



CREATING
A CIRCULAR
FUTURE

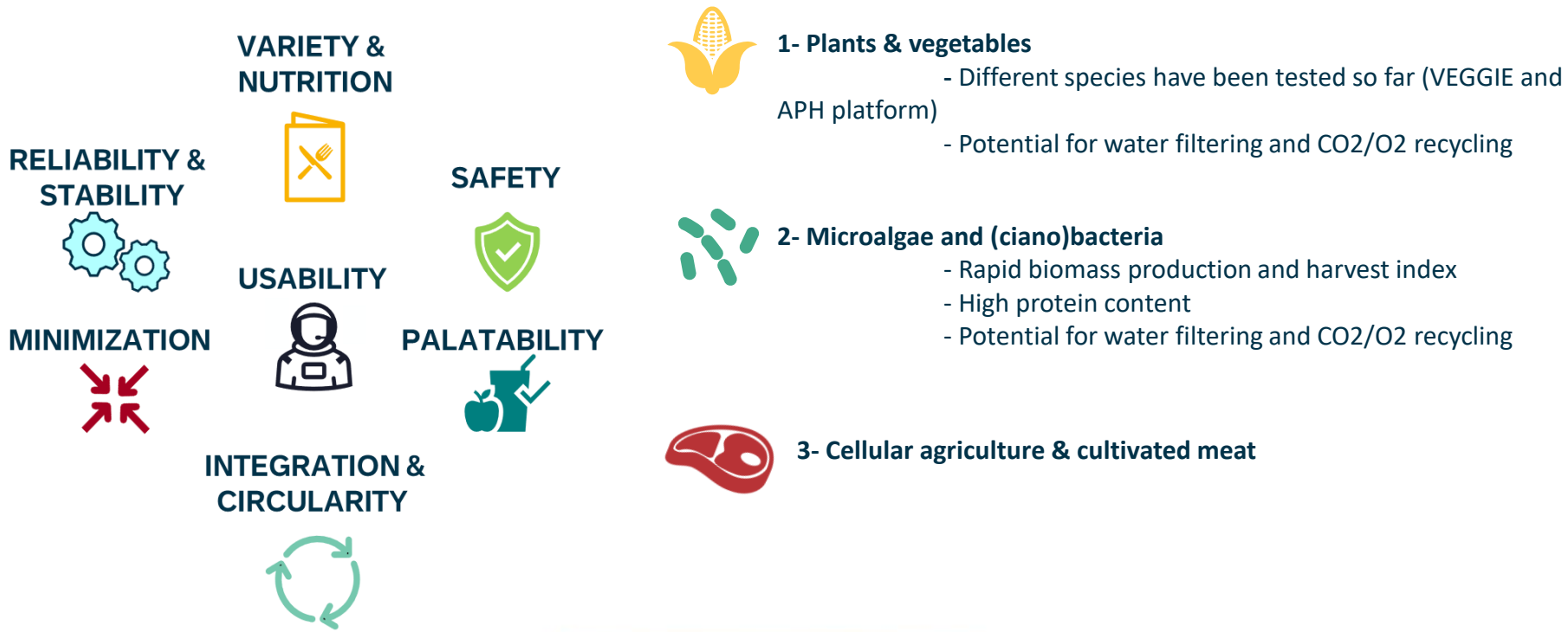
Cultivated meat for human space exploration

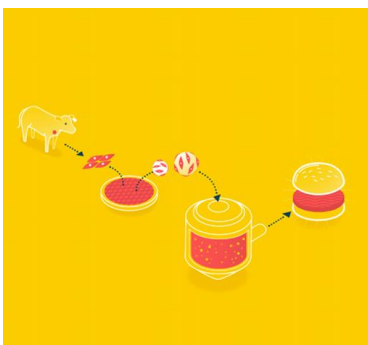
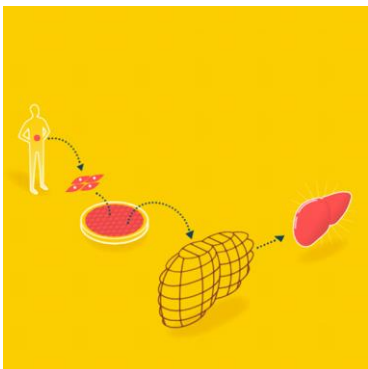


2022 MELISSA Conference
Toulouse, France

João Garcia
Research fellow
European Space Agency

MELISSA Future space food systems



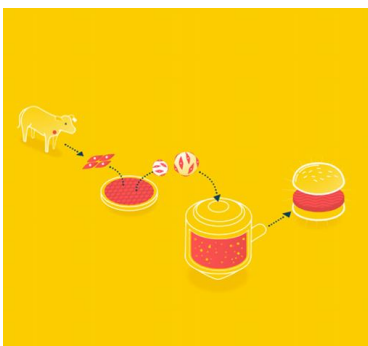
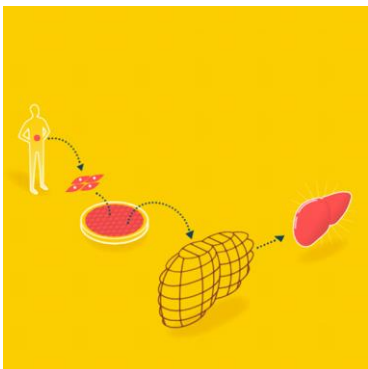


Cellular agriculture describes the process of using cells to produce commodities (meat, leather, milk)

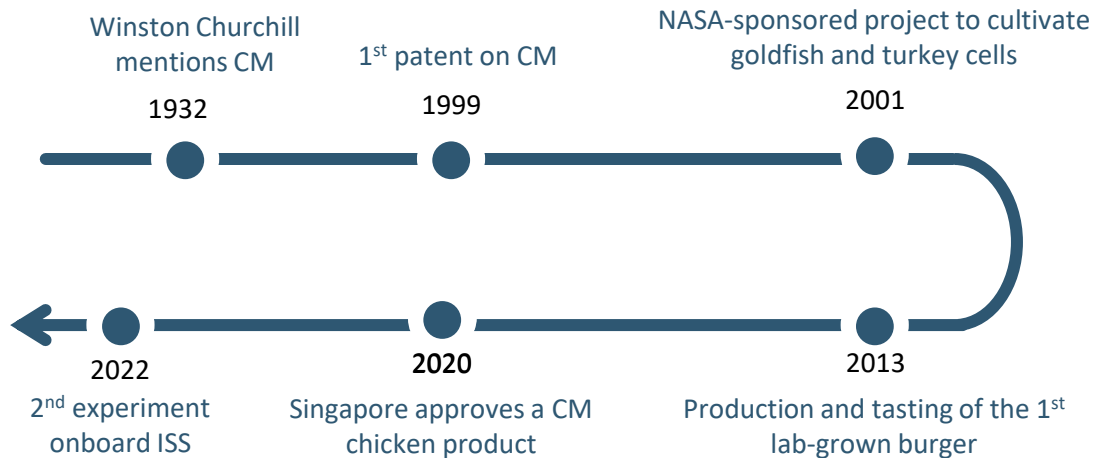
Cultured meat describes the process of growing animal tissues (i.e. muscle, fat) using animal cells:

- Core principles derived from **tissue engineering** and **regenerative medicine**
- It emerged as an alternative to conventional (and **unsustainable**) livestock farming
- Deemed to be **safer**, as there is total control of the production processes, reducing the likelihood of contamination and **foodborne illnesses**
- Requires less **land**, **energy** and **water** when compared to livestock farming

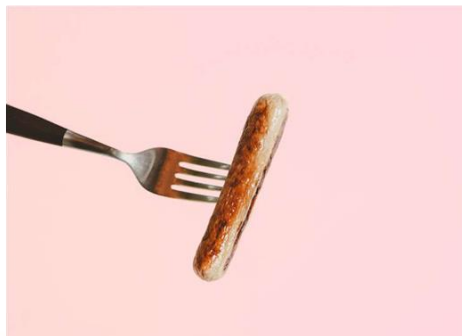
MELISSA Cellular agriculture & cultivated meat



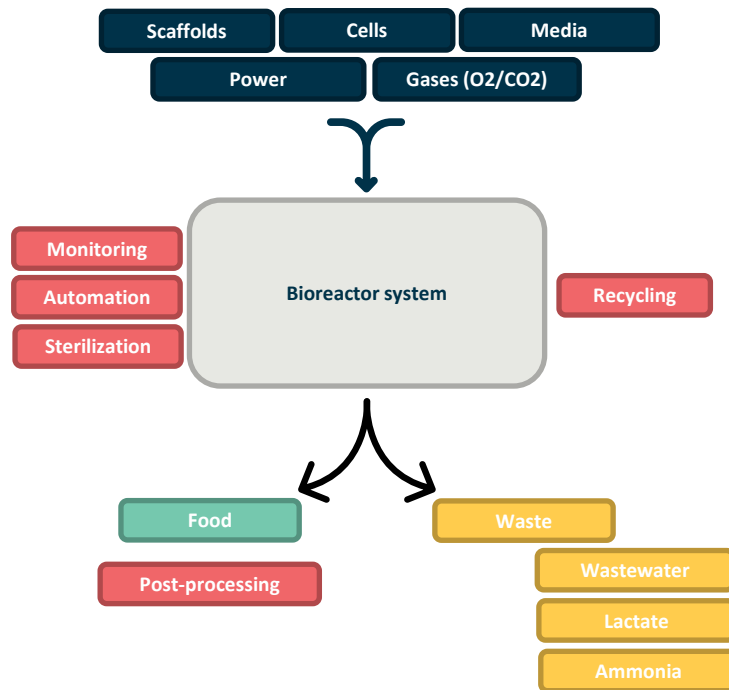
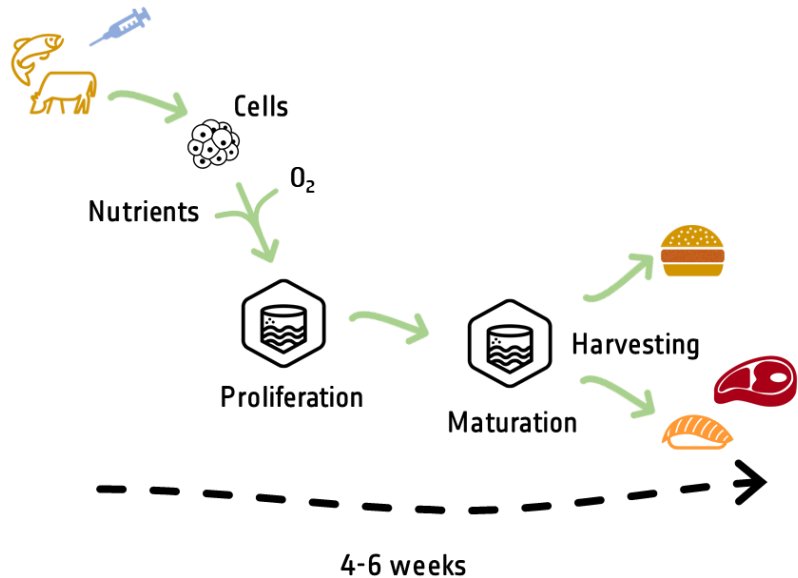
Cellular agriculture describes the process of using cells to produce commodities (meat, leather, milk)



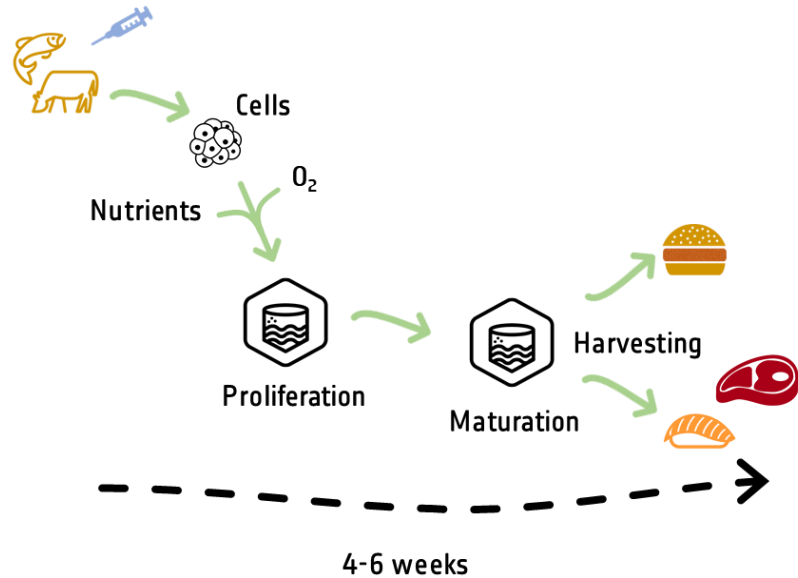
MELISSA Several cultivated meat products are in development



MELISSA Cultivated meat



MELISSA Cultivated meat - From Earth to Space



Space & habitat environment:
1) **Altered gravity**
2) Deep space radiation
3) **Volume constrains**

System:
1) Automation
2) Reliability & stability

Bioprocess:
1) **Resource minimization**
2) **Waste management and recycling**
3) Nutritional profile and palatability



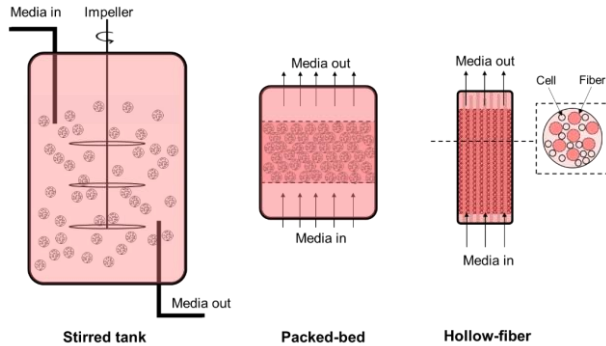
MELISSA Sources and cell types

Parameter	Mammalian & avian cells	Fish & crustacean cells
Temperature (°C)	37	10-30
Oxygen (pO ₂)	1-5% or 21%	<1-21%
Carbon dioxide (CO ₂)	5 %	5 %
pH	7.4	7.4, possibly resisting higher variations

Water species might be more convenient as they are more resistant to variation in environment conditions and can be cultured at lower temperatures

	Cell type	
	Muscle-resident cells/progenitor cells	Pluripotent stem cells
Examples	<ul style="list-style-type: none"> Satellite cells Fibro-adipogenic progenitor cells Mesenchymal stem cells 	<ul style="list-style-type: none"> Induced pluripotent stem cells (iPSCs) Embryonic stem cells (ESCs)
Cons	<ul style="list-style-type: none"> Limited proliferation capacity Long doubling times (>30h) 	<ul style="list-style-type: none"> Difficult to (re)program cells Might require genetic engineering strategies Ethical concerns (ESCs) Can be difficult to obtain
Pros	<ul style="list-style-type: none"> Easy to isolate Differentiation protocols are faster and easier 	<ul style="list-style-type: none"> “Unlimited” proliferation capacity Can differentiate into multiple tissues Short doubling times (<24h)

MELISSA Bioreactors

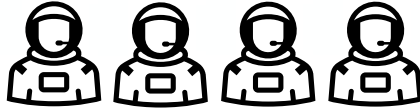


Characteristic	Bioreactor		
	STR	PBB	HFB
Max cell density (cells/ml)	$1 \times 10^5 - 1 \times 10^7$	$1 \times 10^6 - 1 \times 10^7$	$1 \times 10^7 - 1 \times 10^8$
Operation in microgravity	Possible, but mixing of media and cells will need to be optimized	Suitable, since cells are attached to a substrate and media is perfused	Suitable, since cells are attached to a substrate and media is perfused
Reusability	Yes	Yes, but further development needed (edible microcarriers or efficient dissociation methods)	Yes, but further development needed (edible fibers or efficient dissociation methods)
Major limitations	<ul style="list-style-type: none"> • Shear stress on cells • Control aggregate formation/size • Not suitable for differentiation stage 	<ul style="list-style-type: none"> • Scalability • Mass transfer • Cell harvesting • Reusability 	<ul style="list-style-type: none"> • Cell harvesting • Reusability

MELISSA The Mars example



1200 days



4 CM (4x82 kg)

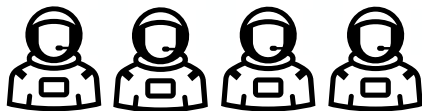


59.04 g protein/CM/day

MELISSA The Mars example



1200 days



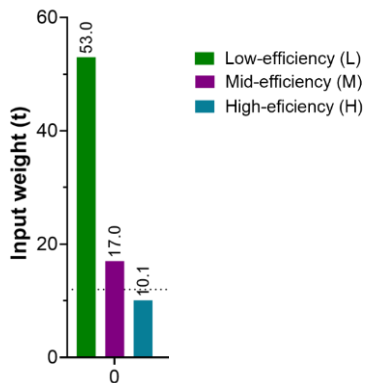
4 CM (4x82 kg)



59.04 g protein/CM/day

	kg media per kg cultivated meat
Low-efficiency	41
Mid-efficiency	23
High-efficiency	6

Roughly 1300 kg of cultivated meat over a period of 1200 days

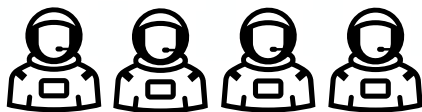


> 97% of media is water

MELISSA The Mars example - Recycling will be crucial



1200 days



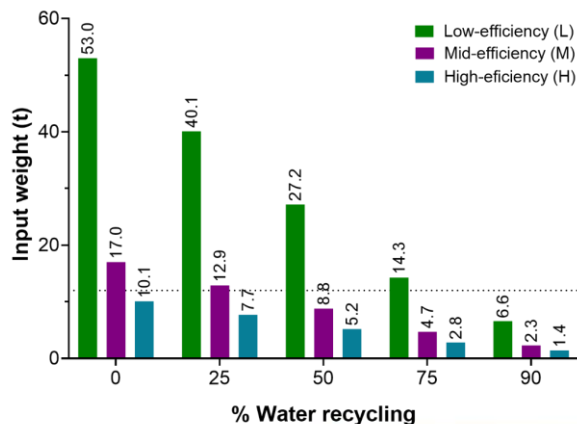
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- Wastewater
- Lactate
- Ammonia



> 97% of media is water

Possibility for integration with water reclamation systems with recovering efficiencies >75%

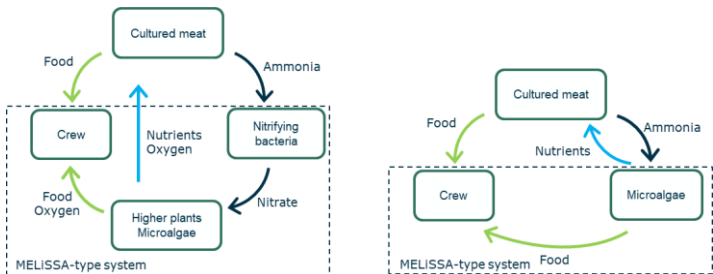
MELISSA The Mars example - Recycling will be crucial

Wastewater

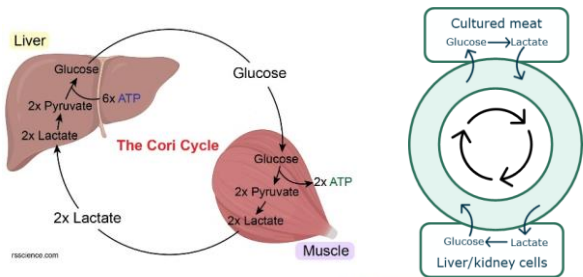
Lactate

Ammonia

Bacteria and microalgae can metabolize ammonia



Certain mammalian cell types convert lactate into glucose



Journal of Biotechnology (2022) 208:673
<https://doi.org/10.1007/s10291-021-01224-9>

ORIGINAL PAPER

A circular cell culture system using microalgae and mammalian myoblasts for the production of sustainable cultured meat

Yuji Haseguchi¹, Yuta Okamoto^{1,2}, Tatsuya Shimizu¹



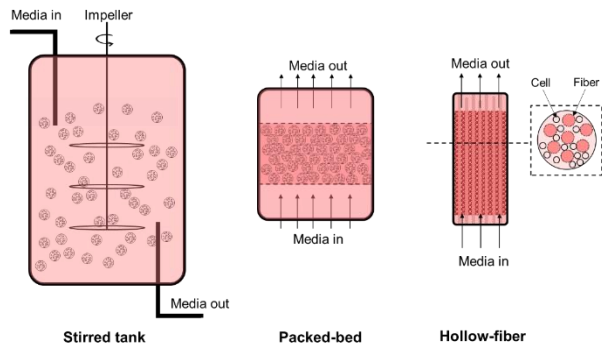
RESEARCH ARTICLE

Proliferation and differentiation of primary bovine myoblasts using *Chlorella vulgaris* extract for sustainable production of cultured meat

Yuta Okamoto, Yuji Haseguchi, Akira Yoshida, Hiromitsu Takahashi, Kunito Yamahata, Naoya Sawamura, Toru Aoki, Tatsuya Shimizu



MELISSA The Mars example – bioreactor sizing



Assuming a batch production cycle of 4 weeks:
 34 kg meat $\approx 1 \times 10^{13}$ cells

	STR	PBR	HFB	Units
Max cell density bioreactor	1×10^7	1×10^7	1×10^8	cells/ml
Internal volume	0.34	0.34	0.034 (0.068)*	m^3

*in HFBs fibers account for 50% of the total volume

Bioreactors that achieve higher cell densities will likely be preferred due to volume and mass limitations

MELISSA Other considerations



Scaffolding

(edible or reusable)



Sensing and monitoring capabilities

(biochemical, chemical and physical parameters)



Sterilization methods

(heat, radiation or chemical methods)



Post-processing

(Product formulation, storage, cooking)

MELISSA Microgravity leads to muscle loss

J Physiol 588.18 (2010) pp 3567–3592

3567

Prolonged space flight-induced alterations in the structure and function of human skeletal muscle fibres

R. H. Fitts¹, S. W. Trappe², D. L. Costill², P. M. Gallagher², A. C. Creer², P. A. Colloton¹, J. R. Peters¹, J. G. Romatowski¹, J. L. Bain³ and D. A. Riley³

Space travel directly induces skeletal muscle atrophy

HERMAN VANDENBURGH^{a,*1}, JOSEPH CHROMIAK^a, JANET SHANSKY^a, MICHAEL DEL TATTO^a AND JULIE LEMAIRE^a

^aDepartment of Pathology, Brown University School of Medicine and The Miriam Hospital, Providence, Rhode Island 02906, USA; and ^bDepartment of Molecular Pharmacology, Physiology, and Biotechnology, Brown University, Providence, RI 02912, USA

Clinorotation prevents differentiation of rat myoblastic L6 cells in association with reduced NF- κ B signaling

Katsuya Hirasaka^a, Takeshi Nikawa^{a,*}, Louis Yuge^b, Ibuki Ishihara^a, Akira Higashibata^a, Noriaki Ishioka^a, Atsuko Okubo^b, Takashi Miyashita^b, Naoto Suzue^d, Takayuki Ogawa^d, Motoko Oarada^c, Kyoichi Kishi^a

npj | Microgravity

www.nature.com/npjgrav

Corrected: Author correction

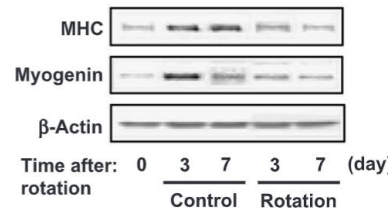
ARTICLE OPEN

Simulated microgravity attenuates myogenic differentiation via epigenetic regulations

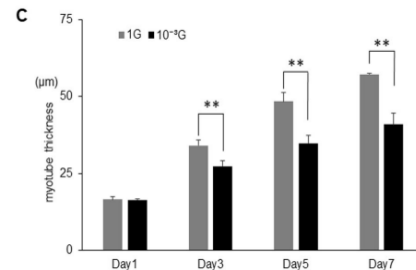
Takuma Furukawa¹, Keiji Tanimoto², Takahiro Fukazawa¹, Takeshi Imura¹, Yumi Kawahara^a and Louis Yuge^{1,4}

Countermeasure strategies are likely necessary for stimulating cells into forming muscle tissue

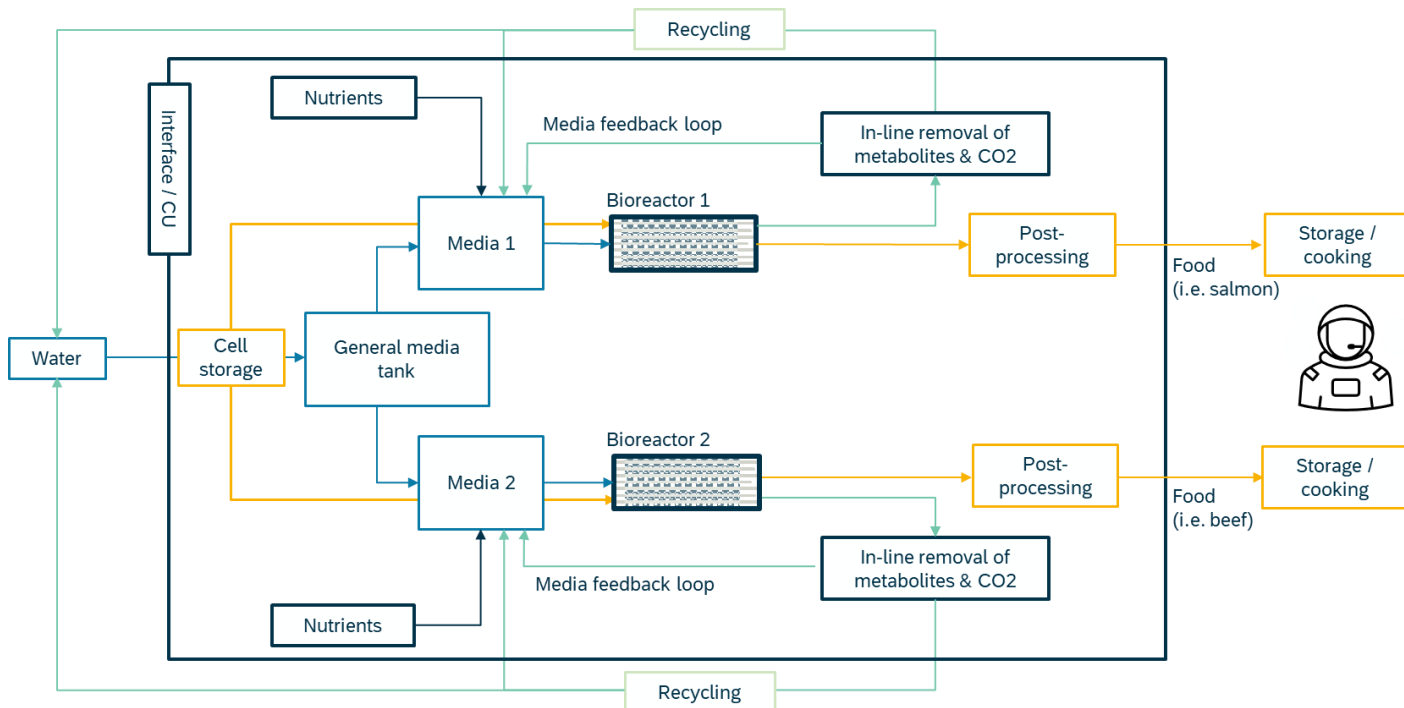
Reduced myogenin and myosin heavy chain production in *simulated* microgravity



Lower expression of myogenic markers and reduced myotubule thickness



MELISSA Concept



- 1) Studies on how cultured meat cells behave in space environment conditions
 - 1) Altered gravity
 - 2) Radiation

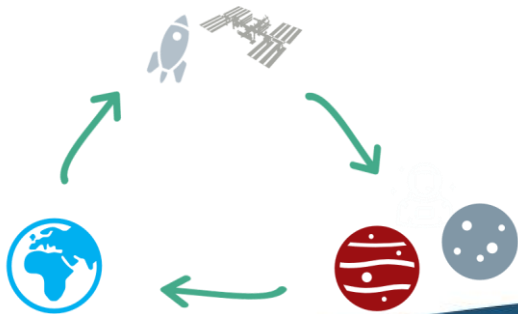


Potential countermeasure strategies to eliminate the effects of microgravity and radiation

- 2) Filtering and recycling strategies
 - 1) Filtering of wastewater, lactate and ammonia
 - 2) Possibility for integration with other life support systems

MELISSA Take home message

- 1) The current space food system is **not suitable** for future long-term & long-distance exploration missions
- 2) **Cultured meat** has the potential to deliver **highly nutritious** and **fresh** food products
- 3) Limitations include **launch volume** and **mass**, **recycling strategies**, and **cell behaviour in microgravity**
 - 1) Bioreactors that achieve higher cell densities will likely be preferred due to volume and mass limitations
 - 2) Recycling strategies (namely water) will be crucial for making cultured meat a viable food source in future human spaceflight
 - 3) Integration with existing life support systems will render the cultured meat systems viable for space exploration



Development and implementation of cultured meat systems for space will improve ground systems, and potentially contribute for a more secure and safe global food system.

MELISSA



MICRO-ECOLOGICAL
LIFE SUPPORT SYSTEM
ALTERNATIVE

What is cultured
meat?



Life cycle analysis of
cultured meat



ESA's presentation @ CMS
Astro



ESA's article on cultured
meat



THANK YOU.

João Garcia

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