B-carotene – a long road to commercial implementation

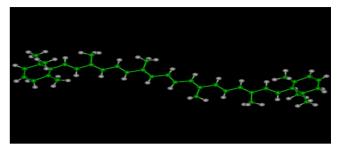
Dusty Gardiner, Raj Lalloo, Dheepak Ramduth, Jozef Dudas, Mark Marcus (NCSA)





Uses of β -carotene (pro-vitamin A)

• Animal feed supplements



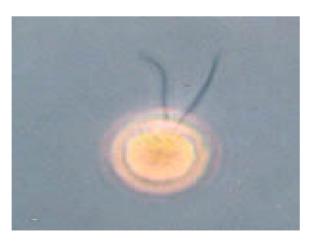
- Food supplements colorant, Vitamin A supplement
- OTC pharmaceuticals (health supplements) anti-oxidant, Vitamin A supplement





Production

- Chemical synthesis
 - BASF, DSM (formerly Roche)
- Algal production (*Dunaliella salina*)
 - Betatene (Cognis), Koor Foods, NCSA
- Extraction from natural sources (carrots, palm oil)
 - DSM (formerly Roche), Hanson and Darius
- Fungal production (*Blakeslea trispora*)
 - DSM







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Research and development – algal production

- Growth and stress phases biomass and carotene production
- "Raceway" pond design selected
- Key parameters:
 - Nutrient optimisation
 - Biomass balance between productivity and self-shading
 - Growth rate temperature and light key parameters
 - Light growth and stressor
 - Temperature growth
 - Surface area light intensity
 - Mixing shading and mass transfer
 - Carotene productivity stressors (salinity, nutrient limitation, light intensity)





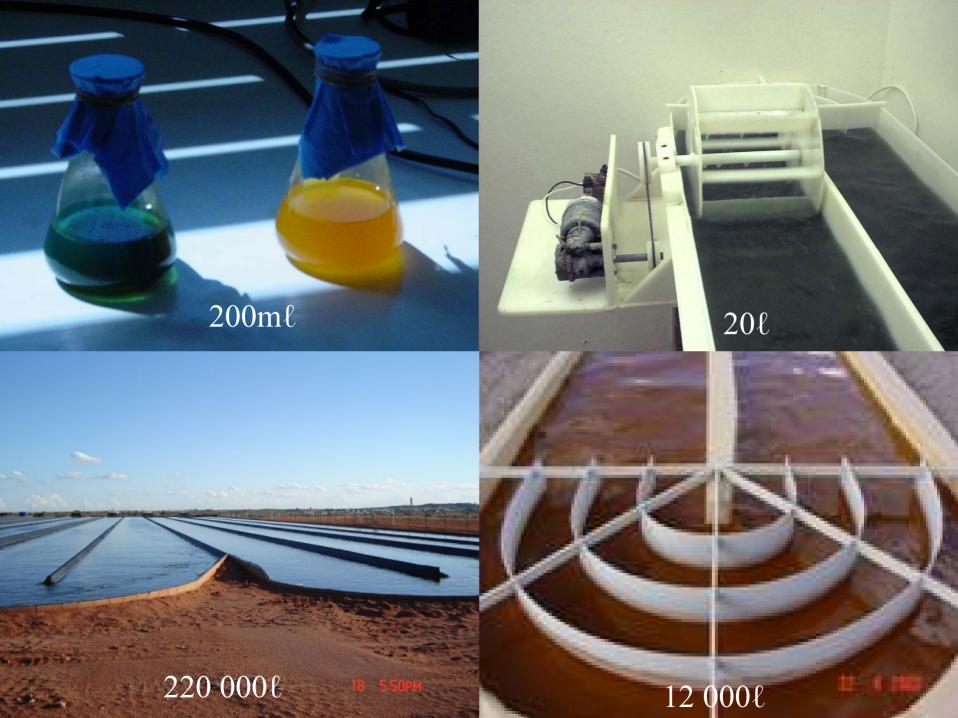
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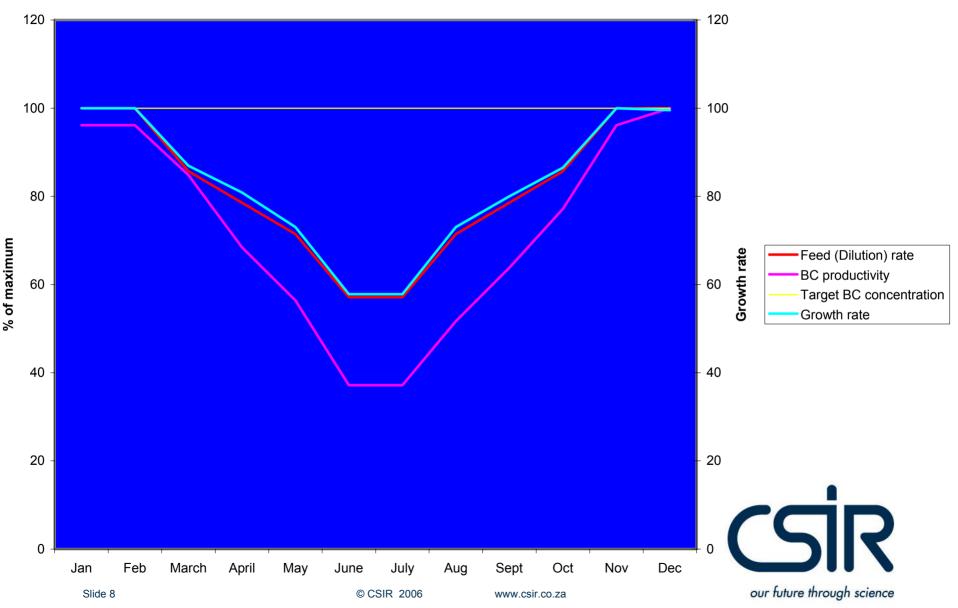
Key challenges

- Many parameters are location specific and require on-site optimisation – pilot plant in Upington
- Transfer of lab research to location specific pilot and production plant
- Integration of upstream and downstream process options
 - Size differentials (250 fold)
 - Upstream process variability impact of climatic variables
- Scale–up
 - Mixing
 - Optimisation against climatic parameters
 - Recycles





Process strategy



Carotene Concentration 200 180 160 140 120 100 80 60 40 20 0 3/2002 2/2002 2/2002 07/01/2002 14/01/2002 04/02/2002 11/02/2002 26/11/2001 17/12/2001 24/12/2001 31/12/2001 21/01/2002 28/01/2002 03/12/2001 10/12/2001 25/ Date

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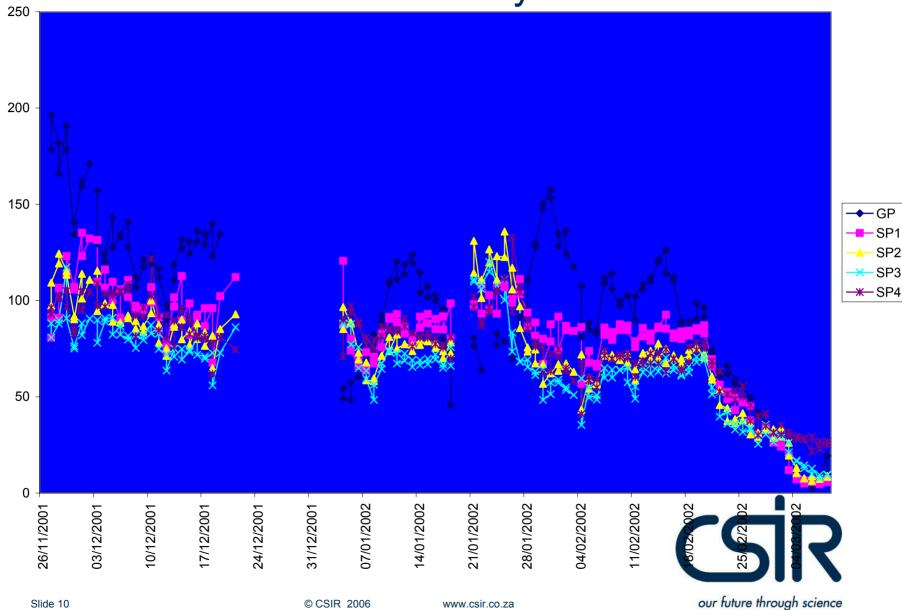
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our future through science

SP1

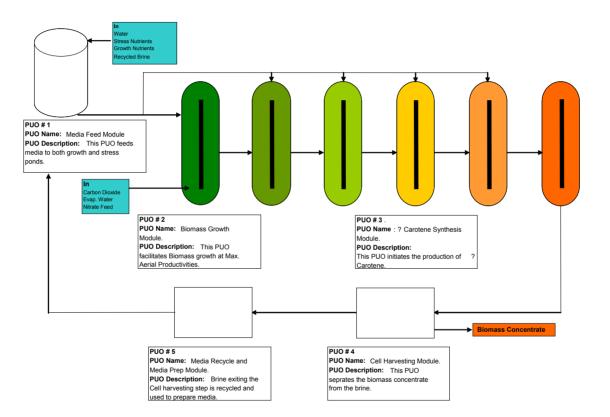
SP3 SP4

Cell density



SP2 SP3

Final upstream process – carotene production



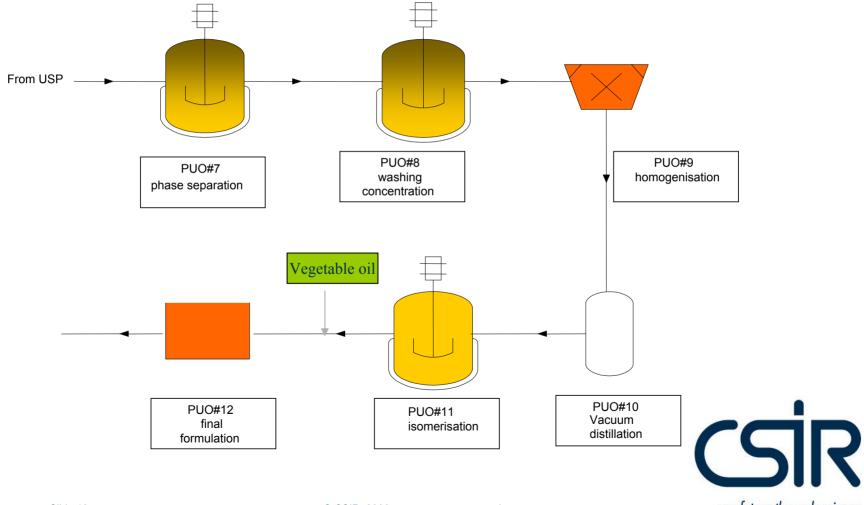


Key R&D challenges – carotene extraction and formulation

- Phase separation
 - 3 layers after solvent extraction
 - "Green layer" containing chlorophyll and mixed carotenoids
 - De-emulsification step introduced
- Solvent removal
 - Product specification < 50ppm food grade solvent
 - Difficult to achieve in two phase product (solid-liquid)
 - Required introduction of homogenisation and additional distillation process step
- Technology integration
 - Process sizing 250X step down from pond to solvent extraction
 - Impact of algal biomass variability on downstream unit processes
 - Solution implemented is to maintain carotene concentration, fix downstream unit operations and vary downstream capacity utilisation



Final process – carotene extraction and formulation



Challenges in technology implementation

- Product specifications
 - Chemical specs vs in house application specific specifications
 - Introduce new customer specific quality specs (require process tweaking)
- Scale up
 - 25 fold scale up from pilot to commercial ponds
 - Mixing and mass transfer dead zones
 - Additional process optimisation at large scale
 - Recycle of brine
 - Impurity build up
 - Light penetration
 - Required brine clarification step
- Technology transfer
 - Transfer of research staff for upstream process

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On site, hands on learning at toll facility



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Commercialisation

- Sasol
 - Non-core business
- AECI
 - Exit from fine chemicals business
- NCSA
 - New start up acquired license to the technology





Key differences between large chemical industry and new start up

- Size/value of new entity
 - R50M R100M minimum turnover vs ~R10M
- Affordable investment in R&D
 - Large industry will invest far more to reduce technology risk
 - Affordability a key issue for start-up enterprise. More reliant on university/government research and funding
- Appetite for technology risk
 - Large industry will attempt to research/engineer out risk
 - Start-up enterprise will take more risk and continue research phase on to commercial entity
- Implementation strategy
 - Large industry will invest in stand-alone plant
 - Start-up enterprises will innovate to reduce investment requirements
 - Phased roll out
 - Outsourcing



Key challenges to commercialisation

• The right entrepreneurs

Affordability and appetite for risk

- Availability of venture capital / angel funding
- Solution was to reduce investment required :
 - Innovative mix of own plant and toll production by CSIR
 - · Phased implementation and investment

Finance

- Investment by Cape Biotech Trust and Bioventures
- Good example of public/private investment
- Market
 - Barriers for new players availability of market samples from pilot plant
 - Customer specific product specifications







What next?

- Expansion of carotene production capacity planning stages
 - Enthusiastic acceptance of product in the market
- Development of Upington "Algal Hub"
 - Astaxanthin (UCT and Cape Biotech Trust)





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Thank you for your attention !!

