#### △♪ナミキ商事株式会社

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# **Flow Chemistry**

**550**+ projects

50+ reaction types

kilo to metric ton scale



### 2023 ACS GCIPR CMO Excellence in Green Chemistry Award Winner

The award-winning project implemented innovative continuous flow process

## Application in safer, more stable, higher-yield processes

- High temperature/pressure
- Highly energetic
- Cryogenic
- Highly reactive and air-sensitive
- Toxic and/or stinky agents
- Unstable intermediates
- Oxidation and/or ozonization
- Diazotization

- Sulfonation
- Esterification
- Halogenation
- Reduction

#### **Reactors**

■ Single-tube ■ Static mixer

Multi-tube

- - Dynamic tubular reactor Photo-flow reactor
- Fixed/micropacked bed Electrochemistry reactor CSTR

# Cases

## **Cryogenic reaction**

Comparison	Batch	Flow
Feasibility of scaling up	×	$\checkmark$
Temperature	-70 to -60°C	-40 to 10 °C
Yield	N/A	84%
Scaling-up risk	High	Low

Completed 260 kg product with 240 mL continuous flow reactor in 30 hours

### **Diazotization**

1	Comparison	Batch	Flow
ľ	Feasibility of scaling up	×	√
	Temperature	N/A	5 to 10 °C
	Yield	N/A	80 - 85%
	Scaling-up risk	High	Low

Completed 200 kg product with a set of 100 mL continuous flow reactor in 2-3 days

#### **Nitration**

Batch	Flow
×	√
20 - 30 °C	30 - 60 °C
N/A	90 - 93%
High	Low
Low	High
	× 20 - 30 °C N/A High

Over 300 kg of product completed with an integrated continuous tubing reactor

# **High temperature**



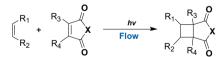
Comparison	Batch	Flow
Feasibility of scaling up	×	√
Temperature	200 °C	220 - 250 °C
Yield	N/A	>94%
Scaling-up risk	High	Low
Automatic leve	Diphenyl ether (BP: 258 °C)	Toluene (BP: 110℃)
Over 100 kg of product completed		

### Oxidation



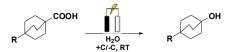
Comparison	Batch	Flow
PMI	15	7
Time	> 4 h	10 min
Yield	88 - 90%	95%
Complexity of work-up	High	Low
	Over 100 kg of product completed	

# **Photocatalytic reaction**



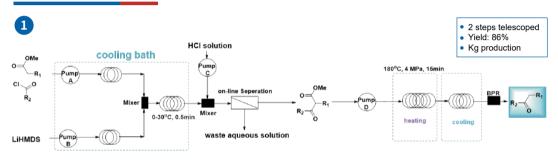
Comparison	Batch	Flow
Feasibility of scaling up	×	$\checkmark$
Time	30 h	40 - 50 min
Light source	Medium pressure mercury lamp	365 nm LED
Scaling-up risk	High	Low
Over 1 MT of product completed		

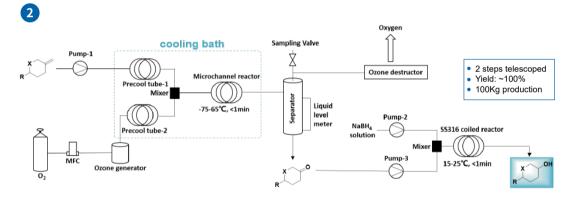
## **Electrocatalytic reaction**



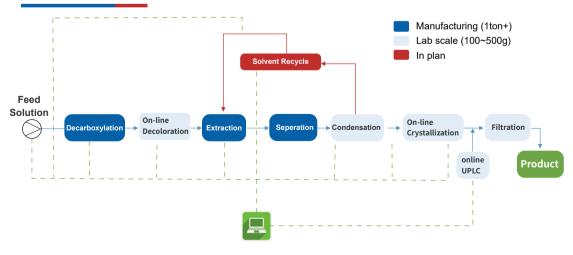
Entry	General Process	Electrochemical Process
Step	3	1
Yield	45%	67%
PMI	135	73
Cost of Material	> 30% cost reduction	
Self-made equipment, Kg scale preparation		

## **Telescoped flow cases**





## **End-to-end solution**







Continuous Manufacturing GMP Workshop at PharmaBlock Zhejiang

【お問い合わせ】

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