



Innovater avec les enzymes

Adebiotech - Paris 27-28 octobre 2014

- **Pharmaceuticals ;**
- **Cosmetics ;**
- **Home & Personal Care;**
- **Agro-industry ;**
- **Environment ;**
- **Energy ;**
- **Others (paper, textile, chemistry ...)**

Fields of Innovation

Consulting

- Desk-based feasibility studies and proposal of strategies

Screening

- Proteus enzymes portfolio
- Micro-organisms library

Optimization

- Enzyme engineering (Mol. Biol.)
- Improvement of enzyme manufacturing process – Formulation

Scale up

- Production of enzymes
- Optimization of fermentation process
- Management of CMOs

Industrialization (PCAS)

- Manufacturing using chemo-biocatalytic processes under GMP or non GMP standards

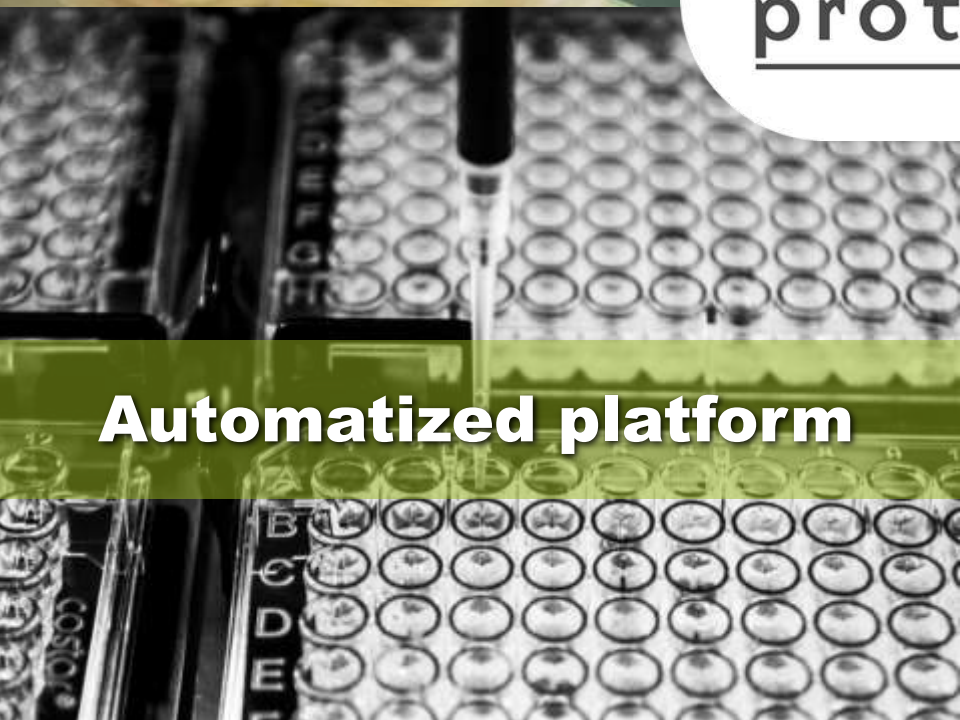


Multi-disciplinary Team



Biodiversity resources

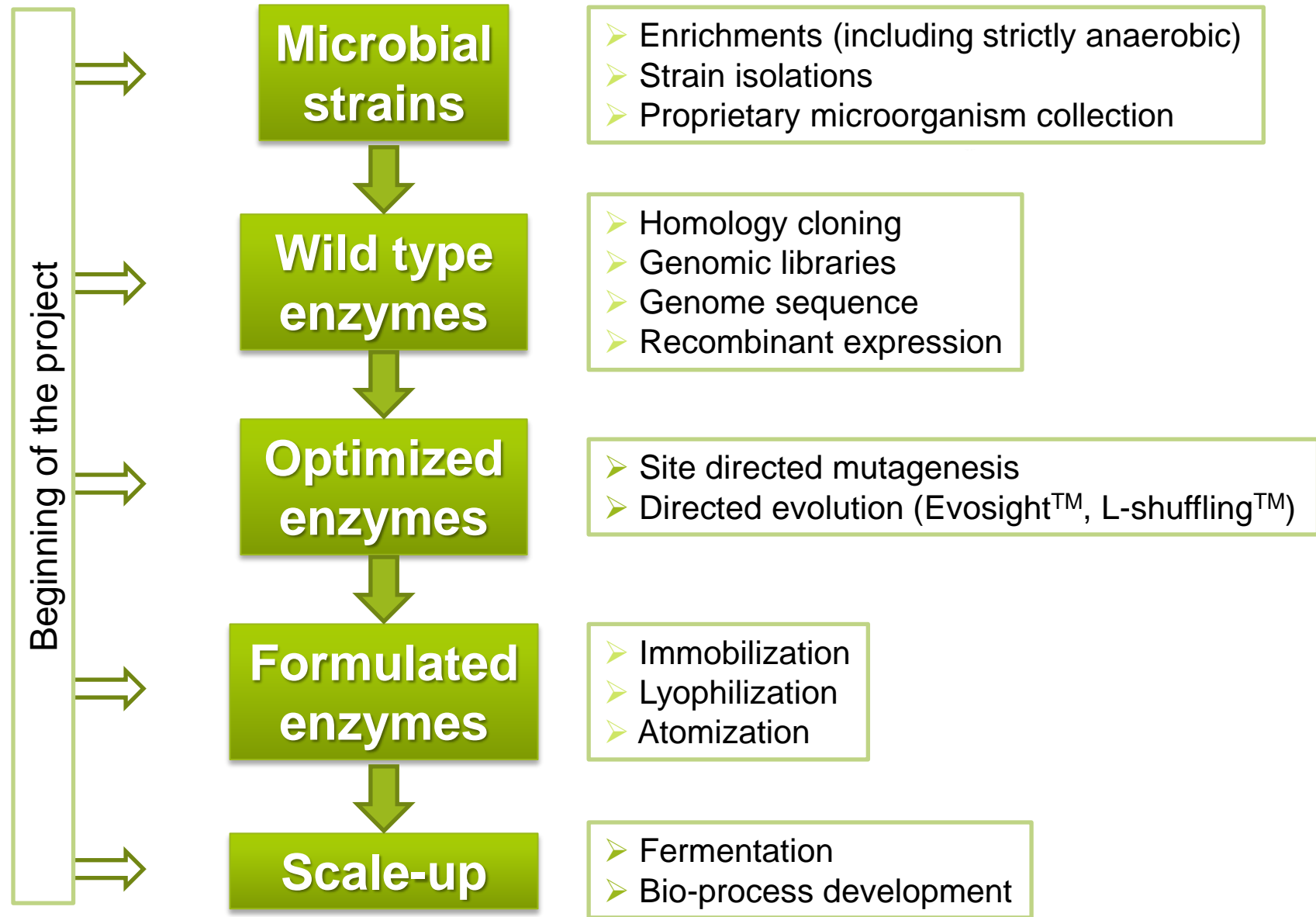
proteus
pcas



Automatized platform



Commercial Production



Strategies for obtention of fine-tuned enzymes



Examples : dehalogenases

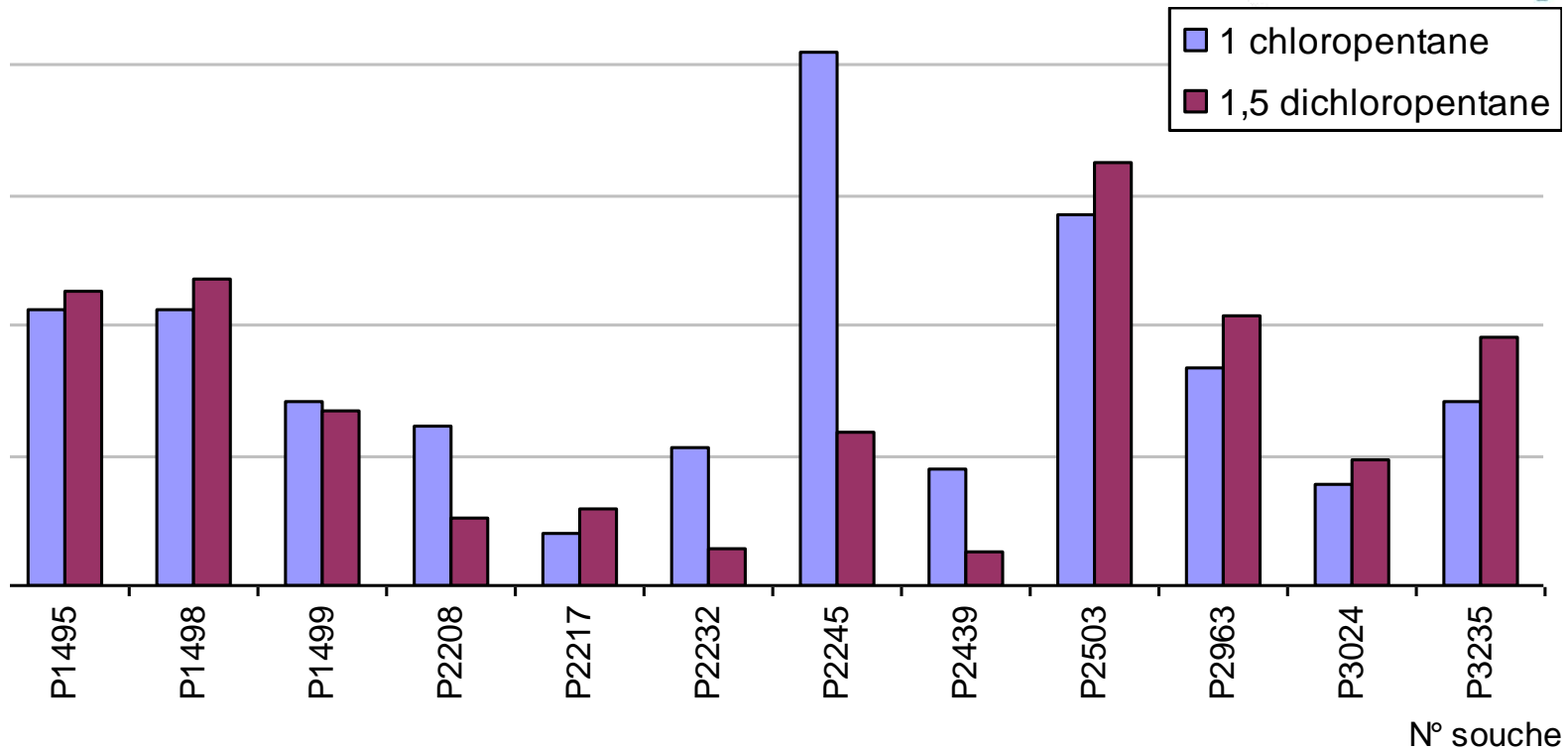
Different strategies for different substrates

Carbon source : targeted substrate
Inoculum = suitable environmental samples (polluted with halogenated compounds)



| substrate | Isolated strains |
|---------------------------|--|
| chloro-3-butyric acid | 29 |
| 3-chloro-1, 2-propanediol | 53 |
| 2-chlorobutyramide | 43 (incl. 28 enantioselective for one or the other enantiomer) |
| bromo-2-hexanoic acid | 14 |

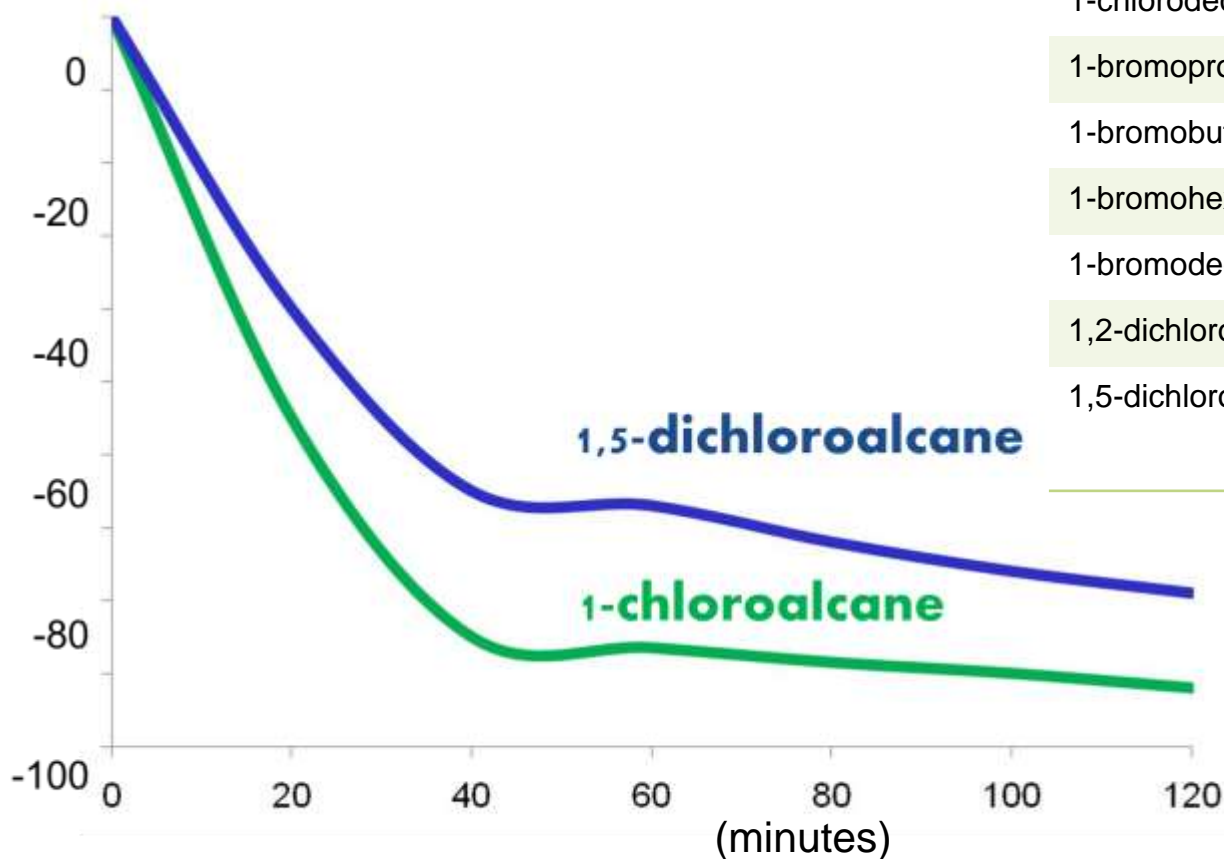
Enrichments and isolations



Screening of the collection of microorganisms

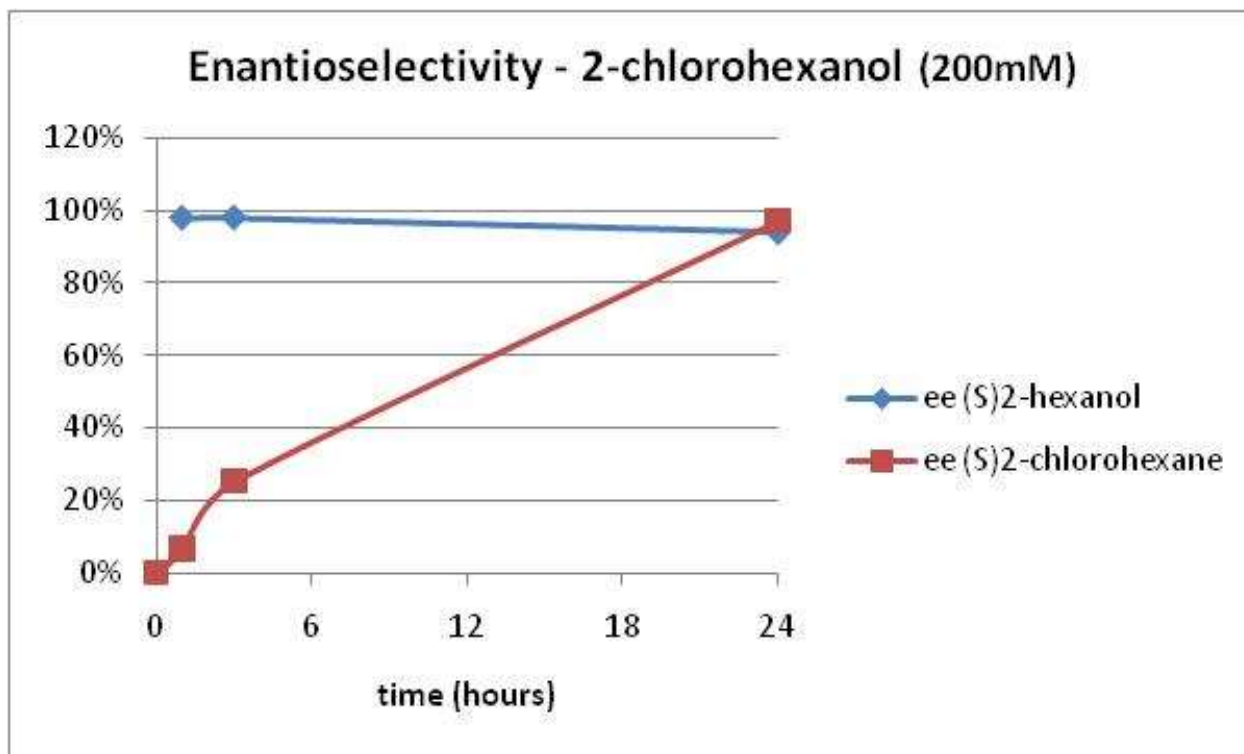
1800 strains screened

| Substrate | Incubation time | % conversion |
|---------------------|-----------------|--------------|
| 1-chloropentane | 19h | 100% |
| 1-chlorodecane | 19h | 35% |
| 1-bromopropane | 4h | 100% |
| 1-bromobutane | 4h | 100% |
| 1-bromohexane | 4h | 98% |
| 1-bromodecane | 4h | 97% |
| 1,2-dichloroethane | 48h | 29% |
| 1,5-dichloropentane | 19h | 100% |



Degradation of Chlorinated Alkanes

- Active with a large range of substrates
- Preference for brominated over chlorinated substrates
- Active on dichlorinated substrates



Enantioselectivity for Chlorinated Alcohols

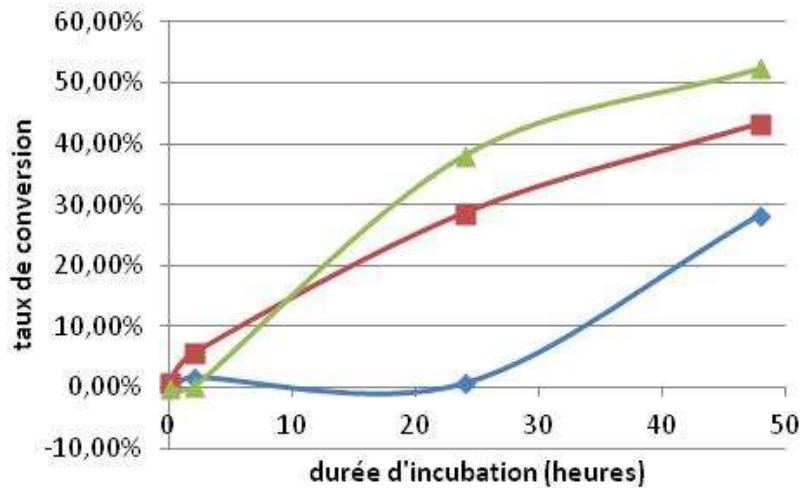
- Subterminal chlorinated substrates
- production of enantiomerically pure alcohols
- Less relevant for bioremediation
- Application in green chemistry is very promising

50mM substrate

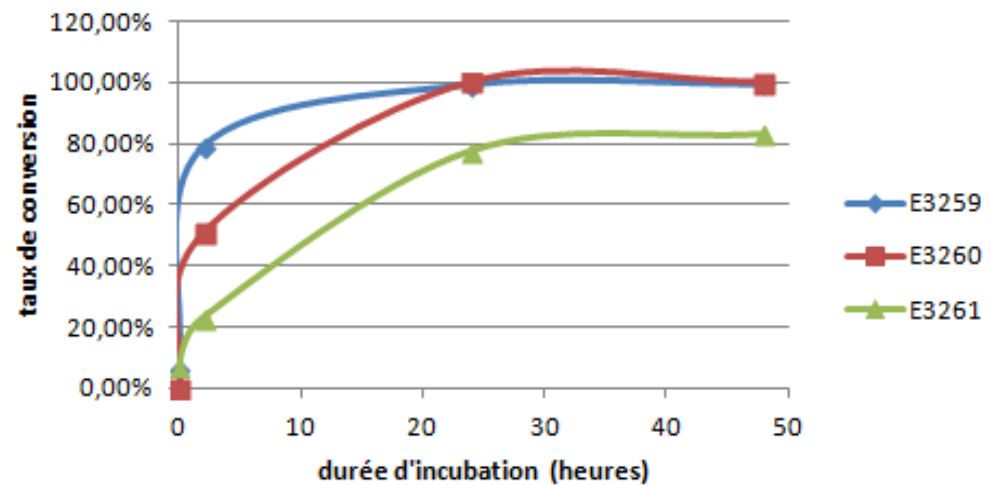
biphasic (buffer / MTBE)

temperature 30°C

Conversion du 2,3-dichloro-1-propène en 2-chloro-3-hydroxy-1-propène



Conversion du 1,3-dichloro-1-propène en 1-chloro-3-hydroxy-1-propène



Screening of the enzymatic toolbox

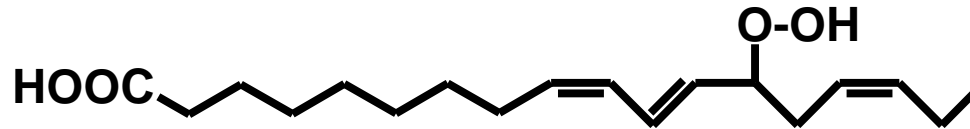


Example : Directed Evolution

Lipids biodegradation
(acyl hydrolases)



O₂ ↓ lipoxygenase



↓ hydroperoxide lyase



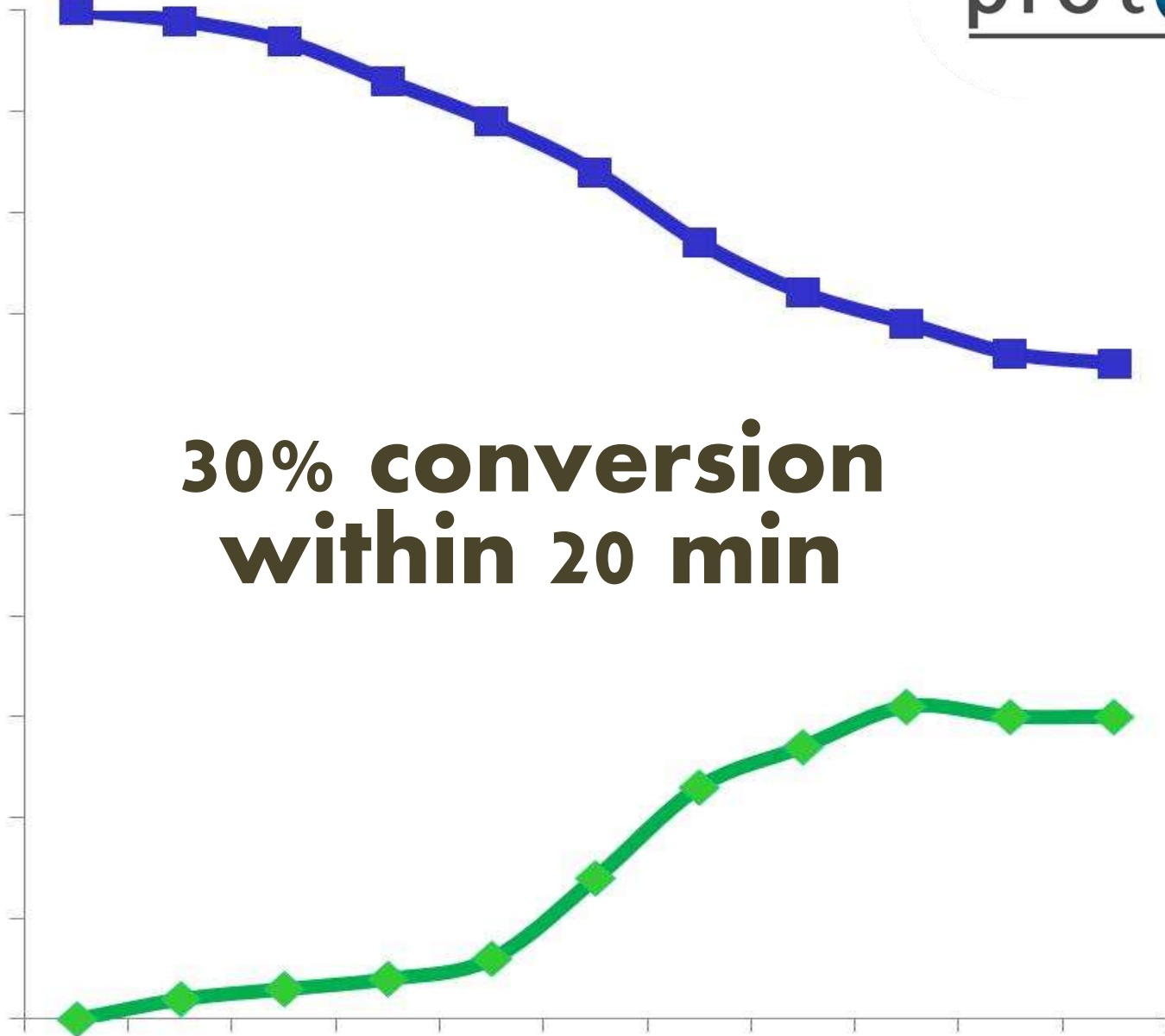
reduction (ADH)
and isomerisation

Green notes

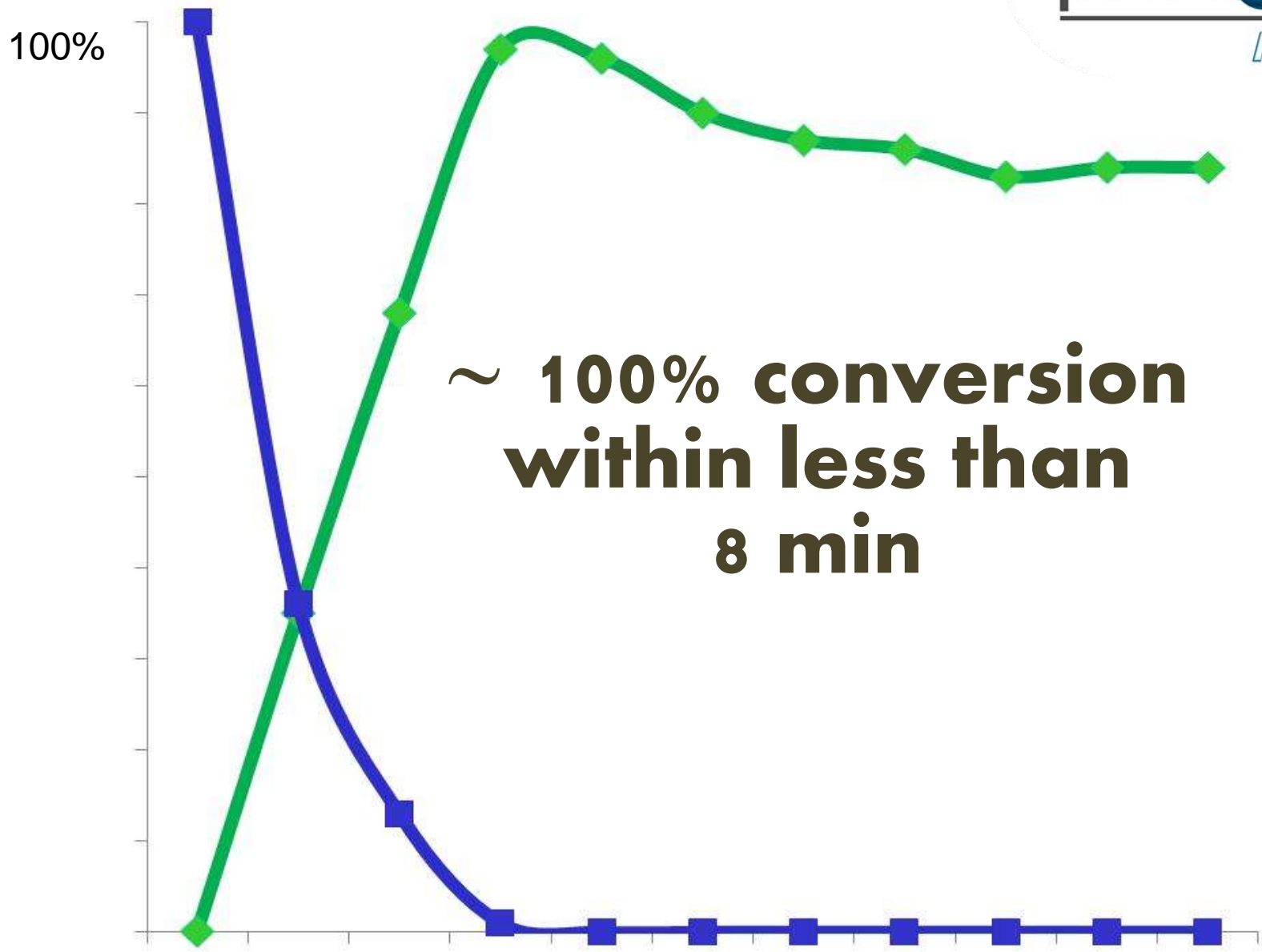
Directed Evolution

100%

**30% conversion
within 20 min**



A collaboration with **Firmenich**



**~ 100% conversion
within less than
8 min**

Green Route to Green Notes

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Pascal Auffray, Laurent Fourage, Denis Wahler.**

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More About This Work



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