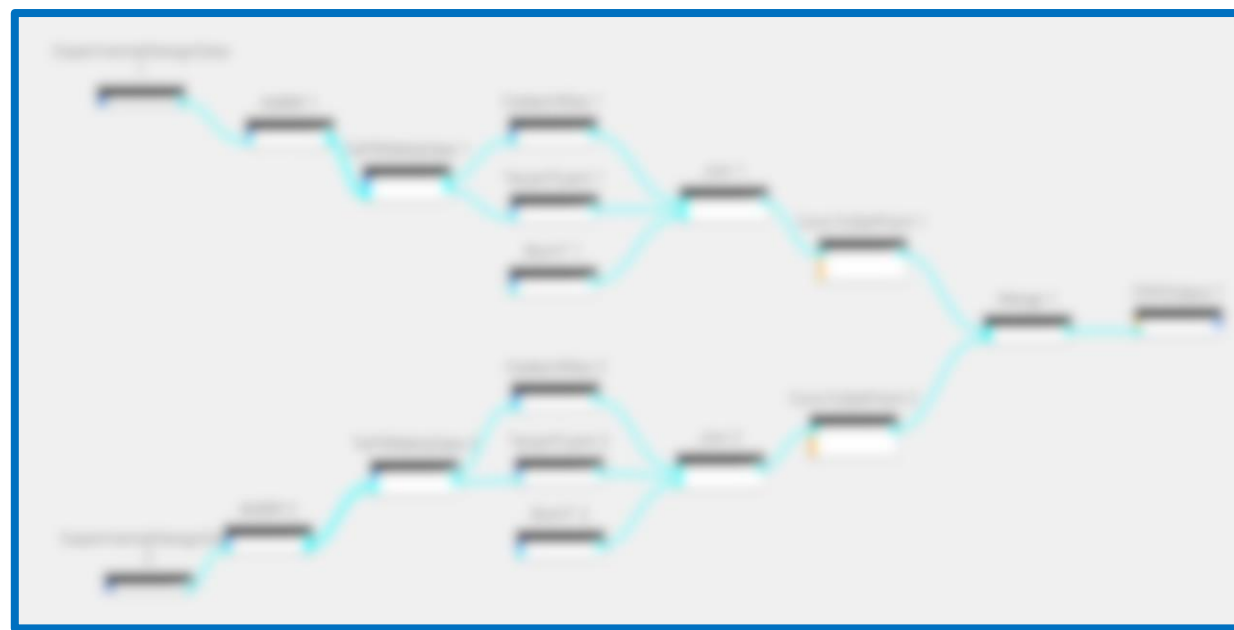
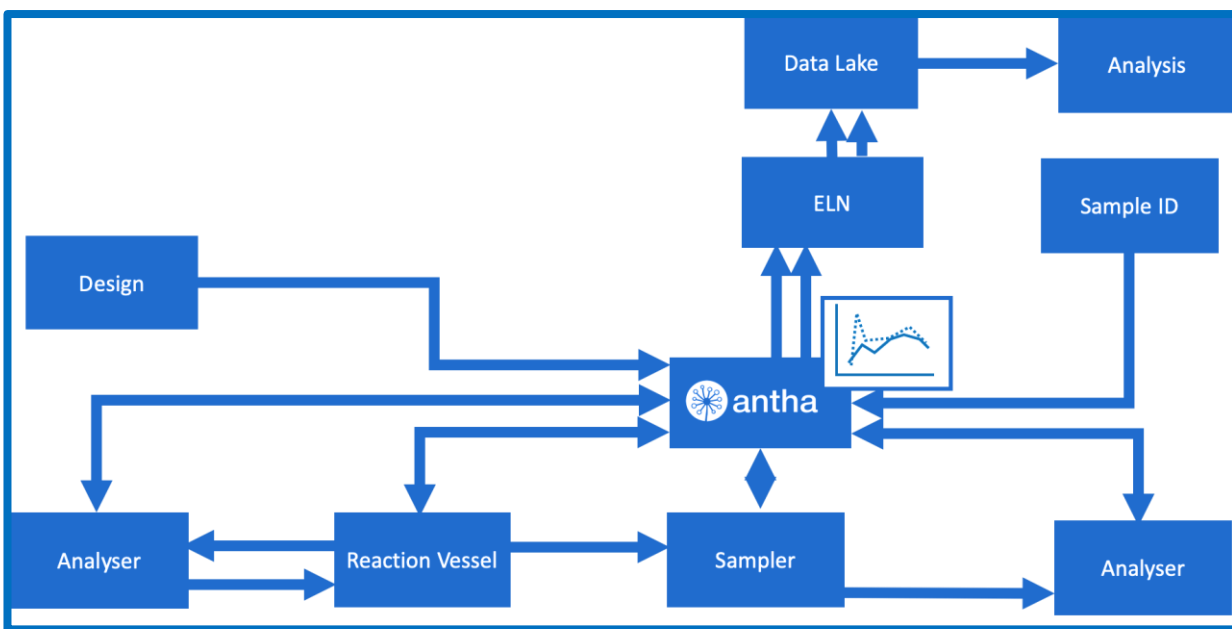
ANALYTICS AUTOMATION EMPOWERS USERS

Set up time per 384 well plate (mins)				
Workflow Step	Manual Process		Antha Automated Process	
	In Lab	Walk Away	In Lab	Walk Away
qPCR master mix preparation	10	-	10	-
Set up robotdeck	-	-	5	-
Robot tip refill	-	-	5	-
Liquid Handling	120	-	-	45
Set up qPCR thermocycling	5	-	5	-
qPCR thermocycling	-	50	-	50
Total Time	135	50	25	95
Total Time to Data Output	185		120	



This workflow is now scalable to allow extensive testing in realistic timelines, increasing throughput by 50%. The resulting data generated is highly reliable and reproducible whilst improving cross contamination control.

FROM UNIT OPERATIONS TO END TO END MODULAR AUTOMATION





Conclusions..

AN ECOSYSTEM OF PLAYERS.... COLLABORATION IS KEY



DIGITAL
Powered by AI



PHYSICAL
Powered by
Automation

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Bio Design

Bio Simulation

Experiment Design

Experiment Simulation

Experiment Execution

Building Blocks

Analysis

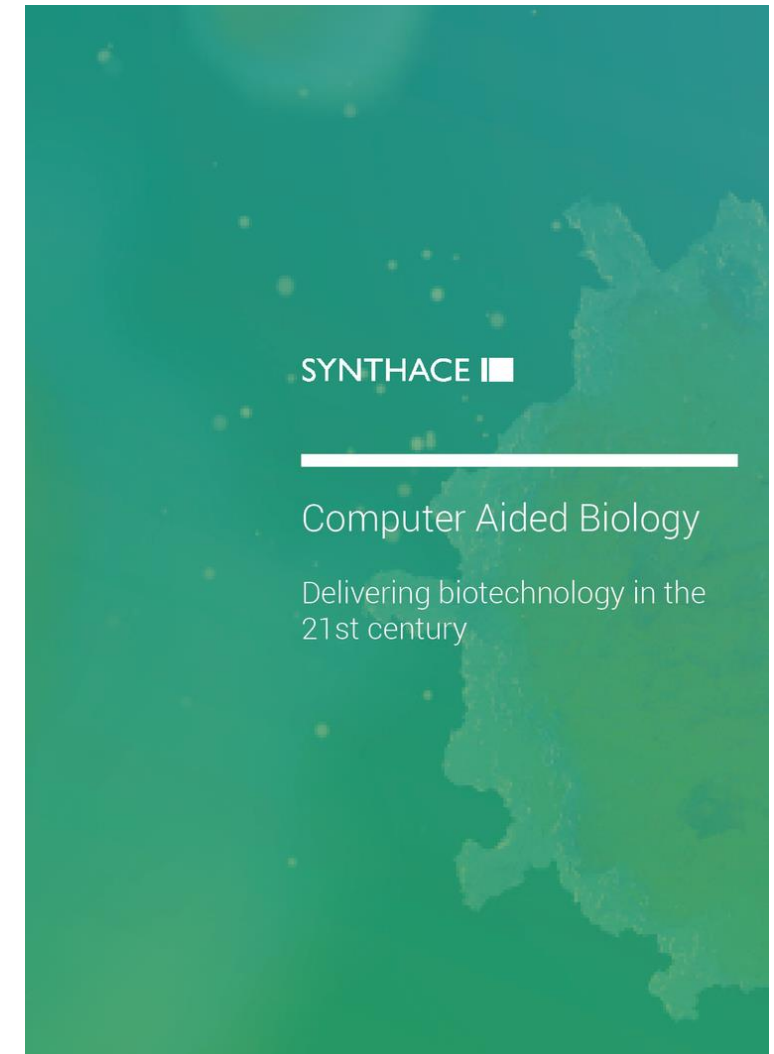
Data Management

WE WORK WITH PARTNERS ACROSS THE LIFE-SCIENCES ECOSYSTEM



WE ARE LOOKING FORWARD TO WORKING WITH YOU ALL..

- **Focus on Digital to Physical Transition**
- White Paper on CAB >>> see website
- Office Hours/Drinks events in our London Office/LOTF (Biology, Bytes and Beers)
- US Office Q2 2019 (Boston)
- Learning more about standards & networks to see how we can add value to community (Allotrope and Pistoia Members)
- Rapidly expanding internal resource so open to collaboration across ecosystem





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