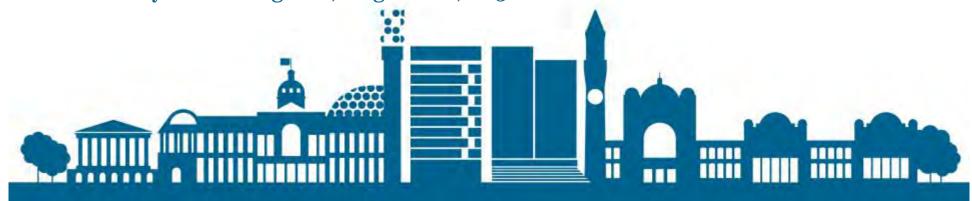


Formulation and Processing: Manufacturability

Peter Fryer

Centre for Formulation Engineering, Chemical Engineering University of Birmingham, Edgbaston, B₁₅ ₂TT



What is formulation?

- Chemical composition the same
- Microstructure controls delivery
- Common across industry sectors





Matrix of skills and industry sectors

	Food, Health and Nutrition	Bioindustries	Home and Personal Care, Pharma	Energy and Chemical Industries
Structured Liquids	Gels and creams	Fermentation broths	Shampoos and creams	Supercritical fluid techniques, paints
Soft Solids	Spreads, emulsions and confectionery	Soft tissue engineering	Crystals, creams and foams	Paints and coatings
Structured Solids	Breads and starch products	Hard tissue engineering	Tablets and capsules	Fuel cells and catalysts: control of interface and structure
Particulate materials	Grains	Separation and delivery systems	Granules and powders	Fuel cells and catalysts

Formulation CDT



























Processing and formulation

- First understand the sequence of events in processes
- □ To build new processes that are inherently more efficient – what are the boundary conditions set by the physics, the biology, and the consumer?
- □ Do you understand the process? The equipment? The usage?



Manufacturability

- First understand the sequence of events in processes
- □ Can you scale it up? No: really.
- □ Do you understand the process?
- □ Can you make it
 - Efficiently
 - To cost
 - Sustainably



Can you scale it up?



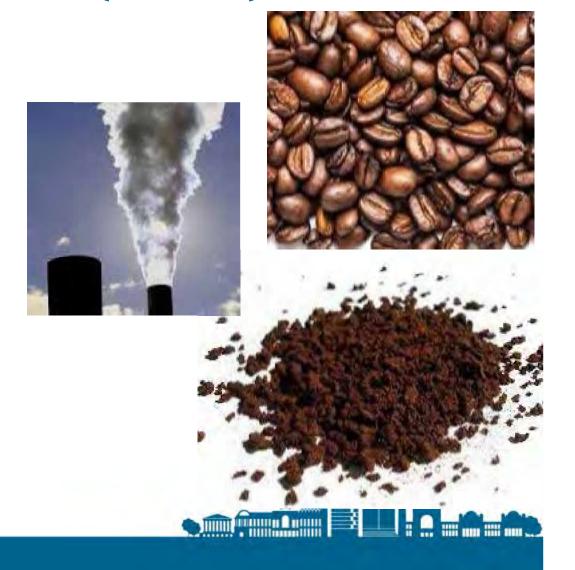
Constraints:

- Microbial
- Quality
- Cost



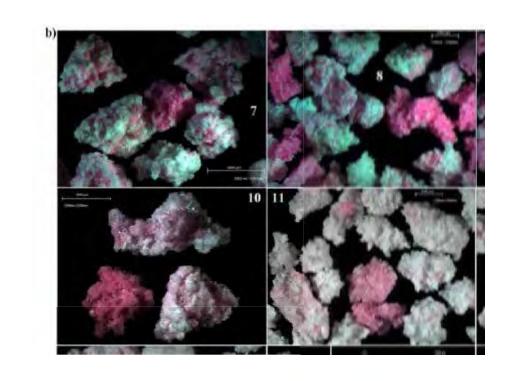
Need to save water (=heat) ...

- □ Final structure controls dissolution
- Constraint on drying method
- □ But all water added has to be evaporated



Scale up: spray drying

- Everything structured in the tower, right?
- So paint the wall layer
- □ And count the coloured particles that come out...

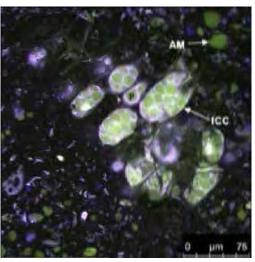


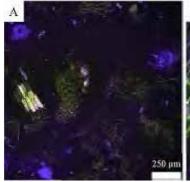
AIChE J, 61: 1804-1821, 2015

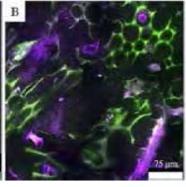


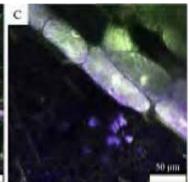
Scale up: grinding, ultrasound

- Ultrasound proposed as process intensification
- □ Works at lab but not pilot scale...





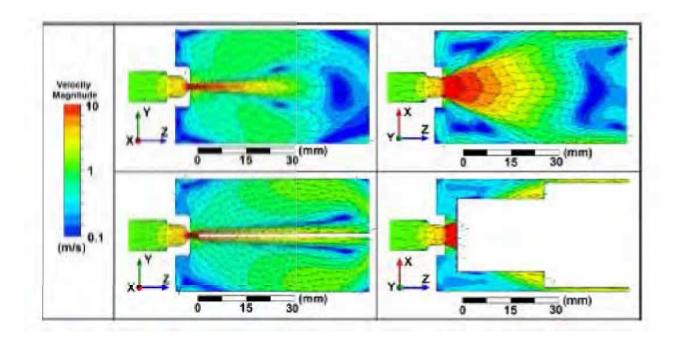




Journal of Food Engineering, 206, August 2017, 1–12



Understand the equipment: liquid whistles

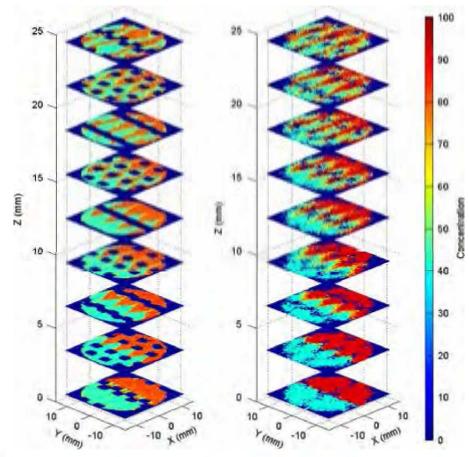


D.J. Ryan et al./Chemical Engineering Science 163 (2017) 123-136



Understand equipment: static mixers

- PEPT follows individual particle paths
- MRI measures concentration of doping agent
- □ Do they agree?...

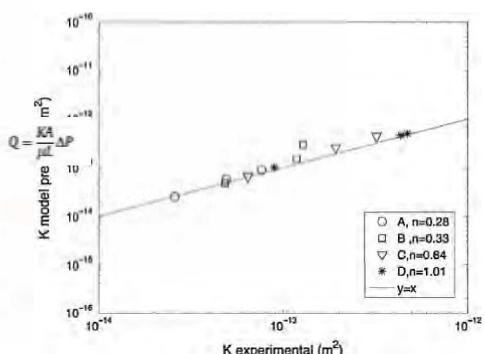






$$Q = \frac{KA}{\mu L} \Delta P$$

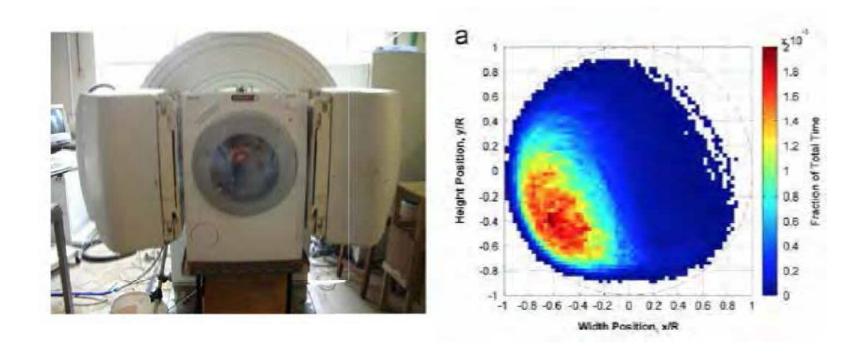




K experimental (m2)

J.Food Eng., 150, 106-116, 2015



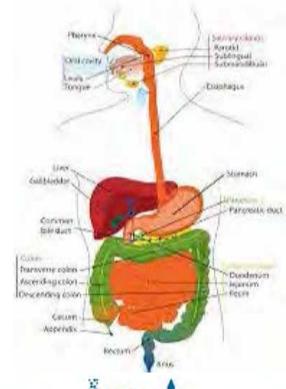


C. Mac Namara et al. / Chemical Engineering Science 75 (2012) 14-27



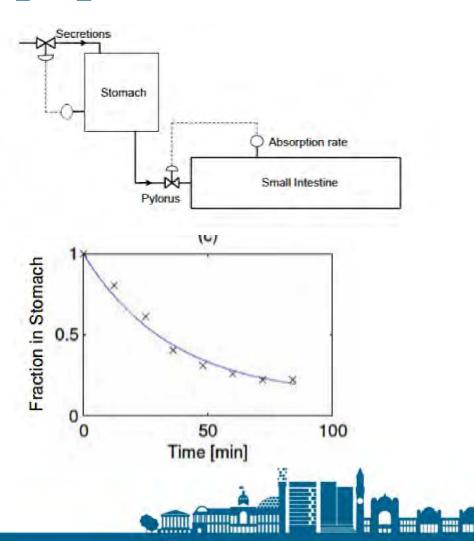
- Dry food is hydrated and acidified
- □ Then mixed and reacted
- □ Water is then removed...







- Dry food is hydrated and acidified
- □ Then mixed and reacted
- □ Water is then removed...

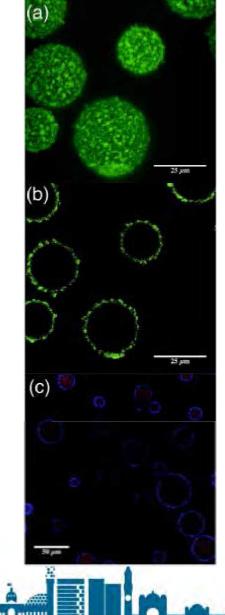


Moxon et al, in press, Chem. Engng. Sci

Make new structures

- □ Fluid gels for encapsulation
- Whey protein encapsulating fat
- New delivery
- □ Applicable to pharma, foods, HPC

RSC Adv., 2015, 5, 60786-60795



Stop new structures

- ☐ Heterotopic ossification (HO)
- Dissolvehydroxyapatite
- □ New delivery
- □ This is inverse dentistry



Eisenstein N, Stapley SA and Grover LM (2016). A novel treatment strategy in preventing and treating heterotopic ossification. *Front. Bioeng. Biotechnol. Conference Abstract:* 10th World Biomaterials Congress. doi: 10.3389/conf.FBIOE.2016.01.01939



Manufacturability

- □ How do we make sustainable supply chains?
- Minimise transport costs?
- New sustainable feedstocks
- Distributed manufacture
- Where is waste in process and after?



Formulate replacements









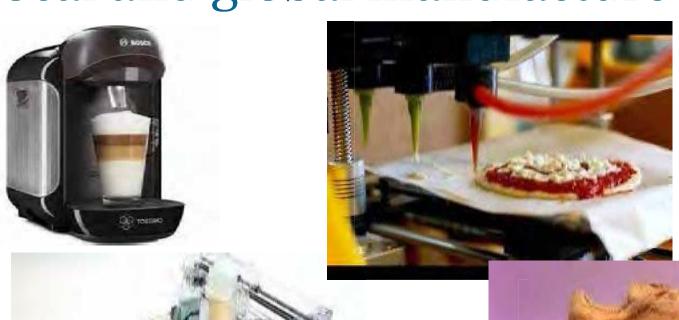
Local and global manufacture

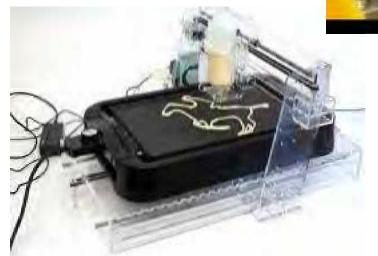






Local and global manufacture



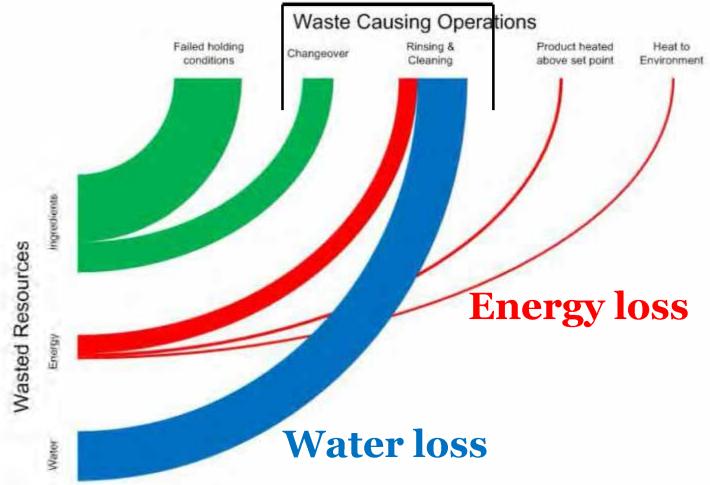






What do you waste?

Changeover and cleaning





How does cleaning work?

- □ Everyone does it... nobody knows how
- □ Tend to be two problems:
 - Product recovery: at the end of a run, the plant is full of product;
 - Cleaning: at the end of product recovery, there is still stuff to remove
- □ Efficient measures of cleaning time at plant scales
- □ To know what *not to do…*



Cleaning with an impinging jet

8mm thick deposit of Carbopol® solution



1000 ml min⁻¹ 16.7 x 10⁻⁶ m³ s⁻¹

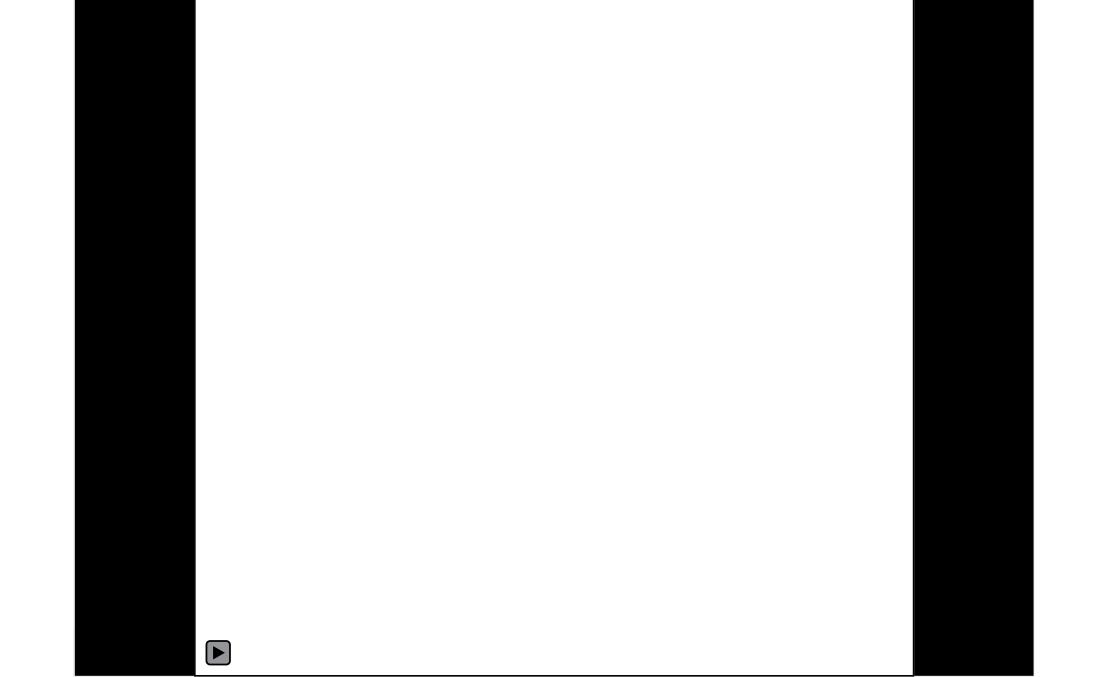
1500 ml min⁻¹ 25.0 x 10⁻⁶ m³ s⁻¹

COFE: Ohio State, Sept 2016

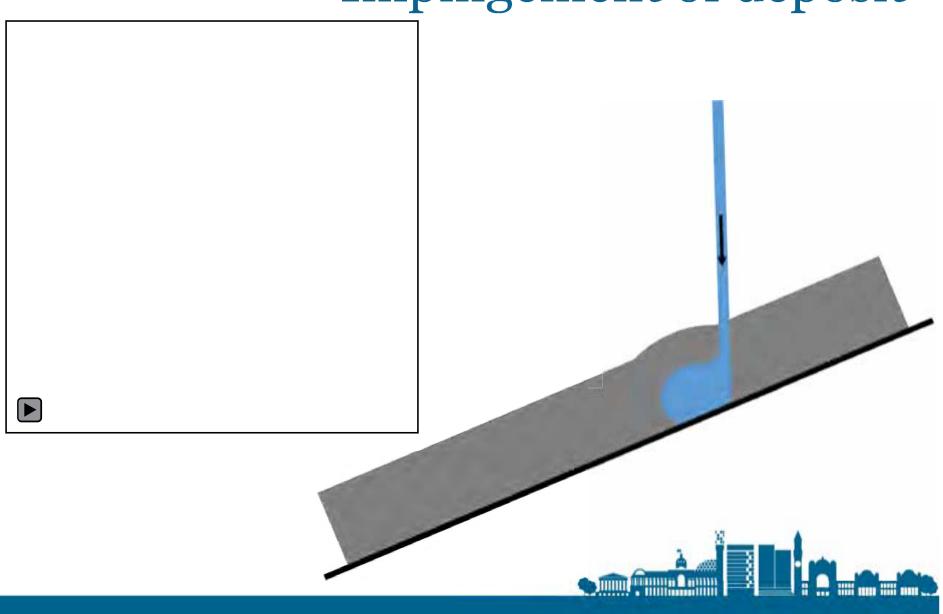


3 seconds later 1500 ml min⁻¹ 1000 ml min⁻¹ Adhesive cleaning Cohesive cleaning

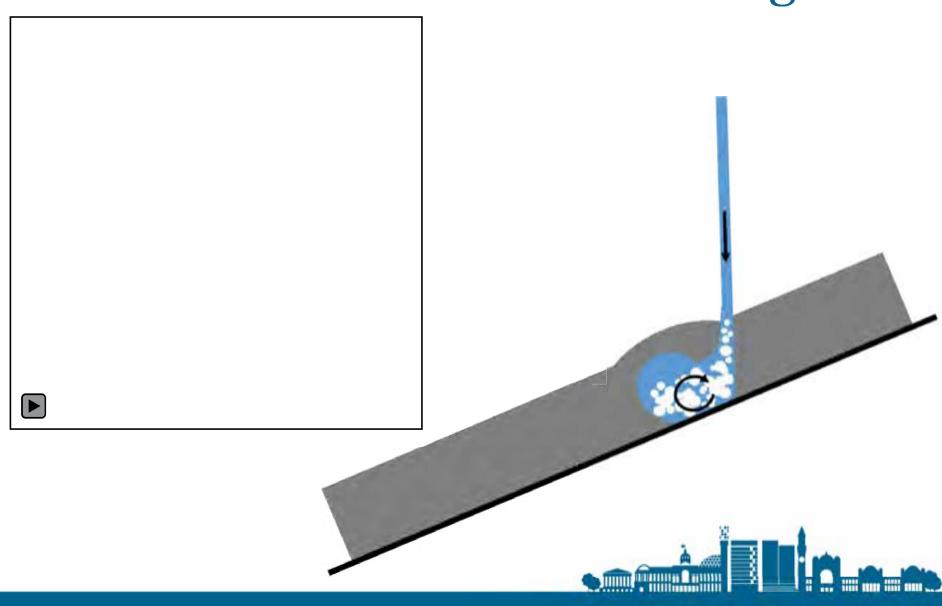




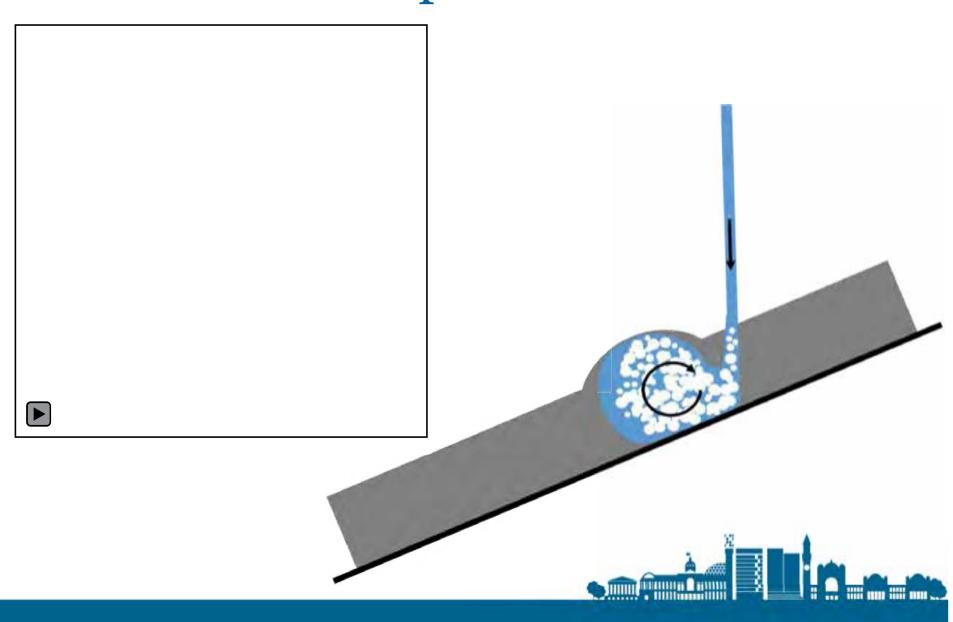
Impingement of deposit



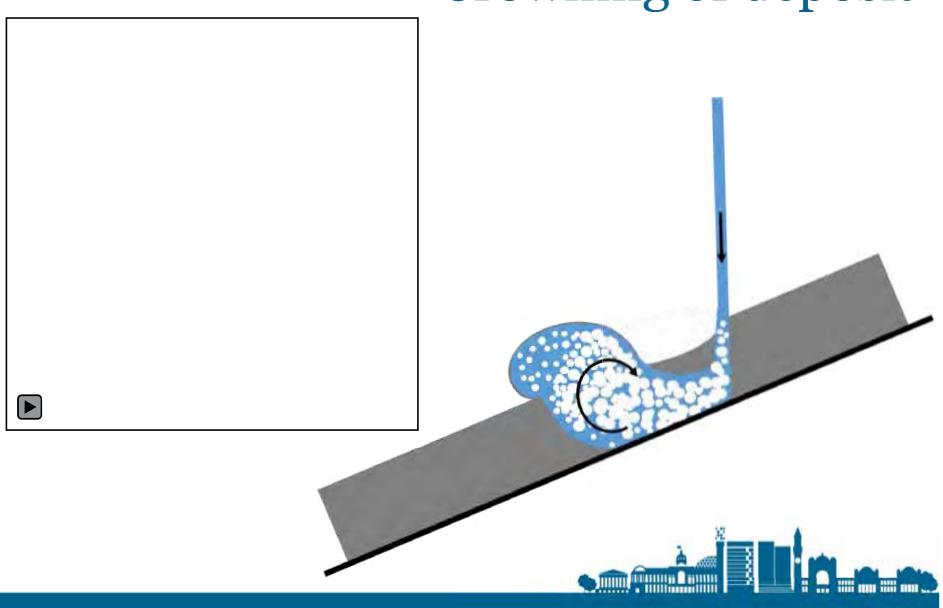
Entraining Air



Expansion of water sack



Crowning of deposit



Conclusion

- □ Manufacturability is...
 - Understanding scaleup
 - Working out how the equipment actually works
 - What does the equipment actually do to the product?
- □ Constraints are complex: microbial safety, product acceptability and cost
- Water and energy are parts of the manufacturing process
- ☐ How do you clean 3D printers?





Thank you

