

Water-Insoluble α -Glucan Polysaccharide Nanoparticles for Controlled Release of Compounds

Kervin O. Evans, Christopher
D. Skory, David L. Compton,
Michael Appell



Presented by Kervin Evans
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Thanks to the Following:



David Compton – ARS Chemist,
Feruloyl Soy Glycerides synthesis



Michael Appell – ARS Chemist,
molecular modeling



Greg Cote – Retired ARS Chemist, ,
 α -glucan polysaccharide synthesis &
purification



Chris Skory – ARS
Microbiologist, α -glucan
polysaccharide synthesis &
purification



Suzanne Unser – High
pressure homogenization, α -
glucan polysaccharide
synthesis & purification,
Technical Assistance



Leslie Smith
Retired, Technical
Assistance



Ryan Cormier, former ARS,
Catalent Pharma Solutions, α -
glucan nanoparticles & films

National Program 306 – Product Quality and New Uses

Problem Statement 1.B – New Bioactive Ingredients and Health-Promoting Foods

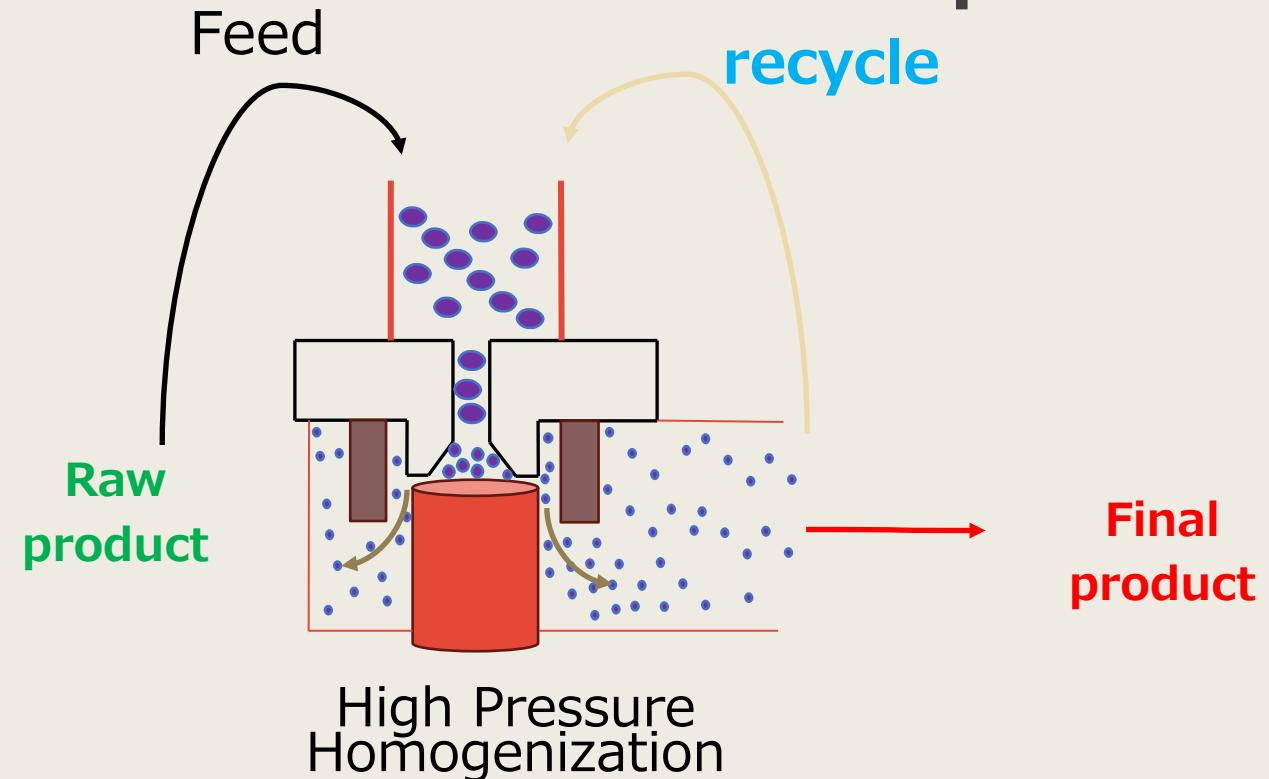
Problem Statement 2.B – Enable technologies to produce new/expand marketable non-food, non-fuel biobased products derived from agricultural feedstocks

CRIS – Technologies for Producing Marketable Bioproducts

Objective 2. Develop innovative lipid and biopolymer-based encapsulation systems for delivering, preserving, or promoting the activity of bioactive ingredients.



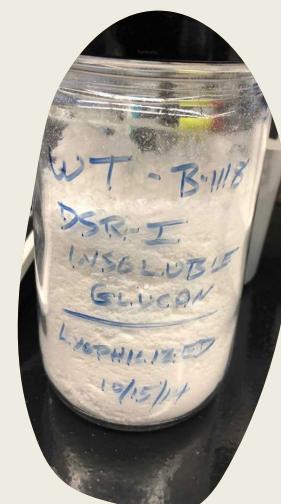
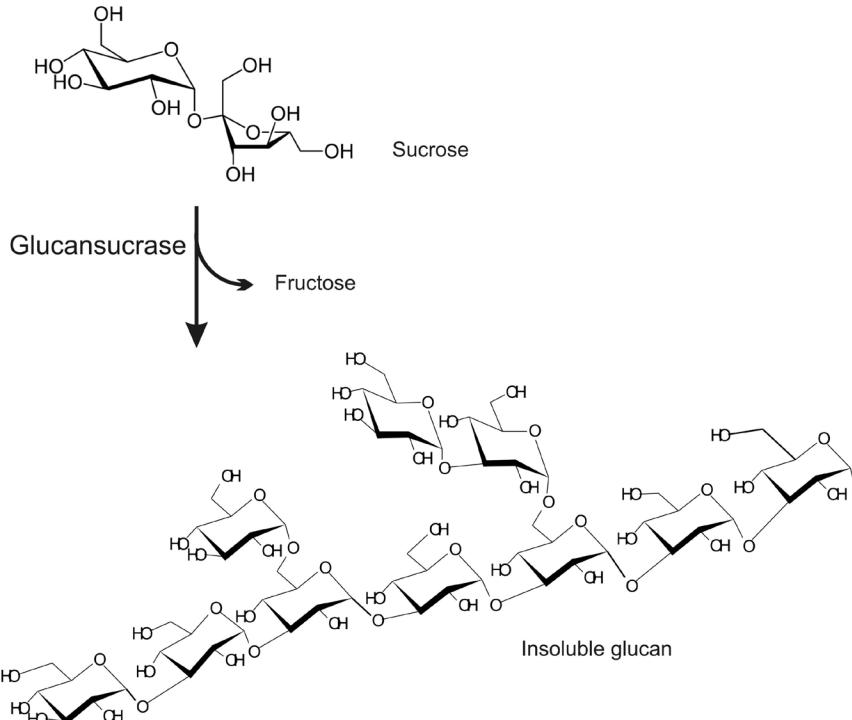
Nanoparticles via High-Pressure Homogenization for Nanoencapsulation



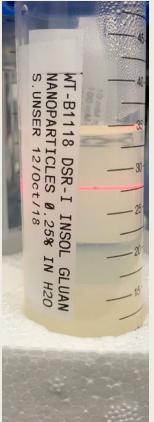
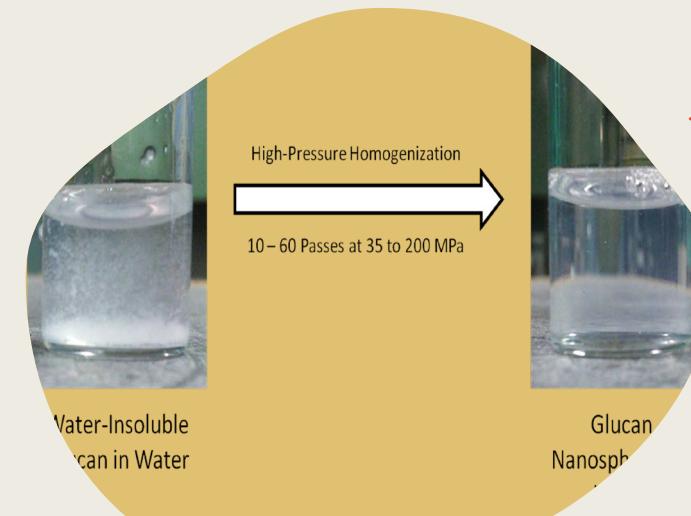
High pressure homogenizer

Sample solutions forced through small openings via high pressure shear forces (MegaPascal) to form small particle (<600 nm diameter)

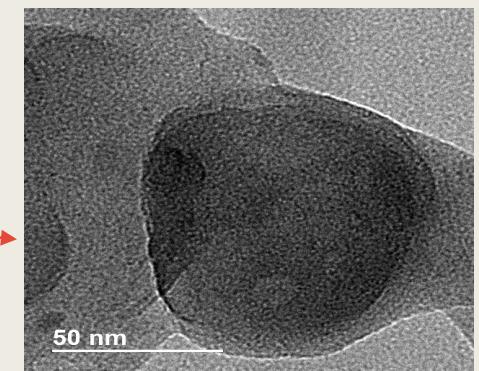
Water-Insoluble Glucan Nanoparticles Formation



MW \sim 1.2 Mega Dalton
(polymer \sim 1.4 Mega Dalton)



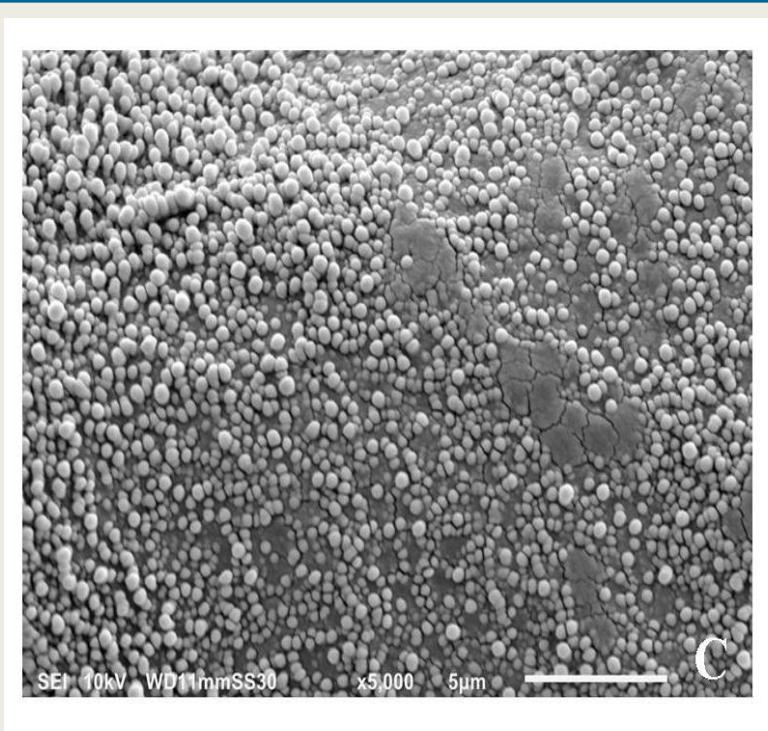
Example of alpha-glucan nanoparticles scattering red laser light



Cryo-TEM

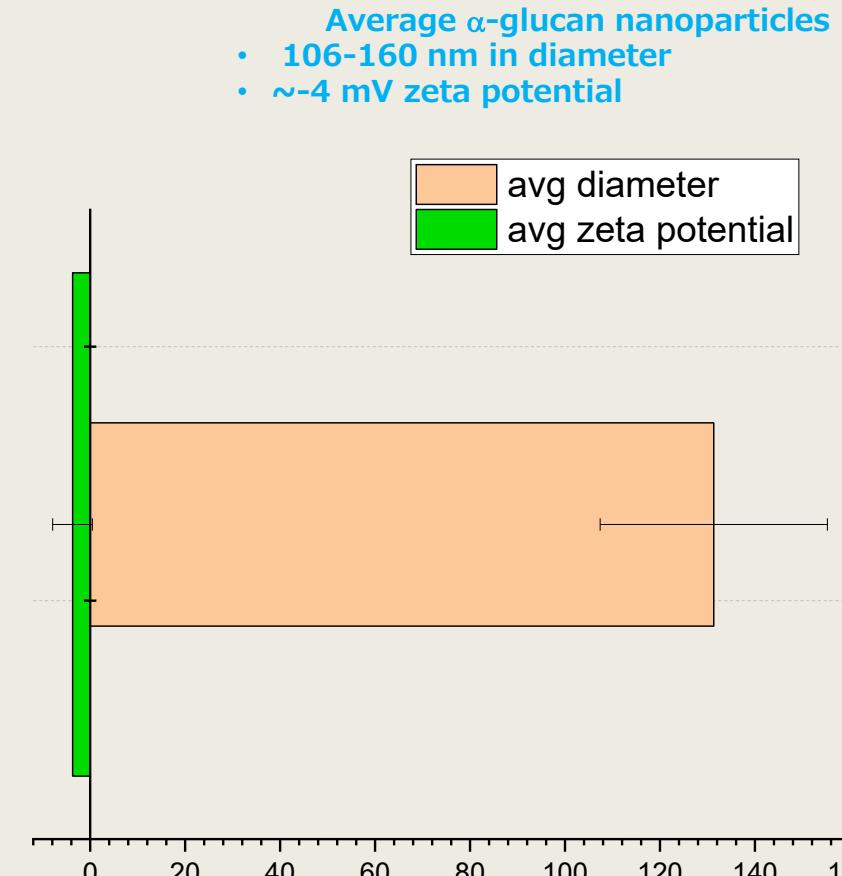
Evans et al, 2020, Molecules 25 (17): 3807

α -Glucan Nanoparticles Physical Properties



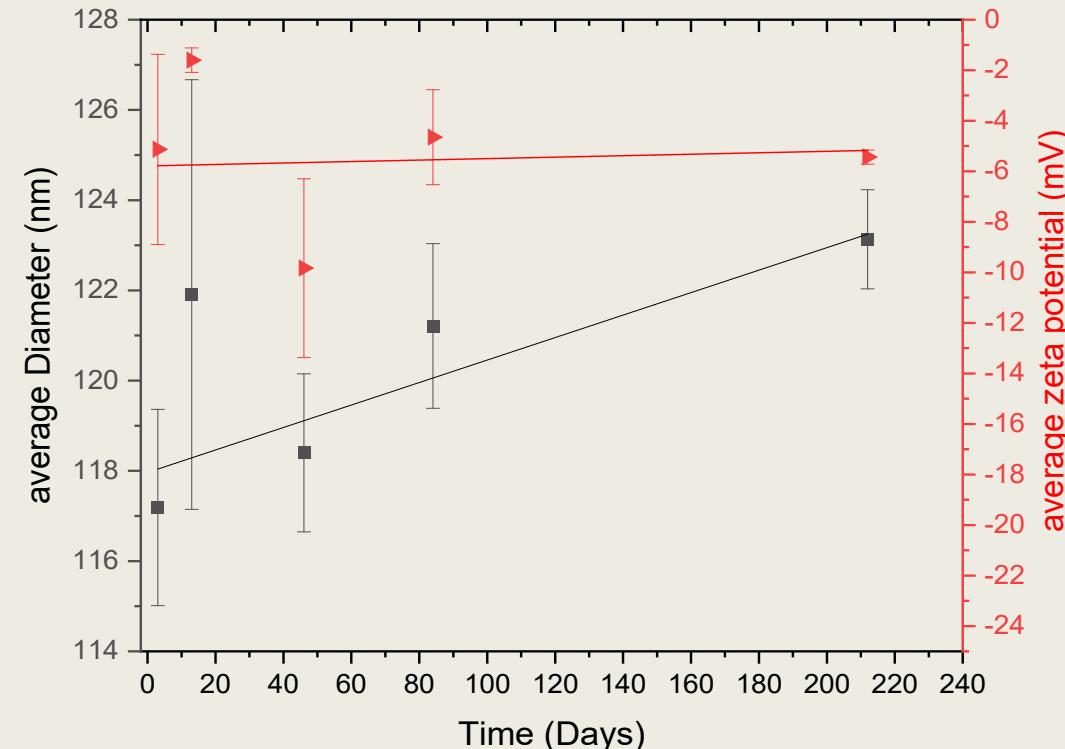
SEM image shows the spherical nature of freeze-dried polysaccharides after high-pressure homogenization

Cormier et al, 2017, US Patent
9,708,417 B2



α -Glucan Nanoparticles Stability

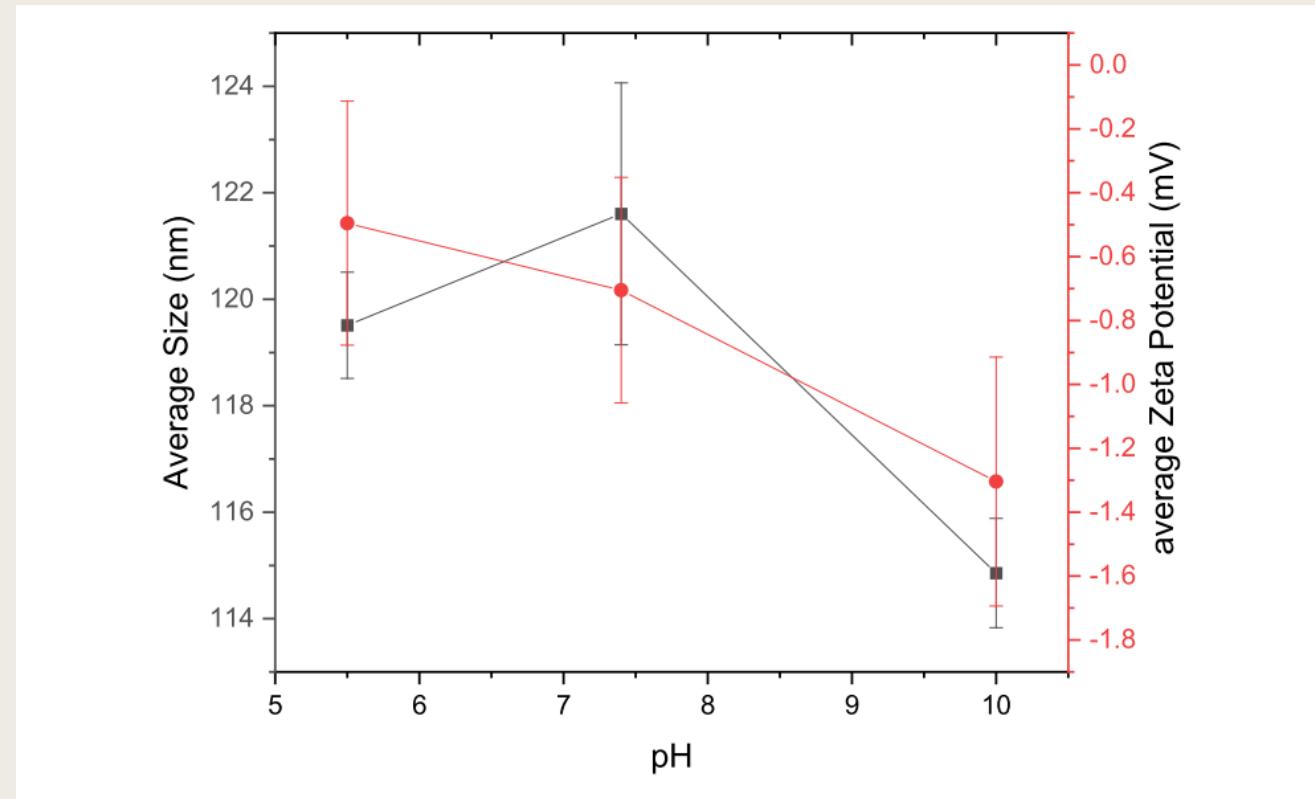
- α -Glucan Nanoparticles show stability for more than 7 months
- Likely due to steric hindrance of highly dense particle



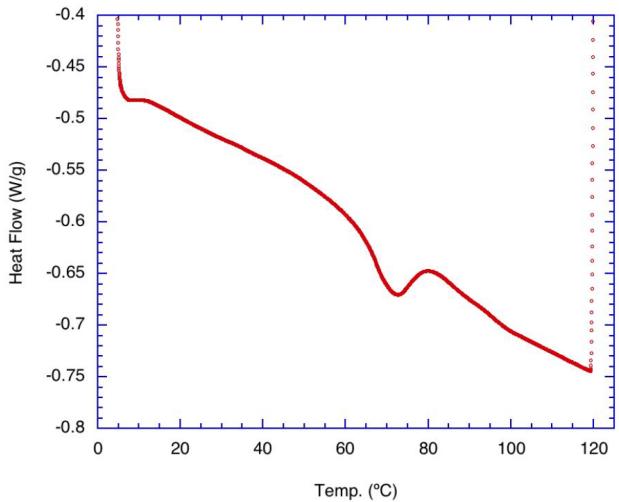
Stability suggests α -glucan nanoparticle can be used for long term controlled-release system

α -Glucan Nanoparticles Under Slightly Acidic and Basic Conditions

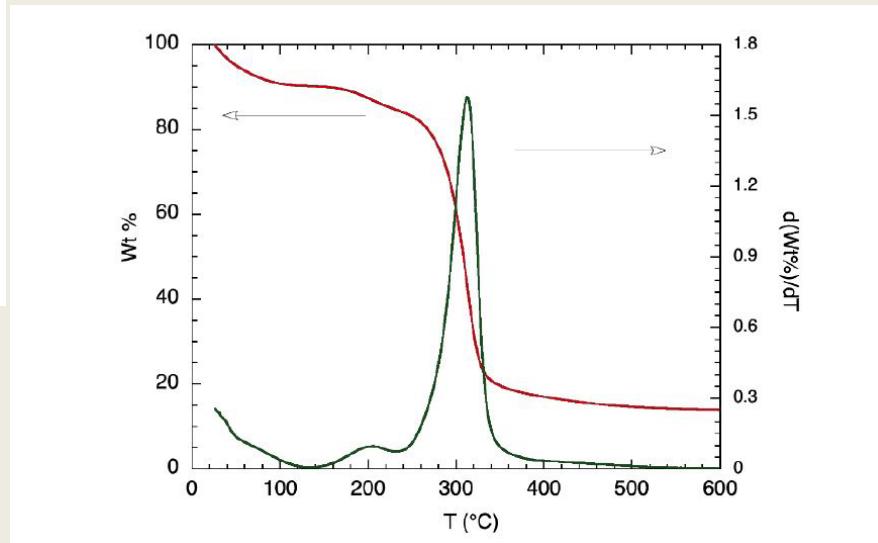
- Alpha-glucans can be made into nanoparticles at pH 5.5, 7.4 and 10 under buffered conditions.



What are Thermal Properties of α -Glucan Nanoparticles



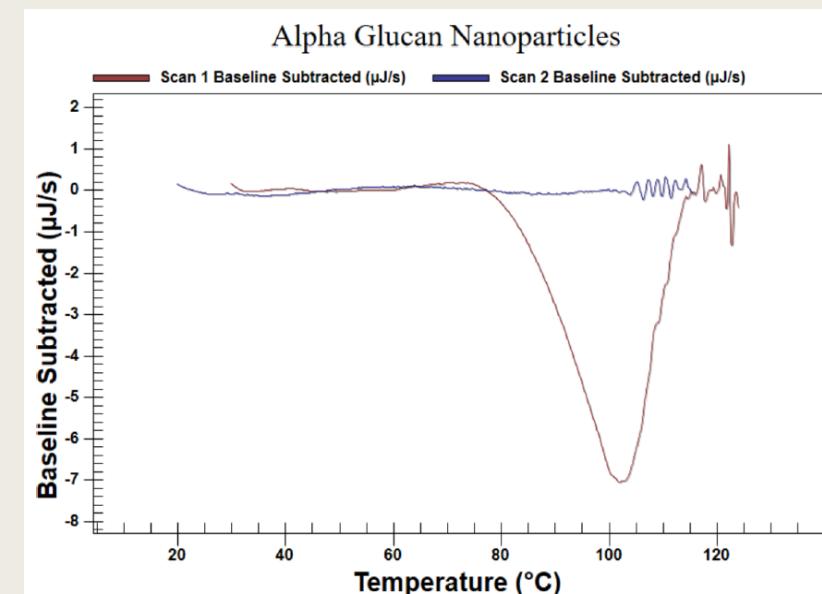
DSC Thermal analysis of α -glucan show structural changes likely to occur $\sim 70^\circ \text{C}$.



TGA shows that α -glucan thermally degrades between 160°C and 340°C .



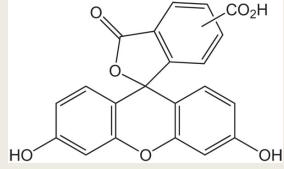
Nano DSC shows that nanoparticles start to unravel at $\sim 70^\circ \text{C}$ and completely unravel at $\sim 104^\circ \text{C}$.



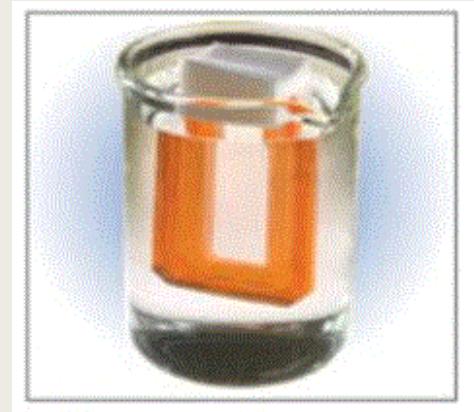
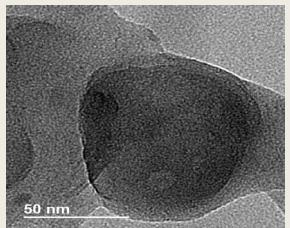
Evans et al, 2020, Molecules 25(17), 3807

Evans et al, 2023, Biotechnology Reports 40, e008107

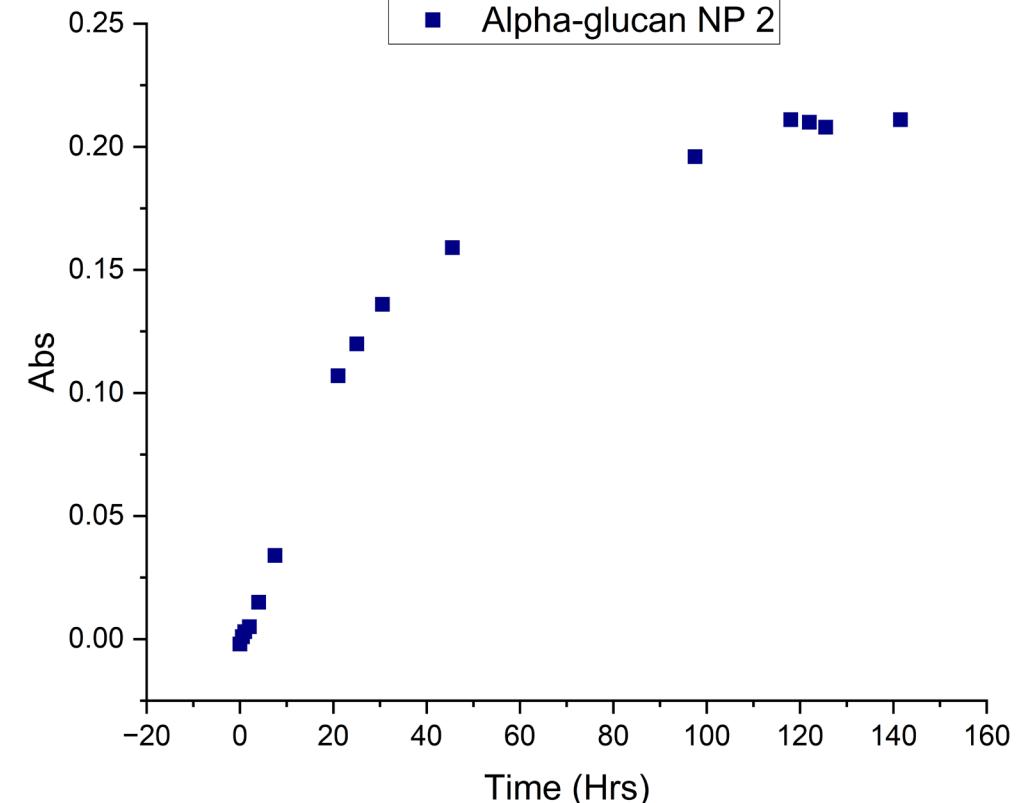
Hydrophilic Encapsulation within α -Glucan Nanoparticles

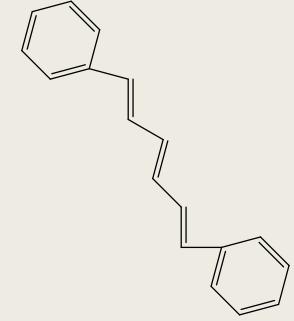


5 (6)-carboxyfluorescein

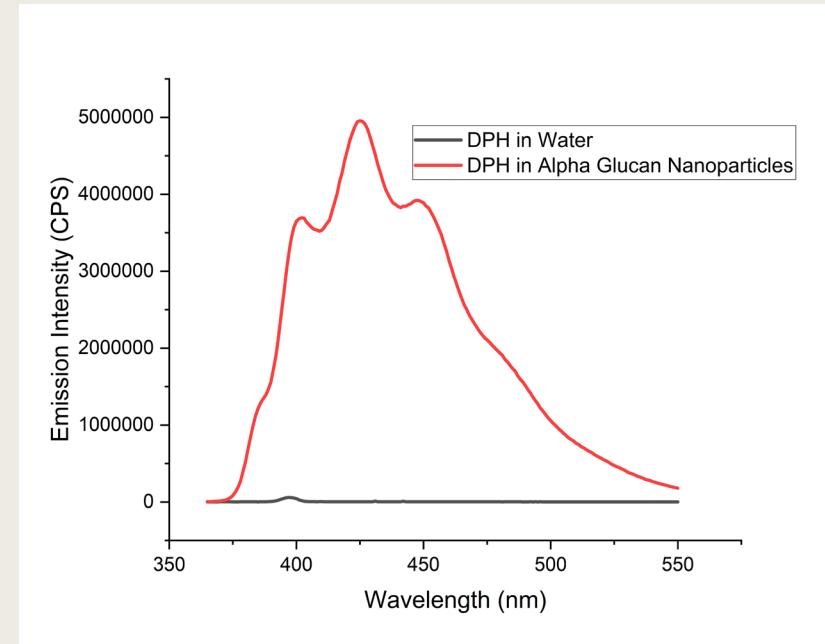


- Alpha-glucans nanoparticles show release of carboxyfluorescein for up to 120 hr during 6-day dialysis.

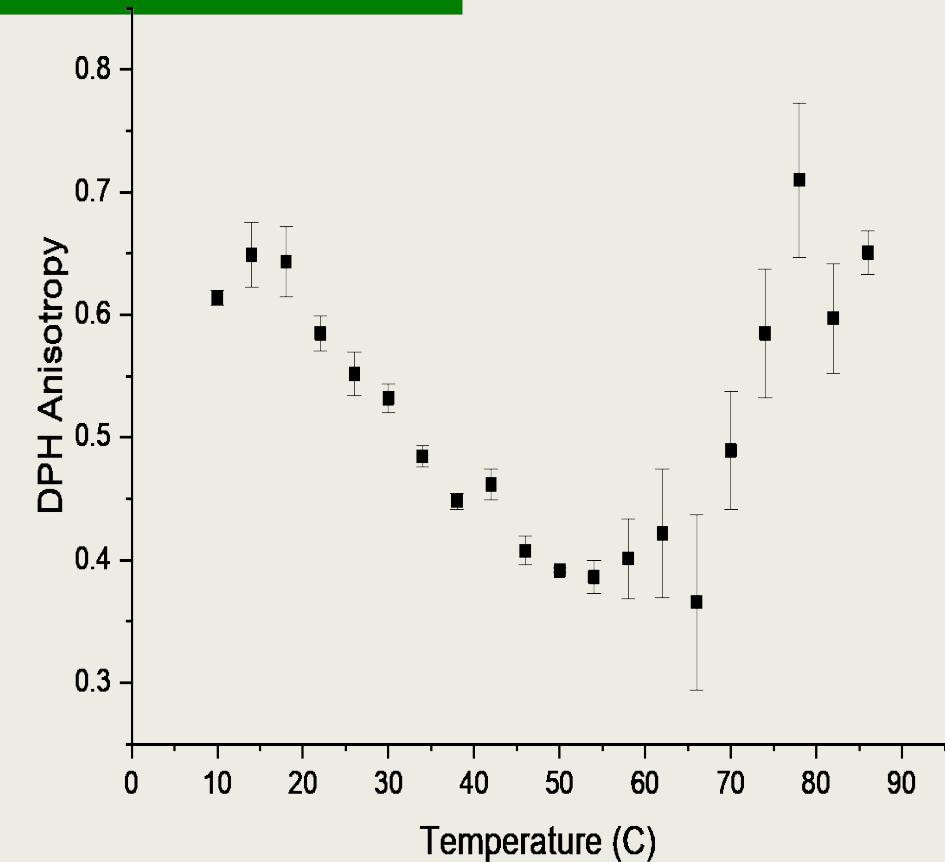




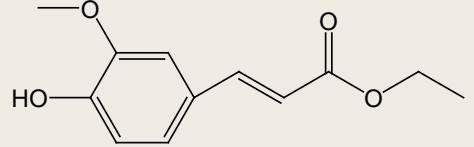
1,6 Diphenyl-1,3,5-hexatriene (DPH)



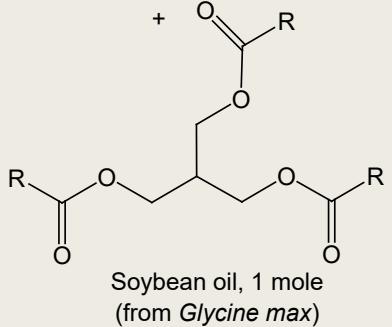
- **Alpha-glucans nanoparticles have internal hydrophobic regions**
- **Anisotropy measurement show structure changes but not complete degradation.**



Can Feruloyl Soy Glycerides (FSG) Encapsulate within α -Glucan Nanoparticles???

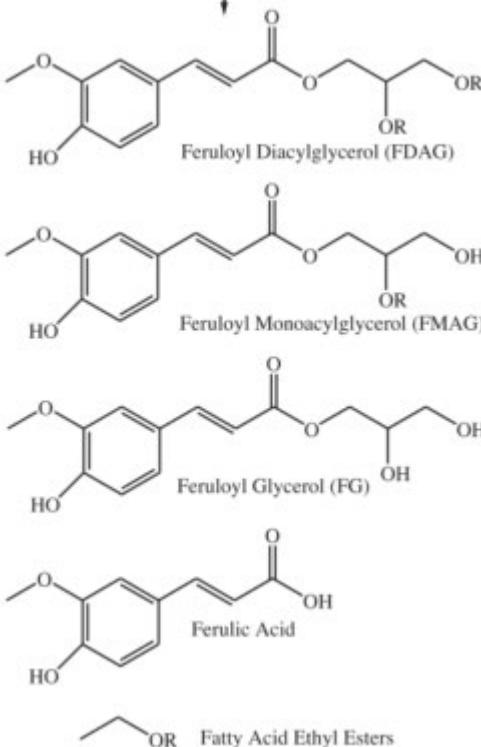


Ethyl ferulate, 1 mole
(CAS # 40446-02-0)



Soybean oil, 1 mole
(from *Glycine max*)

trace amount of H_2O
 $Candida antartica$ lipase B (Novozym 435)
 $60^\circ C, 144 h$

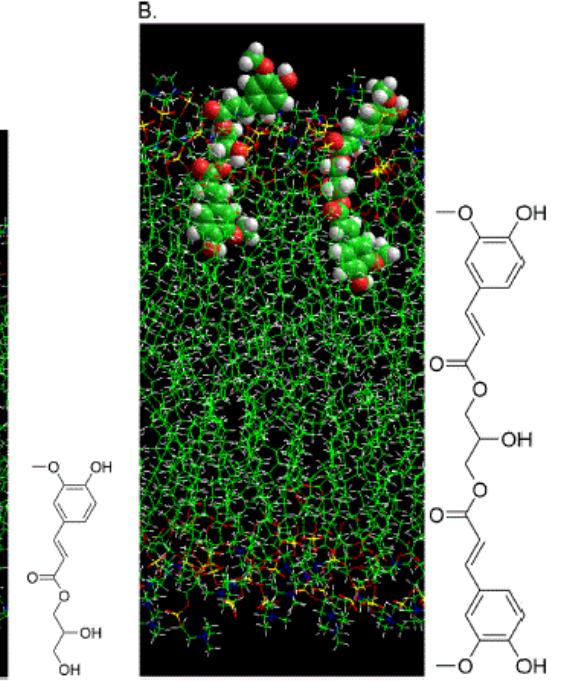
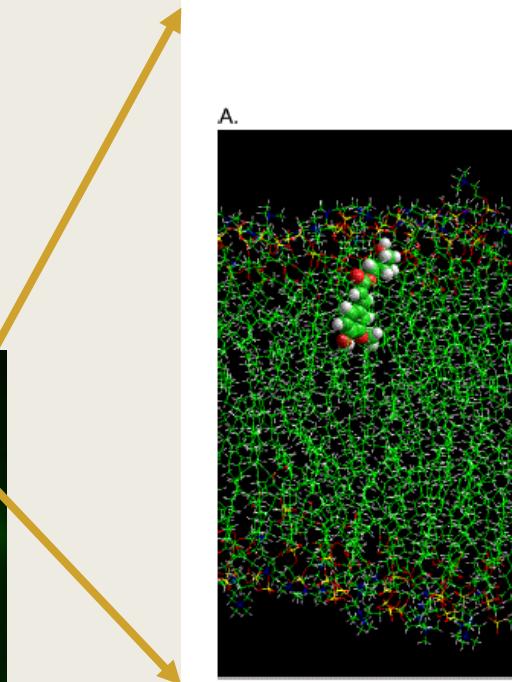
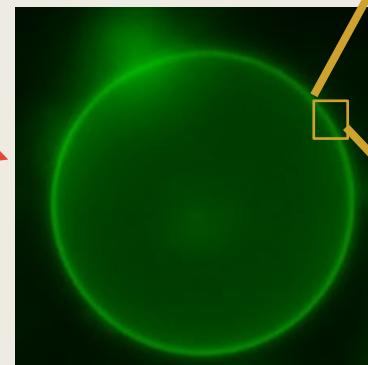


Compton et al, 2015, Industrial Crops and Products 77:787-794

Isolated Components of Feruloyl Soy Glycerides Encapsulate within Liposomal Bilayer. Will they encapsulate within dense α -Glucan nanoparticles?



1-Feruloyl glycerol
1,3-Diferuloyl glycerol

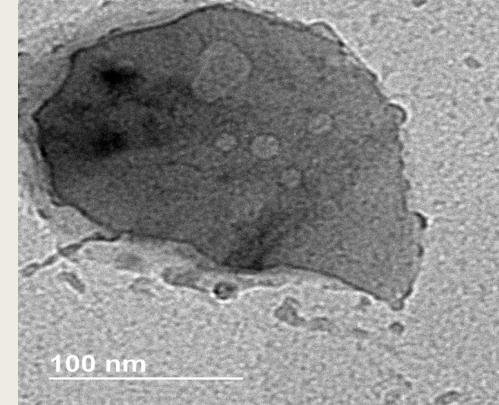
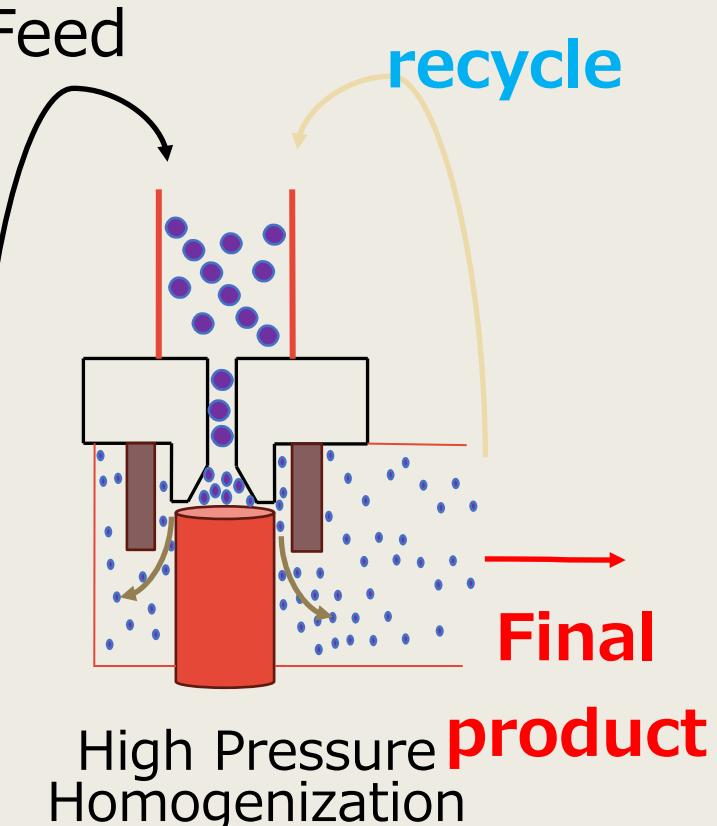


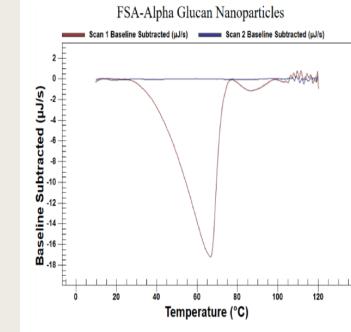
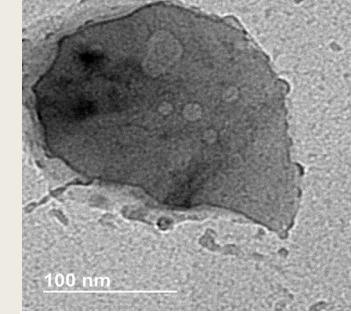
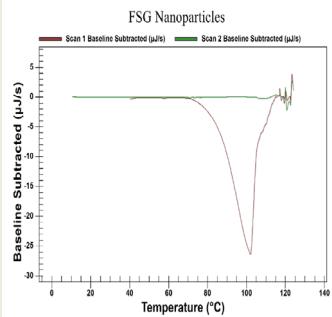
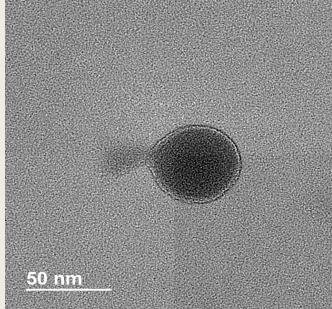
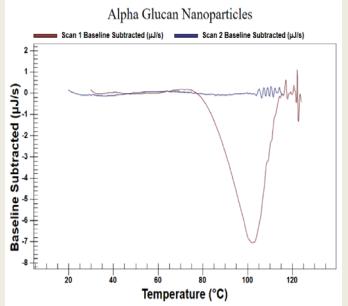
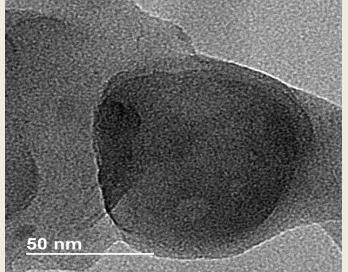
Evans et al, 2016, *Chem. Phys. Lipids* 195:1–11

Can Feruloyl Soy Glycerides (FSG) Encapsulate within α -Glucan Nanoparticles???



α -glucan
+
FSG





Alpha-glucans nanoparticles

- \sim 126 nm diameter
- \sim -5 mV
- \sim 75° C – 102° C irreversible unraveling

FSG nanoparticles

- \sim 190 nm diameter
- \sim -25 mV
- \sim 70° C – 115° C irreversible degradation

FSG/Alpha-glucans nanoparticles

- \sim 216 nm diameter
- \sim -15 mV
- \sim 98% encapsulation
- \sim FSG release 26° C – 66° C

Conclusions

- **Insoluble α -glucan forms nanoparticles stable up to 7 months and 80° C under neutral conditions; can be made at pH 5.5 – 10;**
- **α -Glucan nanoparticles can encapsulate and release hydrophilic and hydrophobic materials;**
- **α -Glucan nanoparticles encapsulate 98% of 5% (w/v) feruloyl soy glycerides;**
- **α -Glucan nanoparticles release feruloyl soy glycerides between 26° C and 66° C;**
- **α -Glucan nanoparticles may be good release system for oils.**



Thank You for Your
Attention!!!

Questions???