



Microfluidics in organoid systems

Juli R Bago

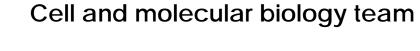


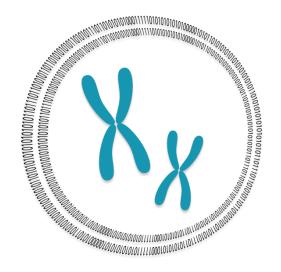




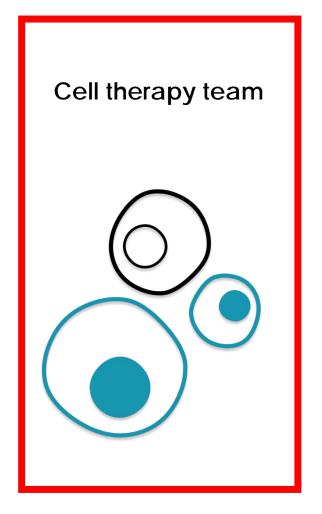
BCRG

Genomic and bioinformatics team















Cell based-immunotherapy: CAR-T cells









Cell based-immunotherapy: NK cells



CART-CELL THERAPY

Expensive=400K \$. Autologous to avoid GvHD

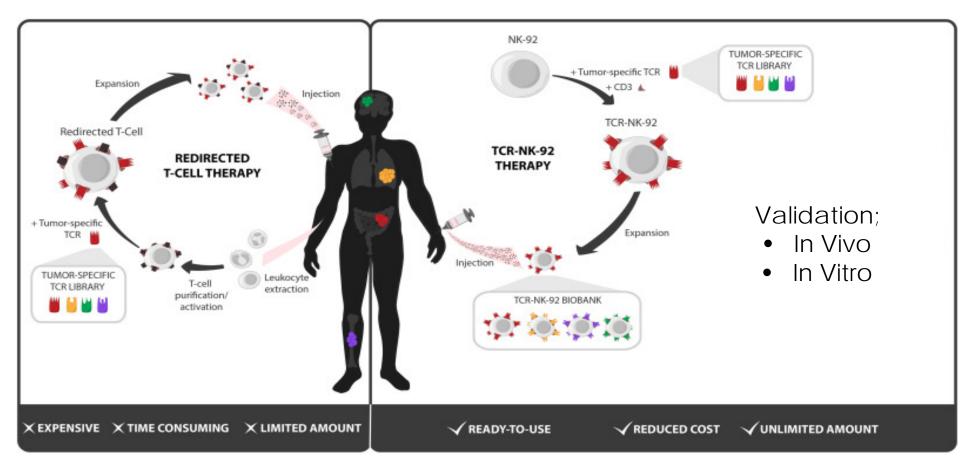
CART cells exert immune reaction towards a specific tumor antigen.







Cell based-immunotherapy: NK cells



Mensali N, et al. NK cells specifically TCR-dressed to kill cancer cells. 2019. EBioMedicine







Cell based-immunotherapy: Validation

IN VIVO



Mouse models of hematological cancer

- Ethical issue
- Differences in physiology and genetics between animals of experimentation and humans.







Cell based-immunotherapy: Validation

IN VITRO

| Culture model | Advantages | Disadvantages |
|-----------------|--|--|
| 2D cell culture | Methodology well established | Static conditions |
| | Simplicity to work with cell monolayer | Uniform concentration of nutrients and |
| | | drugs |
| | | Lack of 3D environment |
| | | Large reagent volumes |

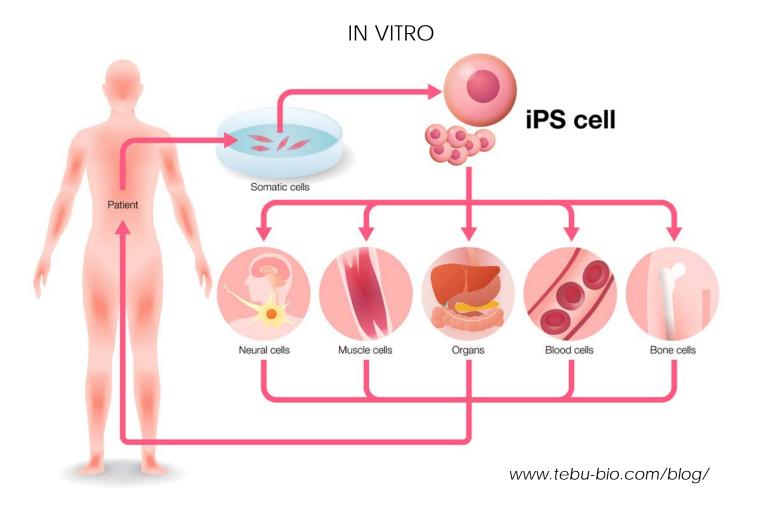


Valente et al. Microfluidic technologies for anticancer drug studies. 2017. Drug discovery today.





Cell based-immunotherapy: Validation







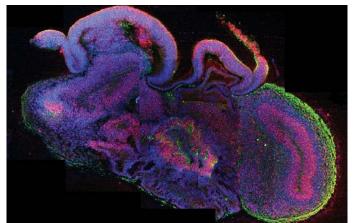


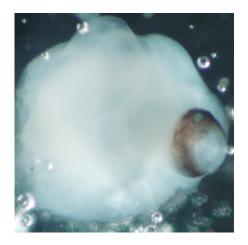
Cell based-immunotherapy: Validation

IN VITRO

MiniBrains







Lancaster, M. A. et al. 2013. Nature.







Cell based-immunotherapy: Validation

IN VITRO



Cell Applications inc

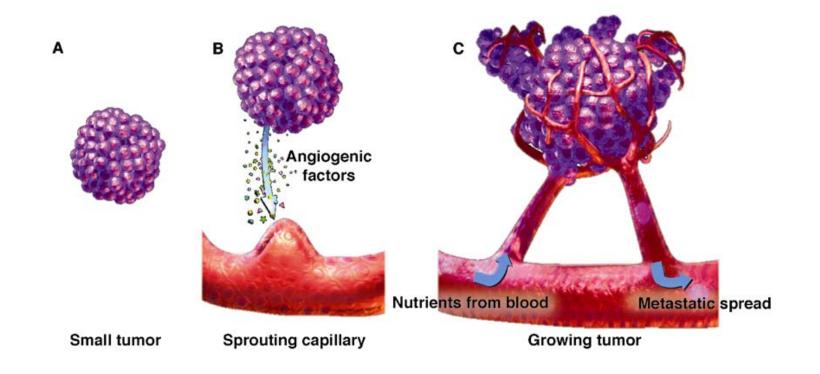






Cell based-immunotherapy: Validation

IN VITRO Vascularization









Cell based-immunotherapy: Validation

IN VITRO

| Advantages | Disadvantages |
|--|--|
| Methodology well established | Static conditions |
| Simplicity to work with cell monolayer | Uniform concentration of nutrients and |
| | drugs |
| | Lack of 3D environment |
| | Large reagent volumes |
| Cell-cell and cell-ECM interactions | Failure to produce dynamic environment |
| Sensitivity to cytotoxic agents similar to in vivo | Lack of fluid flow perfusion |
| | Methodology well established Simplicity to work with cell monolayer Cell-cell and cell-ECM interactions |

Valente et al. Microfluidic technologies for anticancer drug studies. 2017. Drug discovery today.





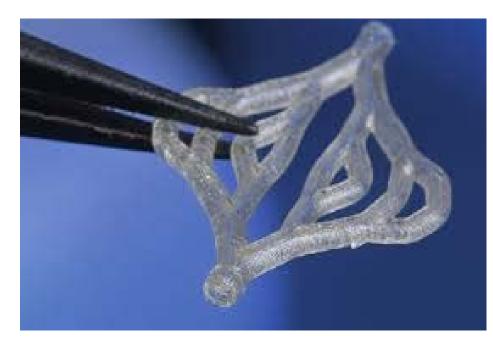


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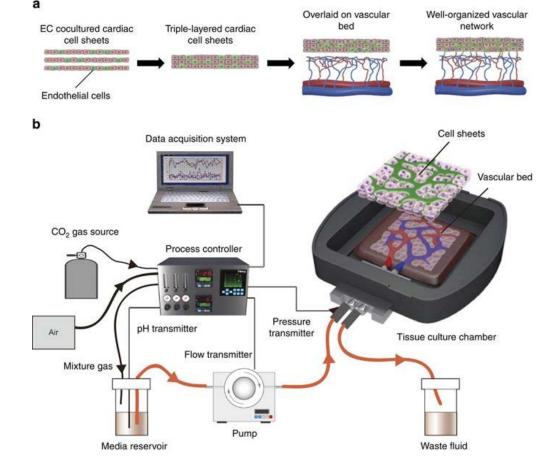
CELL THERAPY TEAM

Cell based-immunotherapy: Validation

IN VITRO Vascularization



Sekine et al. In vitro fabrication of functional three-dimensional tissues with perfusable blood vessels. 2013. Nat Commun.



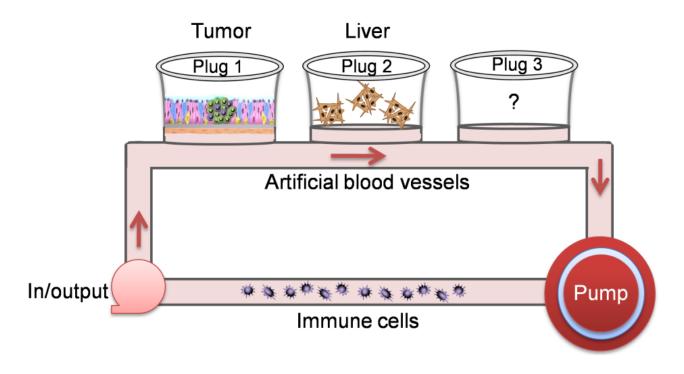






Cell based-immunotherapy: Validation

IN VITRO



Samuel Constant et al. Advanced Human In vitro Models for the Discovery and Development of Lung Cancer Therapies. 2015.







THANK YOU SO MUCH



