



*Journées Techniques*  
*SCALE-UP 2017*  
*Adebiotech – Pôle IAR*

**Aides à la conception du procédé et  
incertitudes (design et aspect économique),  
cahier des charges et données critiques**

**21 novembre 2017, Romainville**

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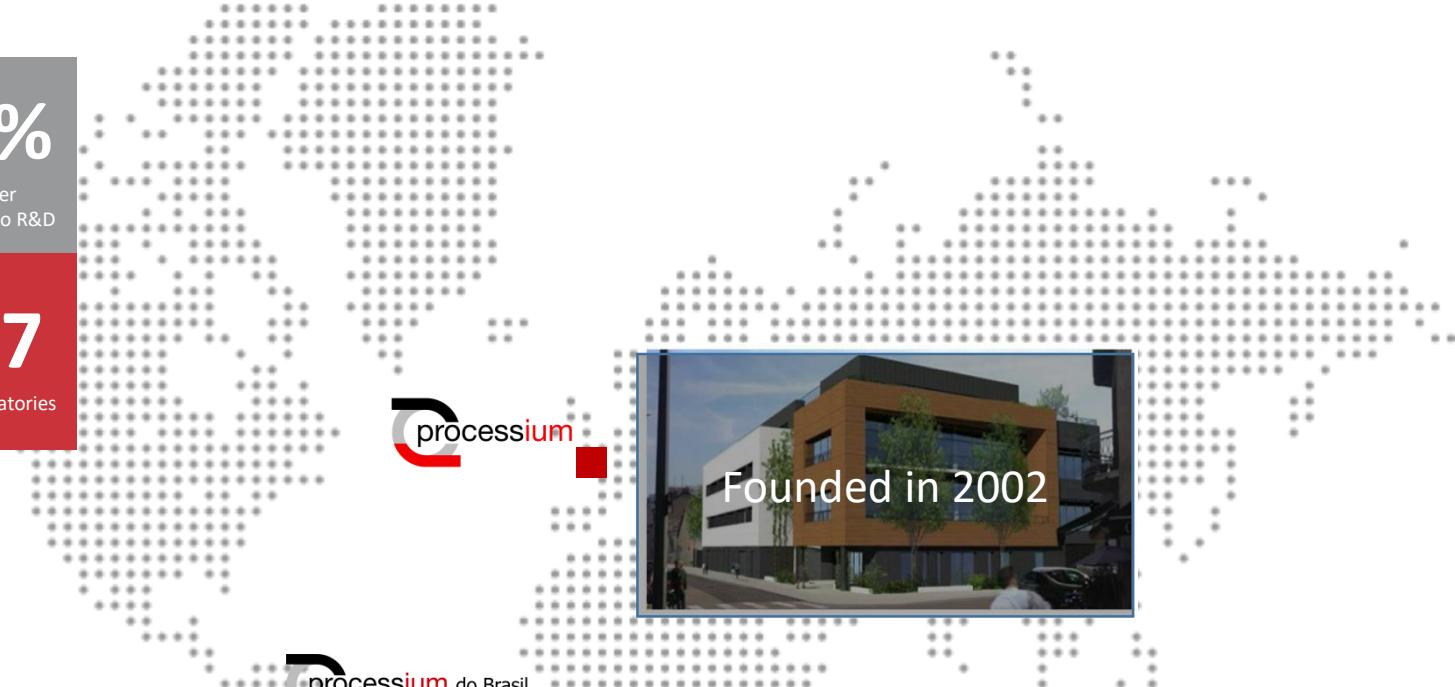
# 1. Processium



# Processium General Overview

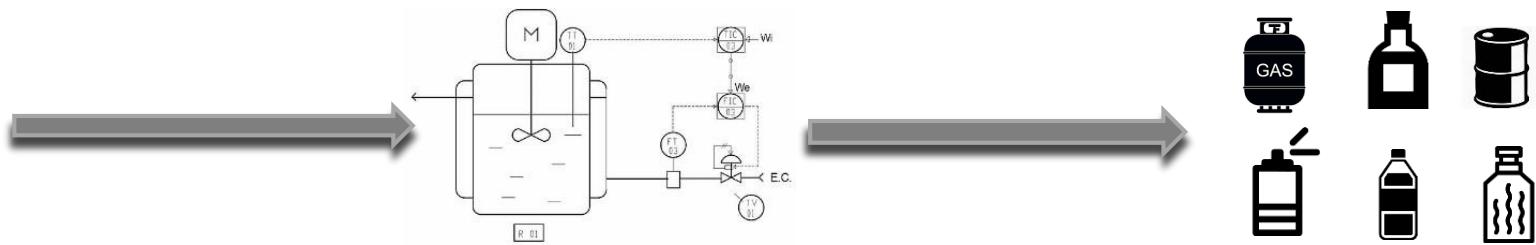
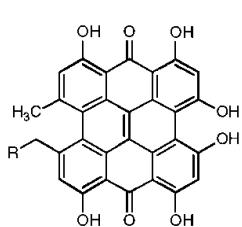
## Who we are

<b>25%</b> PhDs	<b>15%</b> turnover dedicated to R&D
<b>20%</b> turnover outside France	<b>917</b> m <sup>2</sup> of laboratories



# Processium General Overview

## What we do



## **CLIENT PROJECT DEVELOPMENT**

Research

## Process Conception

## Process Scale-up

## Industrialization & production

## Commercialization

# PROCESSIUM SERVICES

**PROCESSIUM  
SERVICES**



# Processium General Overview

## How we work

Laboratory dedicated  
to products



Analysis and  
measurements



Laboratory dedicated  
to processes



Process engineering



Scientific computing



# Laboratory equipments

## Liquid separation

### 1- Liquid-Liquid extraction

(continuous and stirred column)

5 to 25 l/hr, up to 150°C



### 2- Continuous distillation

1 to 3 l/hr, 20 to 250°C, 10 to 1,000 mbar



### 3- Thin layer evaporation

(short pass available)

1 to 3 l hr, 20 to 250°C, min 1 mbar or 10-3 mbar for short pass

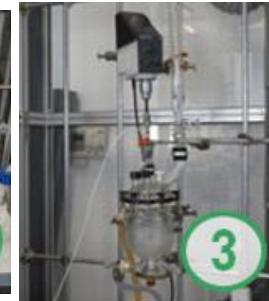


# Laboratory equipments

## Reaction & Separation

### 1- Tangential filtration (ceramic)

MF, UF – Max 4 bar



### 2- Ball mills

### 3- Batch reactor

1 bar, up to 200°C



### 4- Tangential filtration (organic)

MF, UF, NF, RO – Max 60 bar



### 5- Continuous reactor

1 l, micro-wave heating system, up to 40 bar, up to 250°C



### 6- High pressure distillation

1 to 20 bars

Continuous or batch distillation

Reactive distillation

## 2. Feedback from Chemical Industry



# Process Scale-up

## Basic Data Errors

Outcome Variable	Projects without Basic Data Errors	Projects with Basic Data Errors
Percent of successful projects	44	0
Average production months 7-12 vs plan (%)	87	38
Time required for start-up (months / deviation regarding forecast)	7 / +40%	22 / +140%
Slip in execution schedule (%)	16	30



# Process Scale-up

## Key scale-up items

### External: Market, people and environment

#### Critical Scale-up Item

New product ?

Performance product?

New market?

New customers?

New country?

New safety, health, environmental aspects

New suppliers of technology?

New feedstocks? New suppliers?

### Process

#### Critical Scale-up Item

New chemistry?

Novel process?

Novel unit operation? Novel equipment?

Recycle streams?

Solids processing?

New control?

Process operation experience

Phases mass transfer?

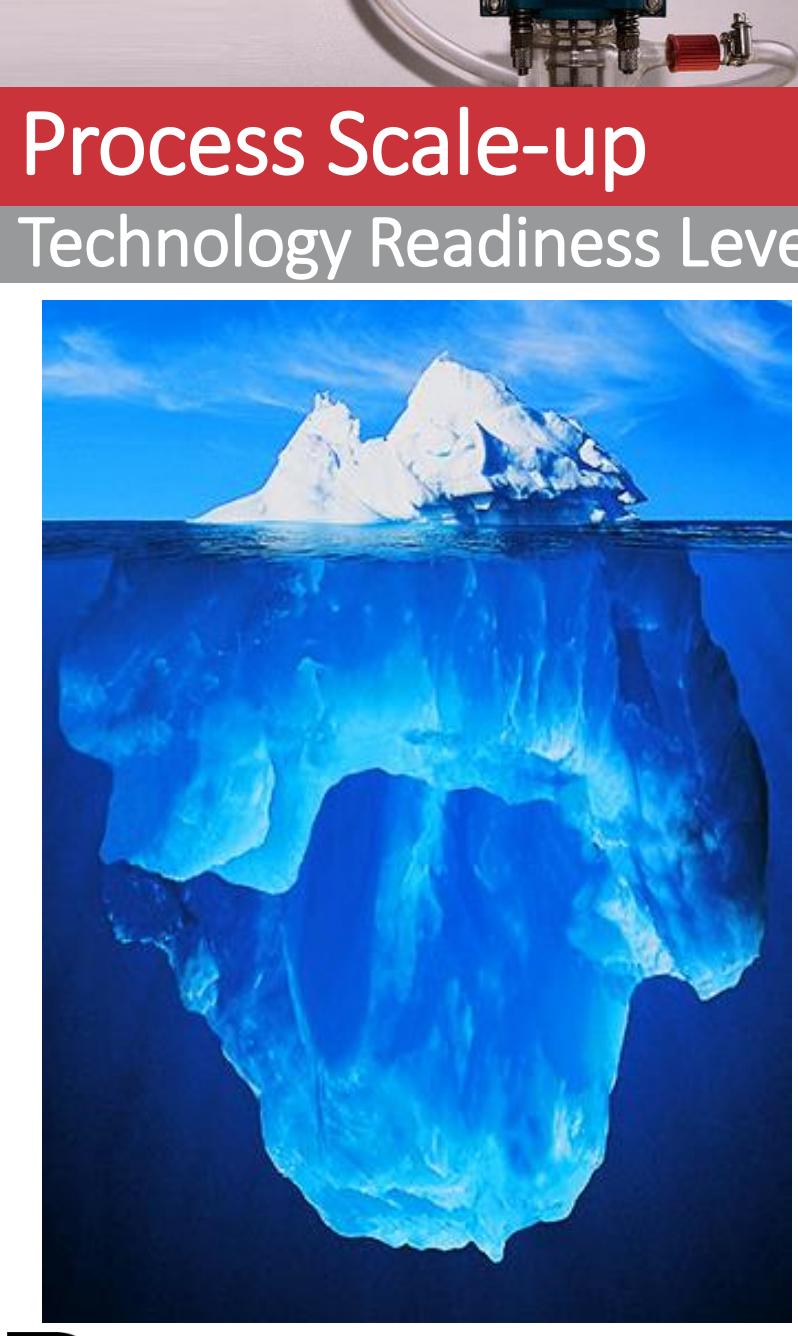
Hydrodynamics scale-up?

G/L or L/L mass transfer?

Residence time distribution?

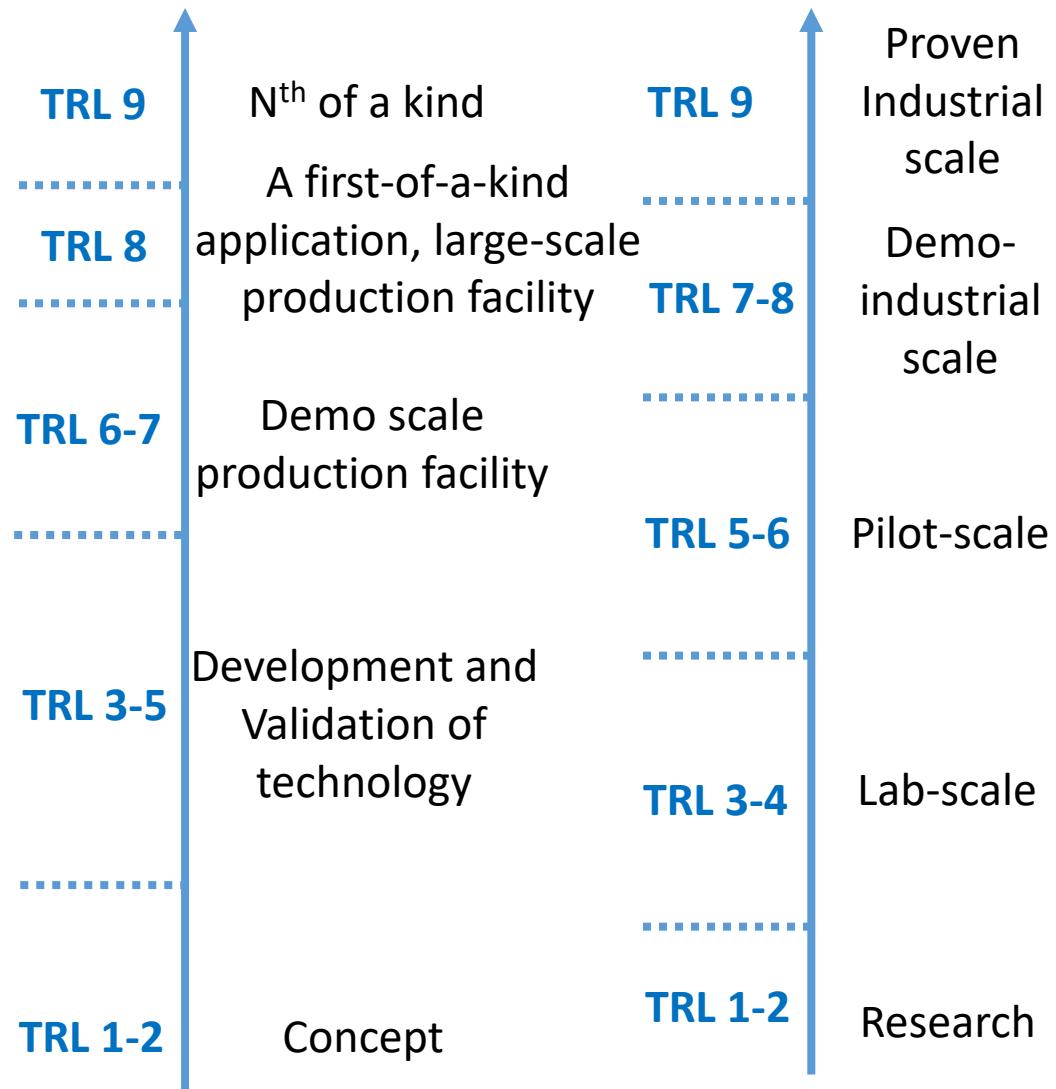
Heat transfer?

New construction materials for streams?



# Process Scale-up

## Technology Readiness Levels (TRL)





# Process Scale-up

## Key steps

	Production rate (kg/h)	Scale Up Factor
Industrial Plant	100-10,000	-
Pilot Plant	1-100	1-10000
Miniplant	0.1-1	100-100,000



# Process Scale-up

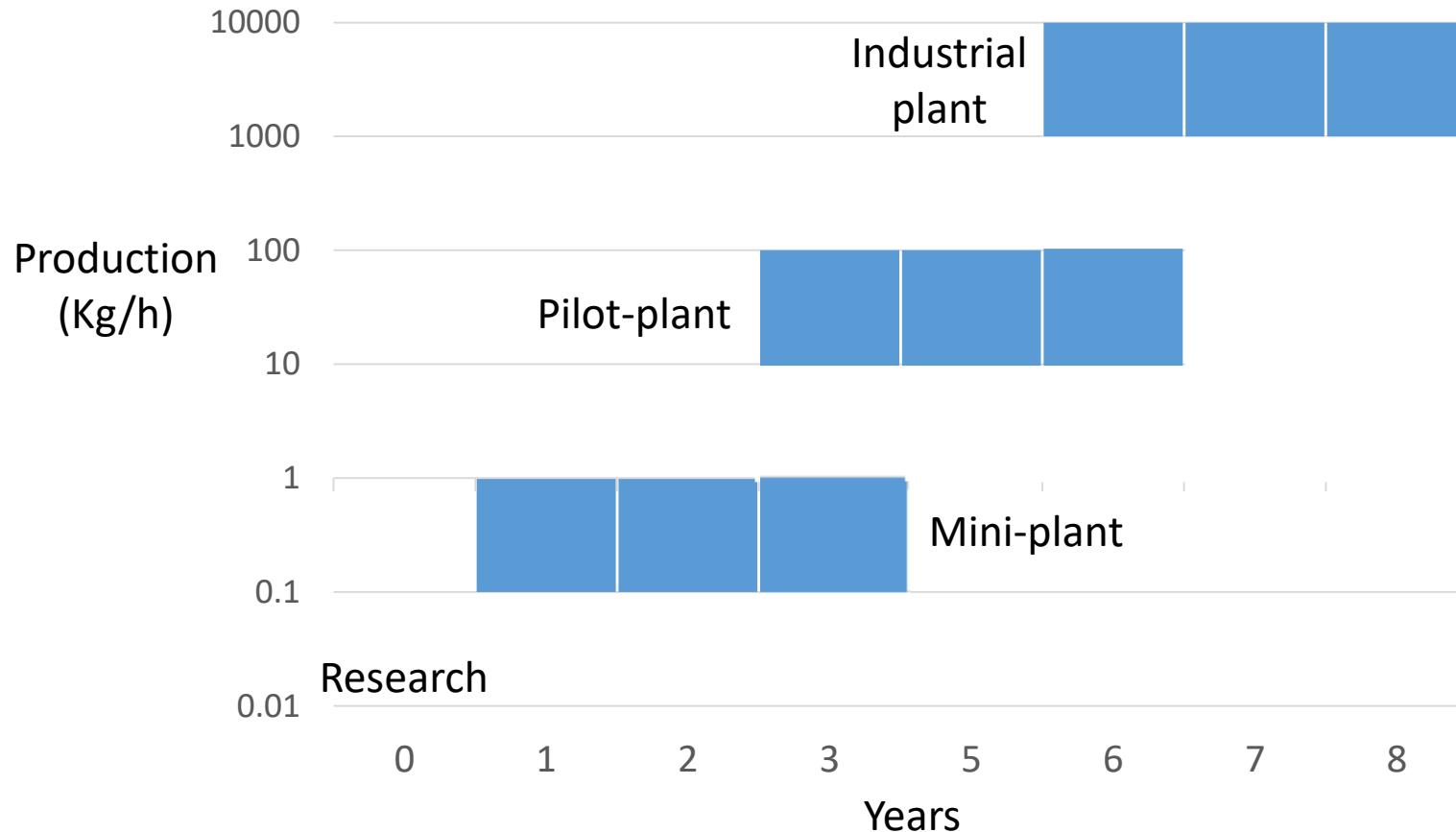
## Scale-up factors

Process Step	Maximum Scale-Up value
<b>Reactors</b>	
Multi-tubular & adiabatic Fixed-Bed reactor	> 10 000 (50 000 achieved)
Homogeneous Tube and Stirred Tank	> 10 000
Bubble Column	< 1 000
Gas-solid Fluidized bed	50 - 100
<b>Separation processes</b>	
Distillation and Rectification	1 000 – 50 000
Absorption	1 000 – 50 000
Extraction	500 – 1 000
Drying	20 - 50
Crystallisation	20 - 30



# Process Scale-up

## Time scale



### 3. Bioprocess scale-up



# Industrial bioprocess scale-up

## A case study : 1,4 BDO

### 1,4-Butanediol: 5 years from concept to commercial (0.5X 1,3-PDO)

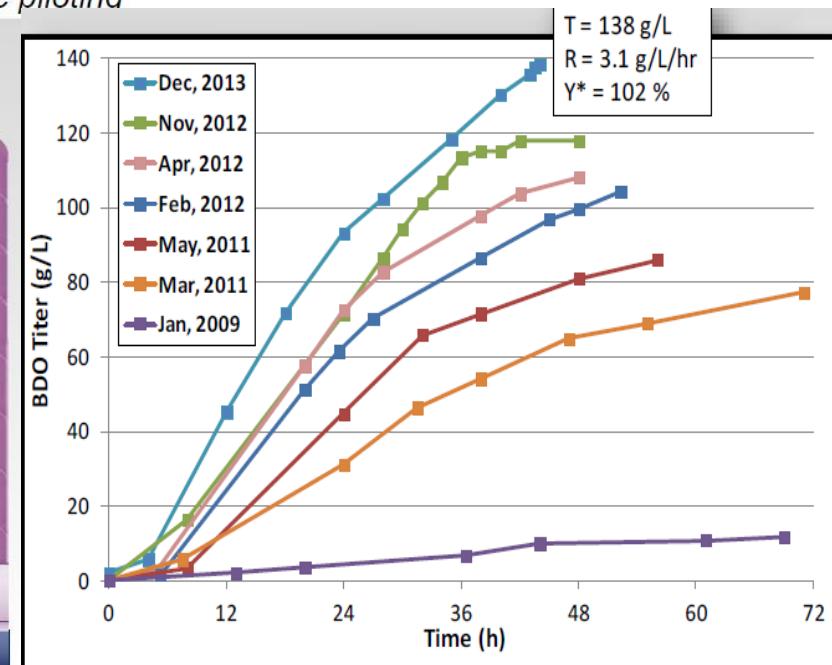
*Process and product validation at commercial scale following extensive pilotina*

#### Process and Product Validation at Commercial Scale

- 50 fermentations run at commercial scale with integrated continuous downstream processing
- **No surprises** - consistent performance, all scales
- 1<sup>st</sup> mass production and sale of an established bulk industrial petrochemical by fermentation
- 2013 Kirkpatrick Chemical Engineering Achievement Award



Scale-Up Partners:





# Industrial bioprocess scale-up

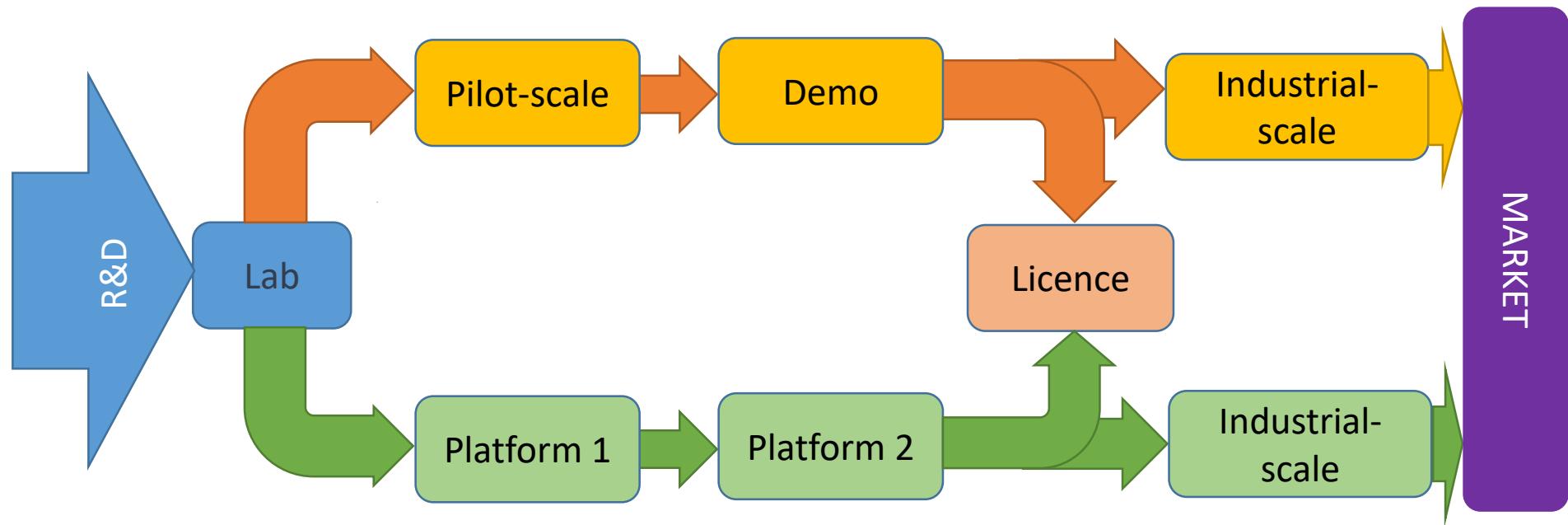
## Biotech vs Chemicals

- Microbes
- Aseptic operation of fermenters
- Control of the fermentation process
- Natural feedstocks
- Low product concentration in aqueous phase
- Selectivity
- Impurities



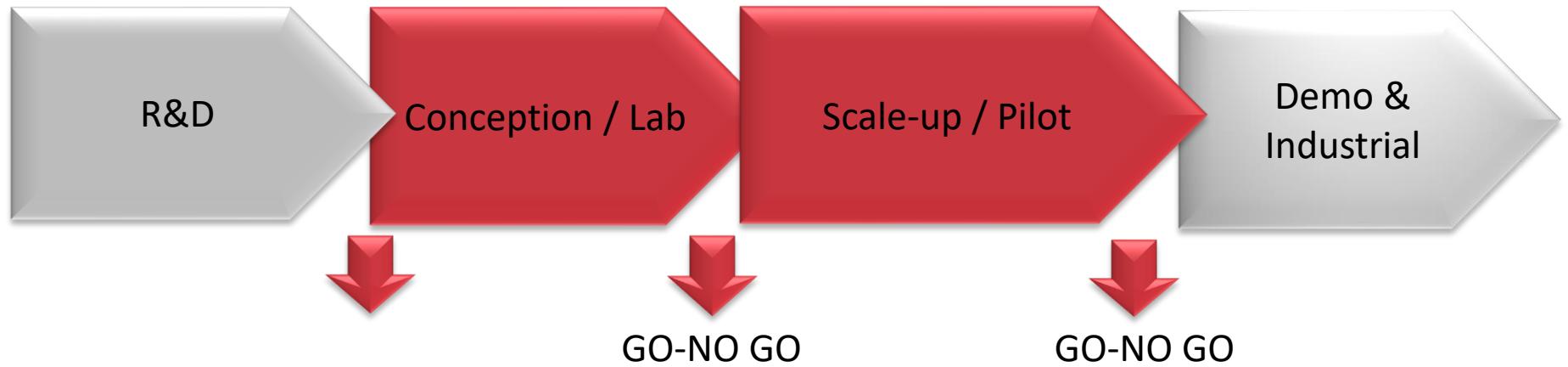
# Product & Process Scale-Up

Two main routes for Process/Product Scale-Up



# Industrial bioprocess scale-up

## Main steps of the conception and scale-up stages



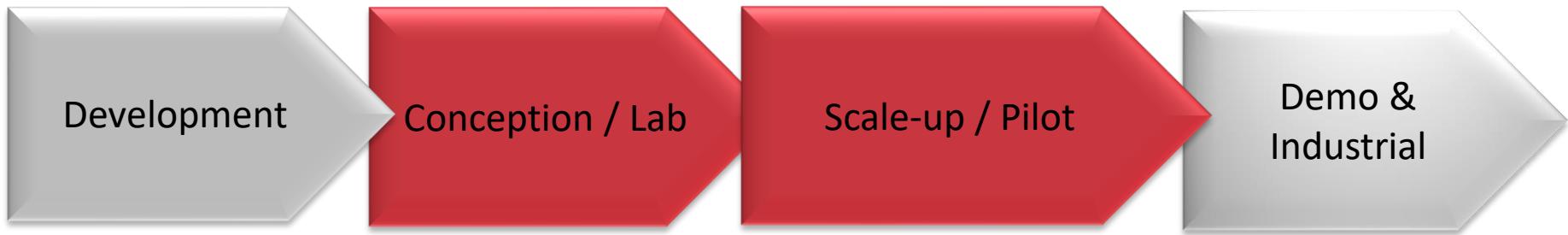
### Deliverables

- Either process book of the demo plant and V1 of the process book of the target industrial unit
- Or update production package
- Risk analysis



# Industrial bioprocess scale-up

## Main results of the conception and scale-up stages



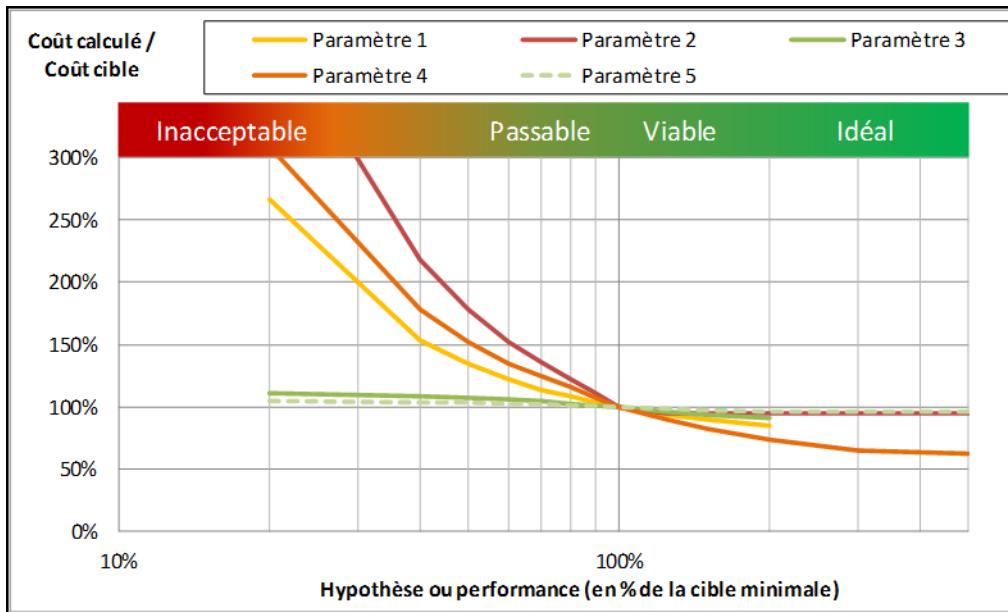
- Process flowsheet
- Mass & heating balances
- Operating conditions
- Sizing of unit operations
- First optimization of the process regarding
  - CAPEX/OPEX
  - Product quality, productivity, yields
  - Environmental impact



## 4. Case studies

# Case study 1

## Preliminary Techno-economical study



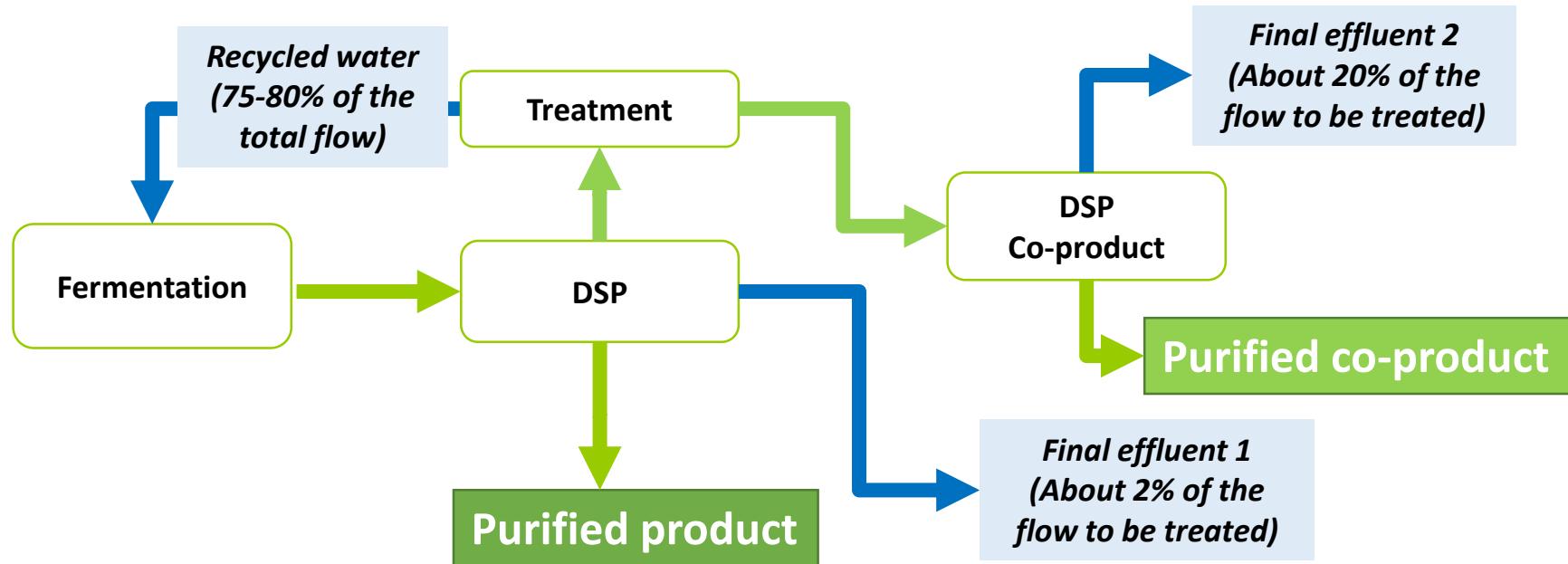
### Results

- First technical and economical estimation of the project
- Fermentation development defined in connection with purification and separation requirements

# Case study 2

## Process development

### Joint development of DSP / USP



- Minimization by fermentation of the most penalizing impurity in purification
- Valorization of a co-product: economy of work in fermentation
- Recycling water with sufficient quality for fermentation



# Case study 3

## Feedstock impact

- One of the key step in the process is to separate the noble product from the salts and sugars
- Development of a DSP with 1G sugars :
  - PCD A (1G): Performance and robustness. Yield 97%
  - PCD B (1G): Low performance, and limited robustness due to fouling problems. Yield  $\approx$  80%
- Development of the same DSP with 2G sugars :
  - PCD A (2G): Low performance and limited robustness
  - PCD B (2G): Good performance, and robustness. Yield 95%



→ Necessity to have a global approach : raw material/USP/DSP



# Case study 4

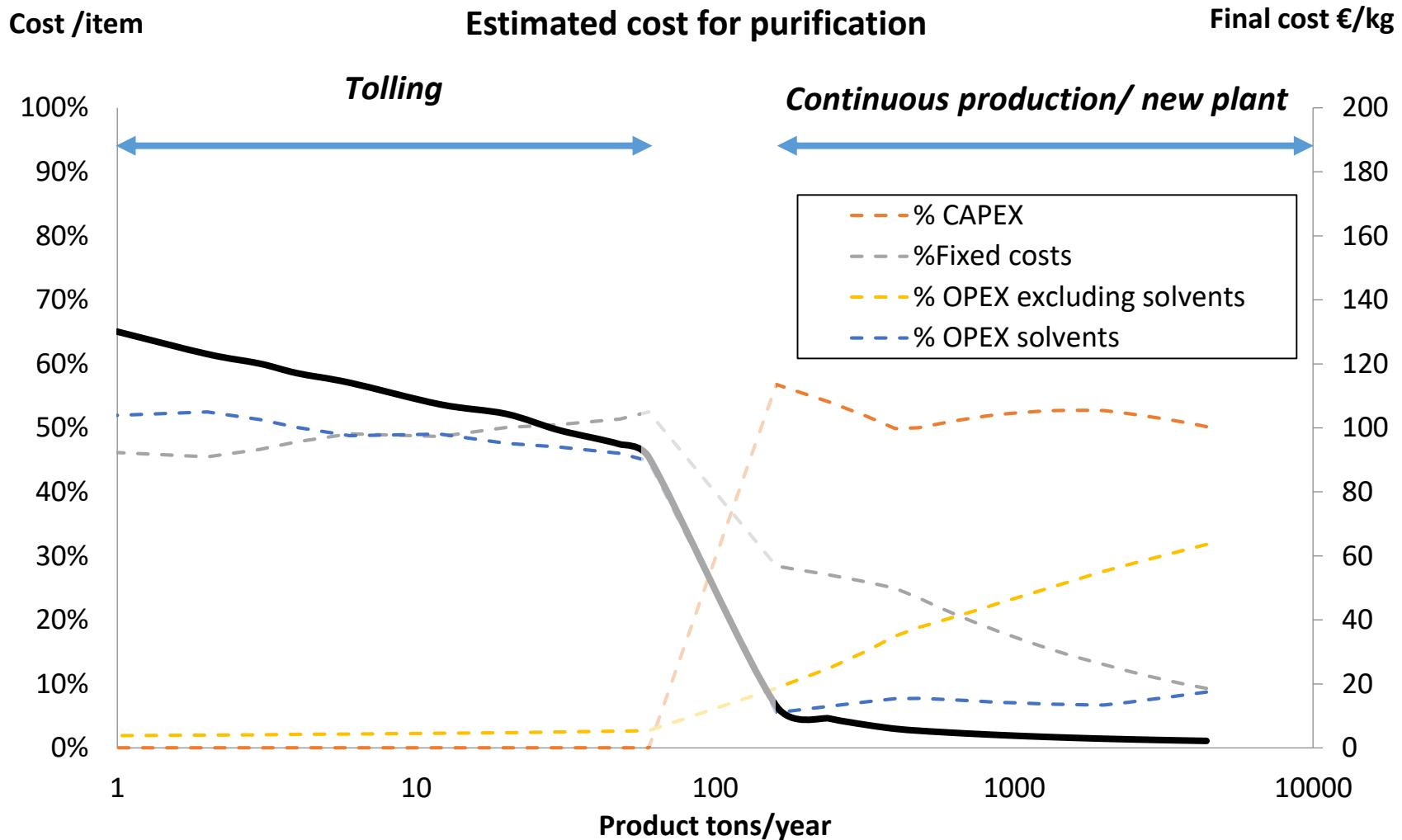
## Downstream process optimization

Estimation for a 30 KT/year plant

	Reference solution	Energy integration solution	Difference
Vapor consumption t/h	44.5	12.5	-72%
Cooling water Consumption m <sup>3</sup> /an	2630	730	-73%
Power consumption (compressor) GWh/an	-	13.56	-
Number of exchangers	16	18	+2
OPEX k€/an	8480	3173	-62%
K€/ t final product	0.28	0.10	-62%
Return on investment time		< 2 ans	

# Case study 5

## Toll manufacturing



# 5. Conclusions



# Conclusions

## Process scale-up

- Objectives
  - To limit the risks
  - To save time
  - To manage CAPEX/OPEX
- How?
  - Identifying market targets
  - Implementing adapted industrial methodologies
  - Looking for key partners
  - Working in parallel, anticipate
  - Defining GO/NO-GO milestones



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