

# Modelling in immuno-oncology

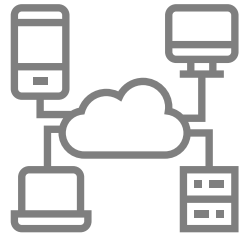
*Biotrinity, April 2020*



# Physiomics brings together a unique combination of capabilities...



Knowledge of  
**cancer biology**  
from industry and  
academia



Understanding of  
pre-clinical and  
clinical **data** sourcing  
and curation



Expertise in  
**quantitative  
pharmacology**



Experience in  
**modelling and AI**  
techniques such as  
machine learning



## ...to answer key development questions for our clients

---

Recommend **efficacy/ toxicity** trade-offs



Predict **biologically effective dose in humans** to support clinical translation



Recommend **combination partner agent** for proprietary in-house asset





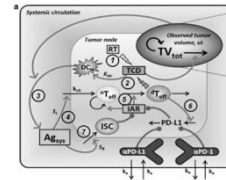
# Physiomics is a consultancy focused on quantitative modelling in oncology

## PK/PD analysis



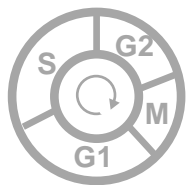
- Data analysis and interpretation
- PK/PD modelling (NCA, PopPK)
- In-vitro, in-vivo, clinical

## Quantitative Systems Pharmacology



- Pathway models
- PBPK
- Literature, bespoke or hybrid

## Virtual Tumour™



- In-silico cell-cycle model of tumour growth
- Predicts tumour regression
- Mono or combination therapy

## Personalised medicine



- Grant funded
- Predict response to treatment for individuals



# We've completed over 70 commercial and grant funded projects

---

**Big pharma**  
Merck, MSD, Lilly,  
Bayer and others

**Grants**  
Innovate UK, EU-FP6  
(TEMPO), Carbon Trust



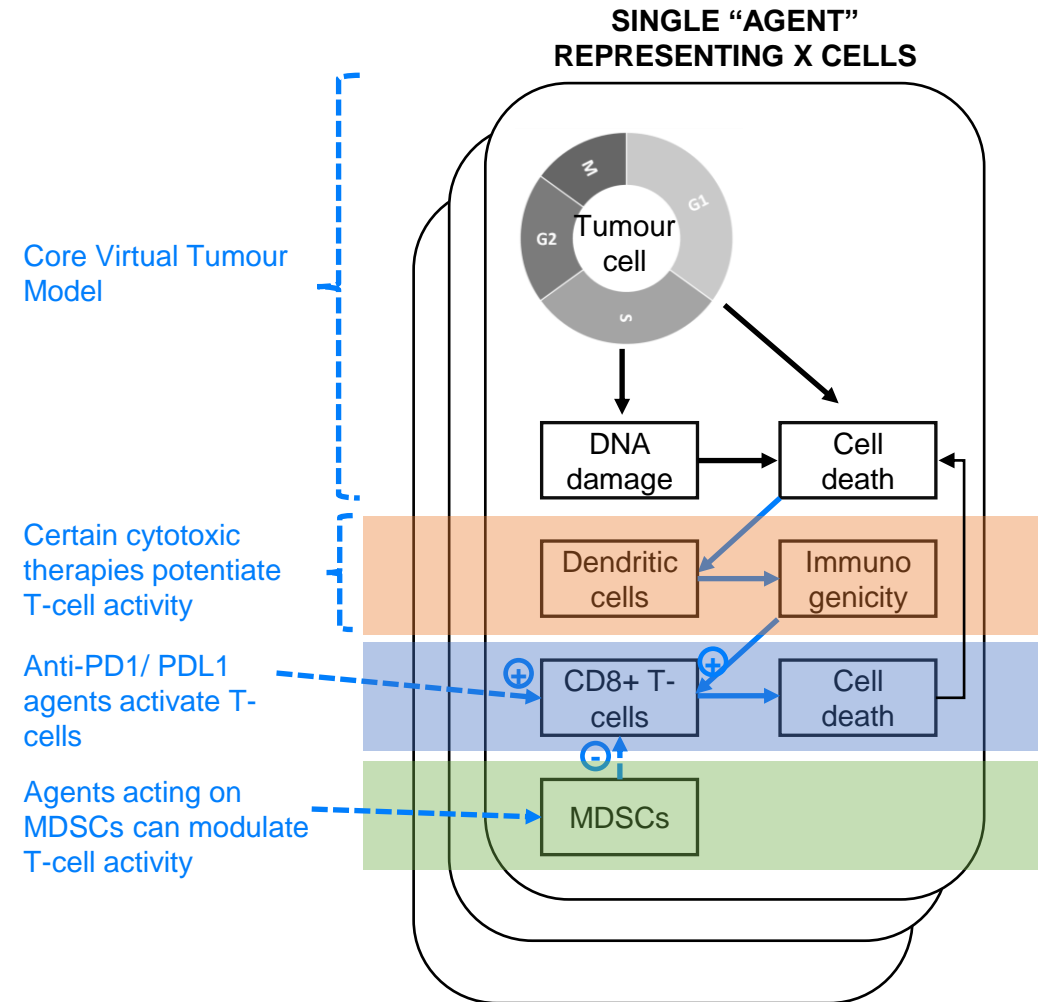
**Biotech/ Midcap**  
Bicycle Therapeutics, CRUK  
Cellcentric, Convert Pharma,  
and others

**Academic**  
Medicines Discovery Catapult, NIH, Oxford University,  
Oxford AHSN, Institute of Cancer Research, CRT



# Our Virtual Tumour™ platform has been extended to enable modelling of immuno-oncology combination treatment

- Various abstracted immune pathways are incorporated:
  - Stimulatory effect of anti-PD-(L)1 on T-cell activation levels, leading to tumour cell death
  - Potentiation by some cytotoxic agents via induction of immunogenic cell death and recruitment/maturation of dendritic cells
  - Inhibitory effect of MDSCs on T-cell activity (allowing consideration of drugs that act on this cell type)
- This enables flexible simulation of combinations of anti-PD(L)1, DDR agents, RT, immunogenic cytotoxics, agents acting on MDSCs and more

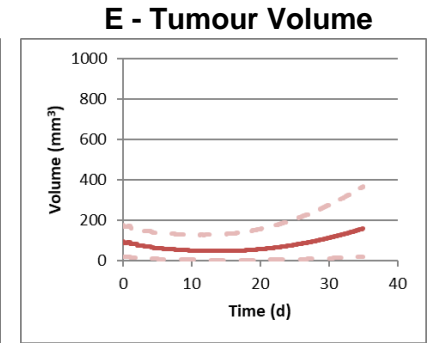
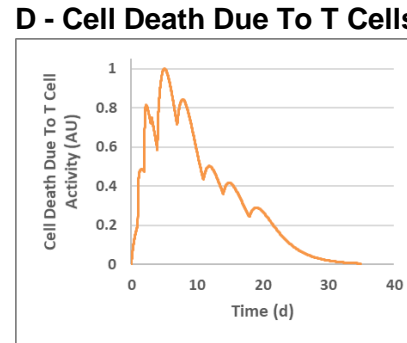
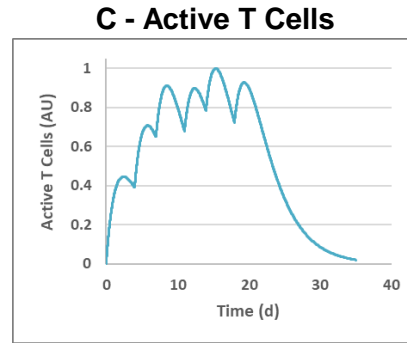
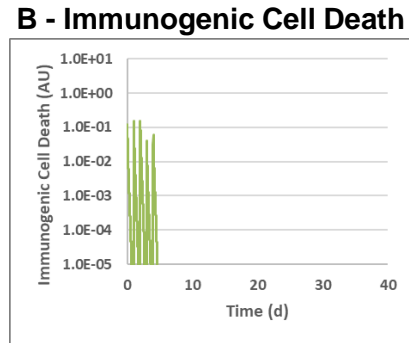
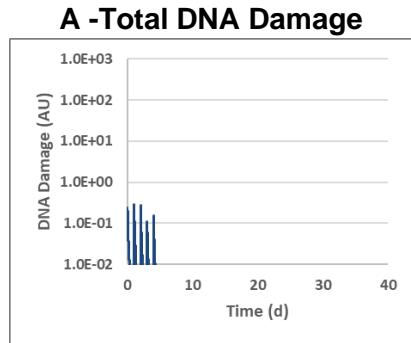




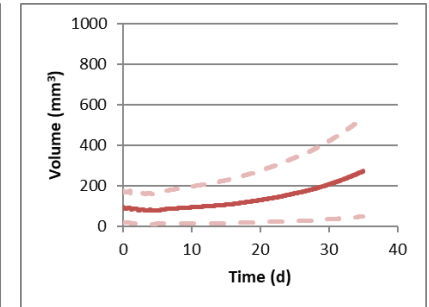
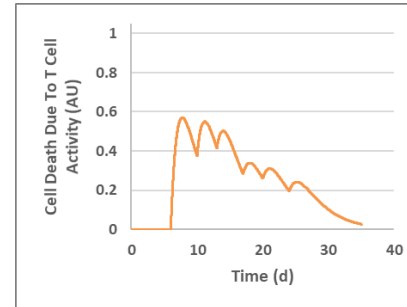
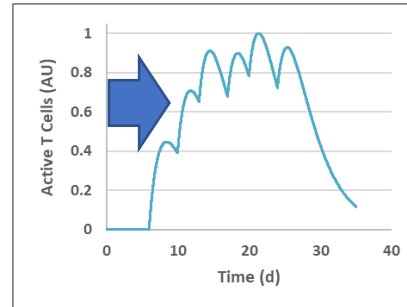
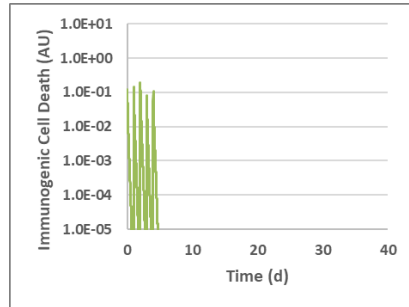
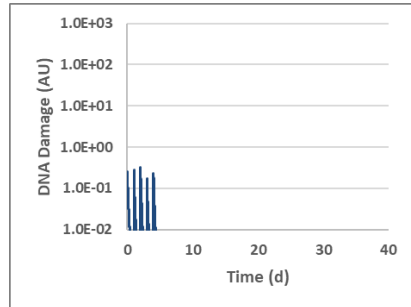
# This enables us to predict and explain timing effects in terms of species modelled – for example, RT + anti-PD-L1

Offset of first dose of anti-PD-L1

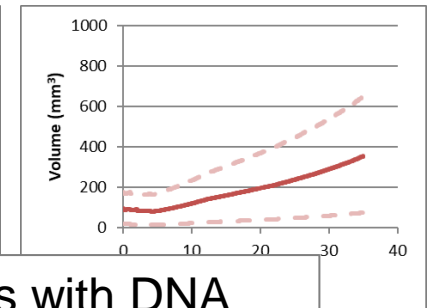
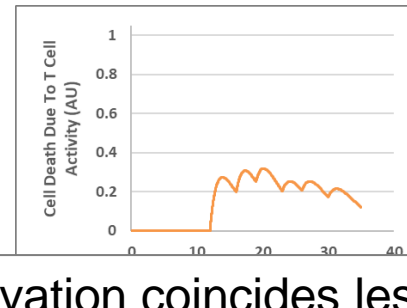
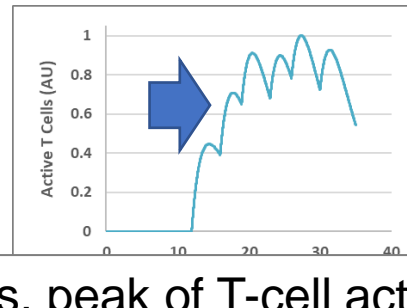
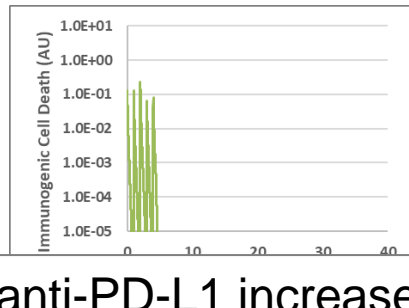
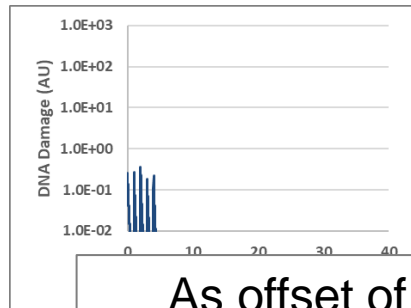
0 d



6 d



12 d



As offset of anti-PD-L1 increases, peak of T-cell activation coincides less with DNA damage, reducing opportunity for potentiation

Decreasing T-cell induced cell death and tumour regression



# Why work with Physiomics?

---

- We help solve real development challenges
- We supplement existing client resources to increase capacity and shorten timelines
- Exclusively focused on cancer
- Deep expertise
  - >70 big pharma, biotech and grant funded projects
  - Significant academic as well as industry experience
- Can bring proprietary (Virtual Tumour™) and other industry standard models to bear on R&D challenges
  - Experience with large number of targets, cell lines, PDX
- We are a small team that offers a dedicated, flexible service



