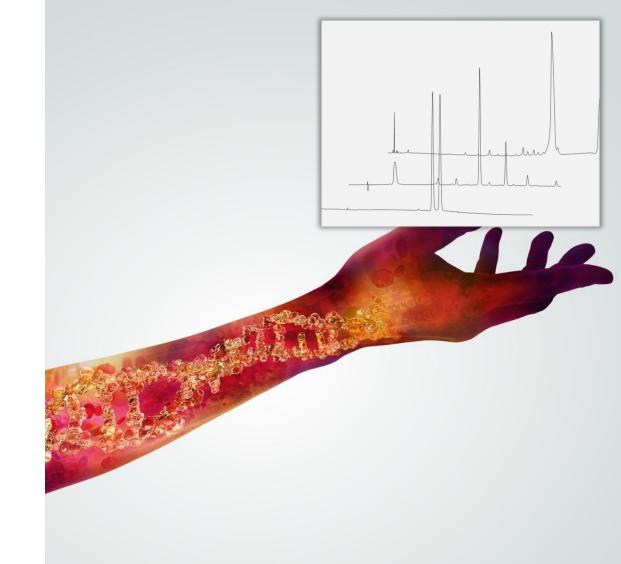


Strategy for chiral SFC separations at AstraZeneca

Anna Jonson

Separation Science Laboratory, Early Chemical Development, Pharmaceutical Sciences, Gothenburg



AstraZeneca Gothenburg



The Gothenburg site in Sweden is one of AstraZeneca's six strategic research and development centers.

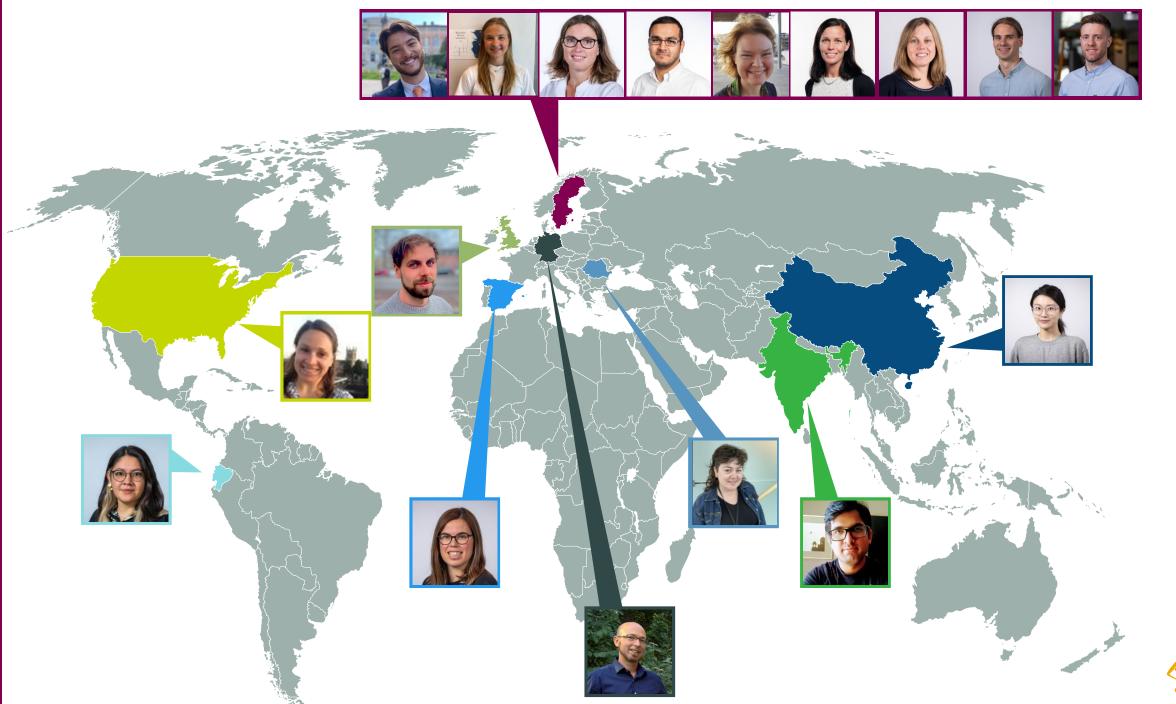
~3000 employees, from more than 50 nationalities.

Covering early R&D to GMP manufacturing

Separation Science Laboratory (SSL):

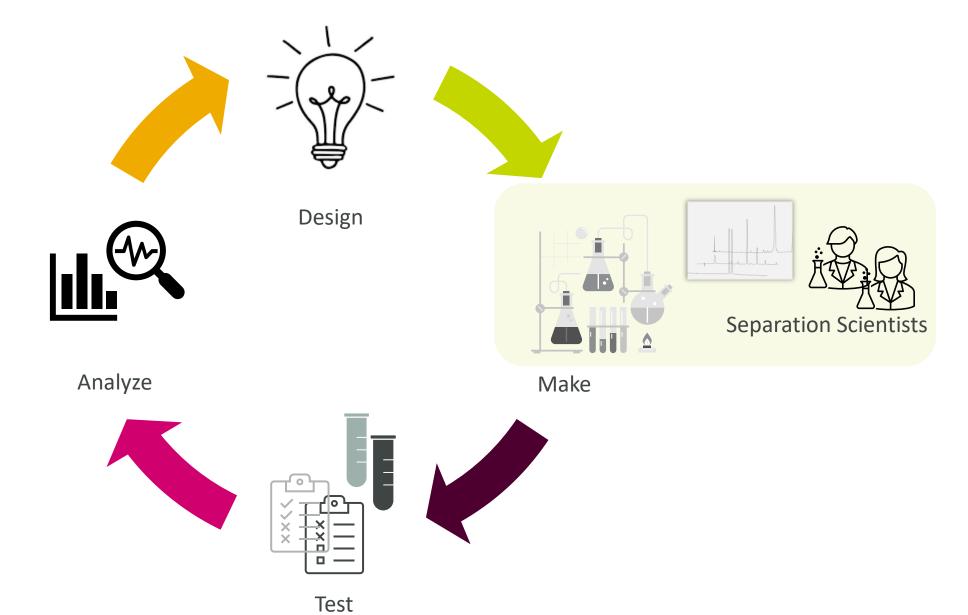
- Support projects from early to late phase
- Support from small molecules up to large molecules and New Modalities
- From mg to kg





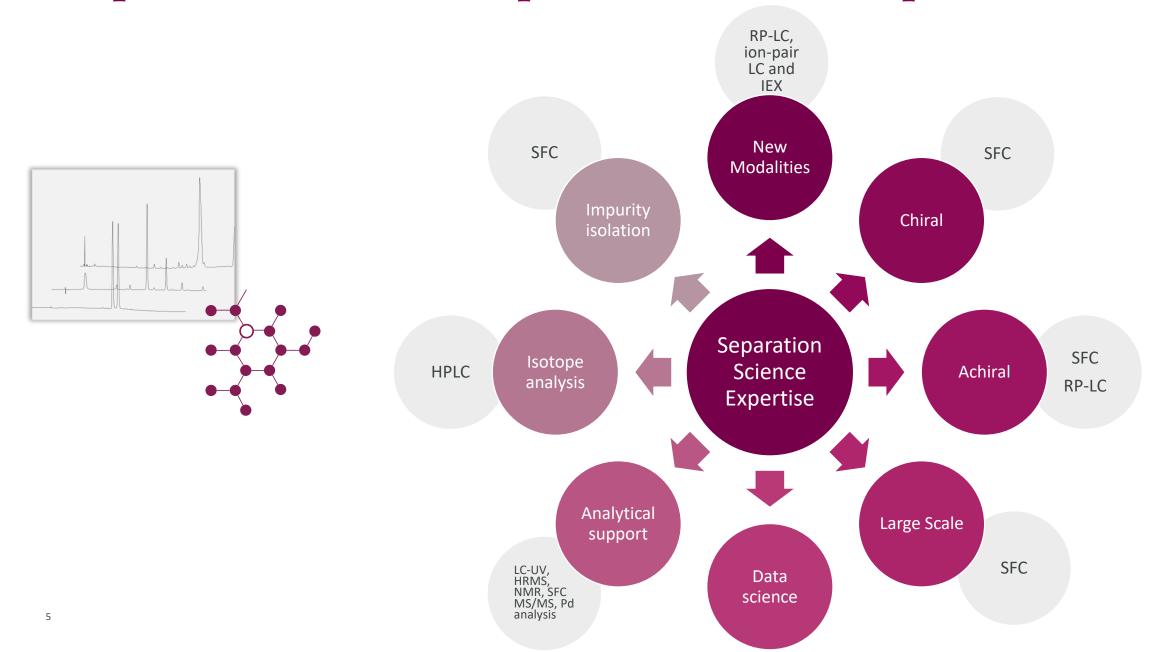


Drug Development Cycle – DMTA Cycle



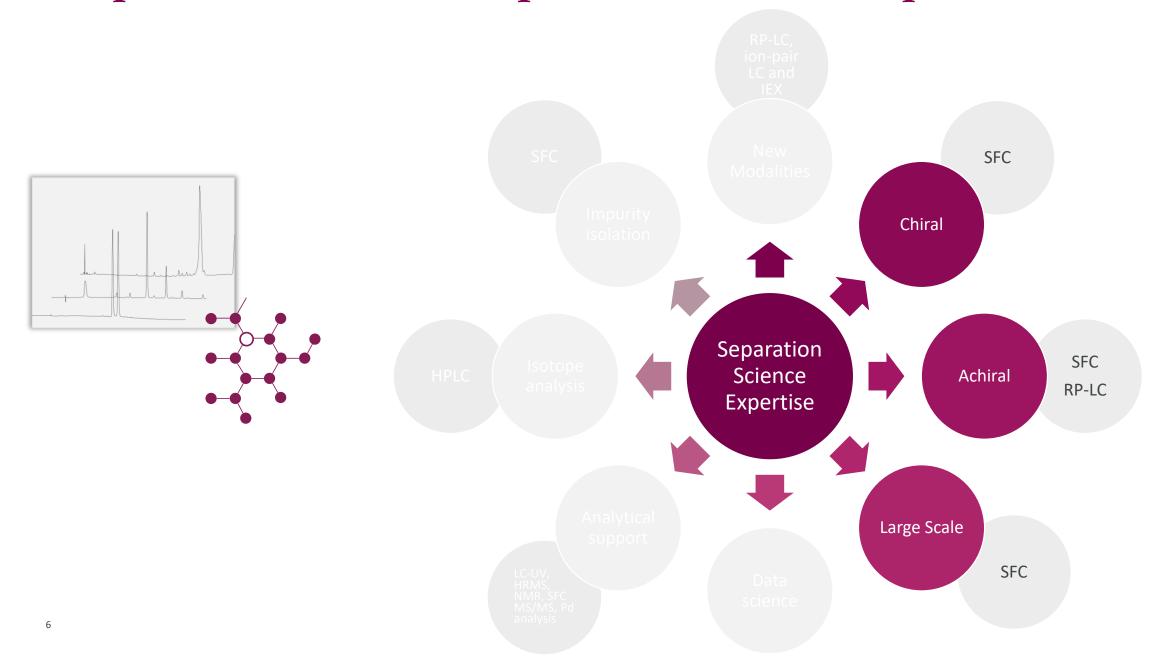


Separation Science Expertise and Techniques



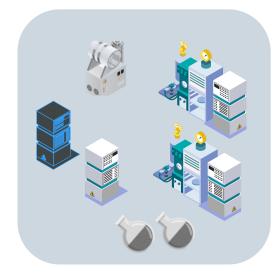


Separation Science Expertise and Techniques

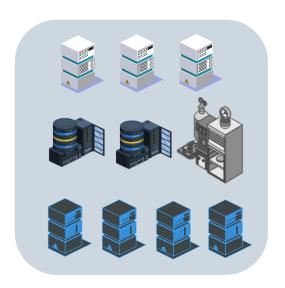




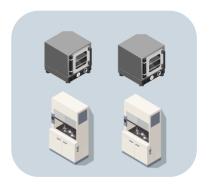
Laboratory Instrumentation Footprint



Large Scale Lab







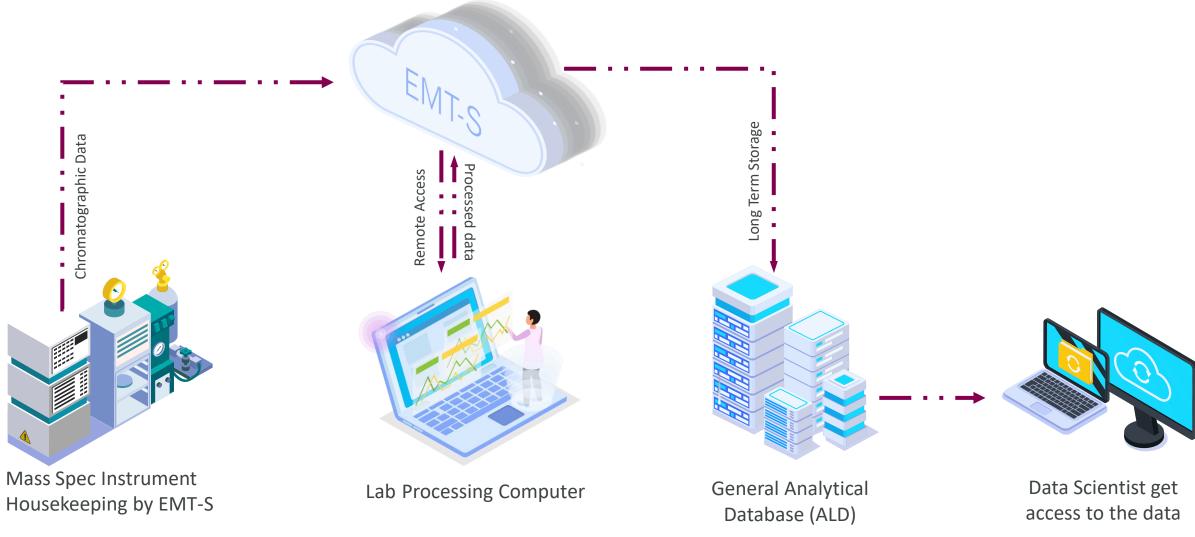
Drying Module





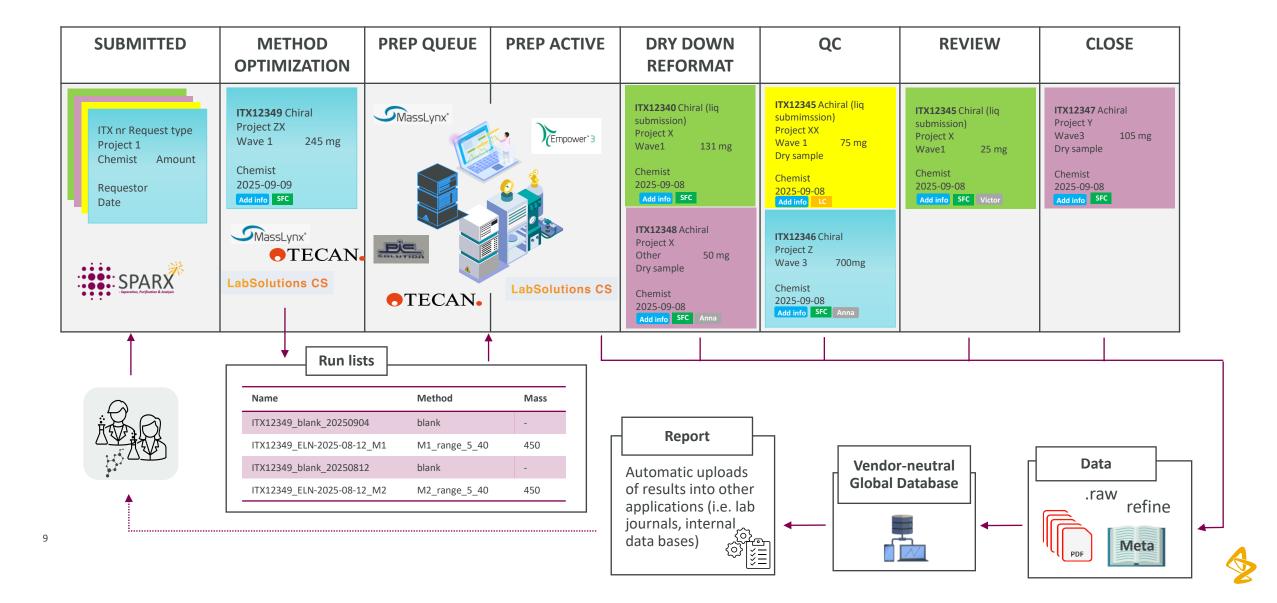


Data Management

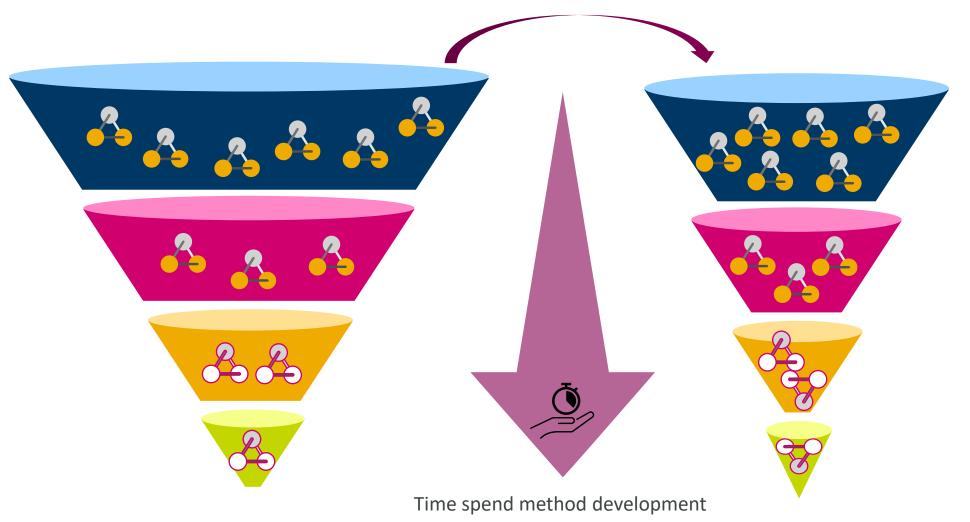




Tracking Multiple Molecules and Data Simultaneously



The method development process



- Digitalization
- Automation
- Prediction
- Save time
- Save resources
- More sustainable operations



Analytical Chiral Screen Setup

Instrument





A: IPA/NH₃ 20 mM B: EtOH/NH₃ 20 mM C: MeOH/NH₃ 20 mM

D: ACN/NH₃ 20 mM



Oven 1

Chiralpak IA

Chiralpak IB

Chiralpak IC

Chiralpak ID

Chiralpak IE

Chiralpak IF

Oven 2

Lux i-A3 (IG)

Chiralpak IH

Chiralpak IJ

Chiralpak IK

Chiralpak IM

(S,S) Whelk-O1

Oven 3

Lux A1

Chiralpak IN

Lux C1

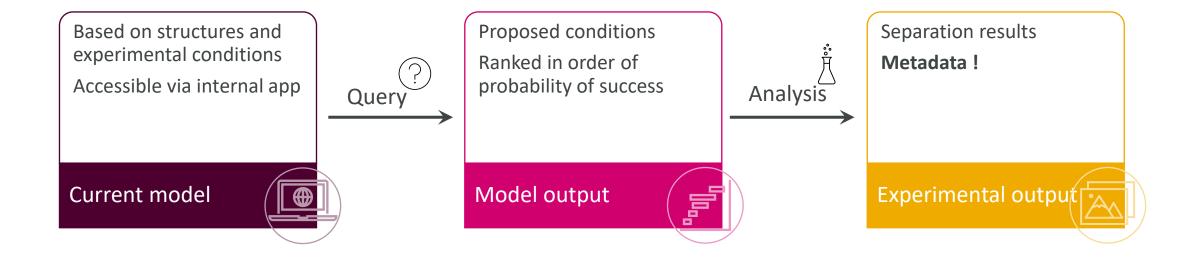
Lux C2

Lux C3

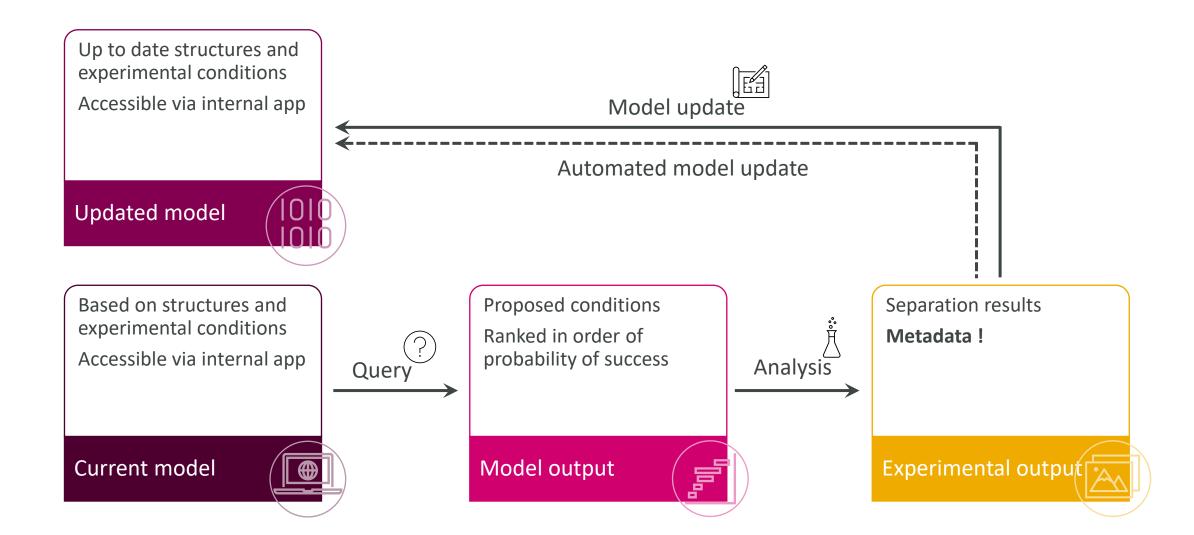
Lux C4



Analytical Chiral Prediction









Green Aspects in our Analytical Chromatographic Process

Minimizing Instrument Time and Electricity Usage

Enhancing Efficiency



Decreasing Solvent Usage and chemical waste

Optimizing workflows and adopting sustainable practices, to decrease our carbon footprint





Analytical aspects

Preparative aspects

Separation results

Metadata!

Experimental output

Experimental conditions

- Chromatography
- Loadability?

Upscale



Run the prep

Deliver material at required purity

Additional useful information

Purification outcome

Column behavior

Compound behavior

Loadability

Experience





Approach for Preparative Chromatographic Setup

Instrument



Shimadzu Nexera Prep

Injection: vial 4 ml

Fractionation:

Open tubes (25 ml)

Duran bottles (500 ml)

Flow: 70-130 ml/min



PIC 600

Injection: 500ml bottle -10L

tank

Fractionation:

5L bottles, 25 L or 50 L tanks

Flow: Up to 600 g/min

Connected to continuously

rotavapors

CO₂ infrastructure



- 6 m³ tank
- Recycled from waste gas in other manufacturing industries



Solvents



Acetonitrile

Methanol

Ethanol

2-propanol

Ethyl acetate

Methyl tertbutyl ether

Tetrahydrofuran

Heptane

Water



Strategic Approach to Recycling in Large Scale



Solvent recycling

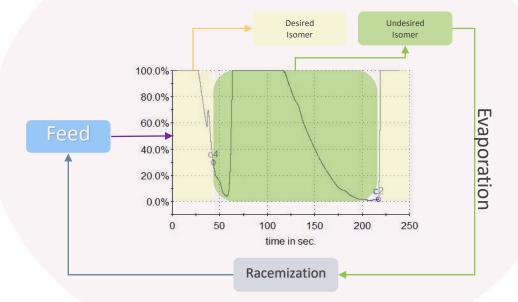
Modifier recycled from evaporation while purifying



Compound "recycling"

Undesired isomer recycled when possible

Sustainability & Resource Optimization





Carbon dioxide recycling

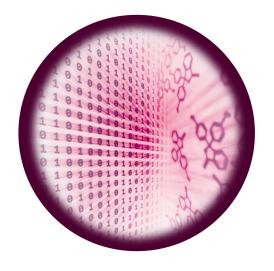


Stationary phase reuse/recycling



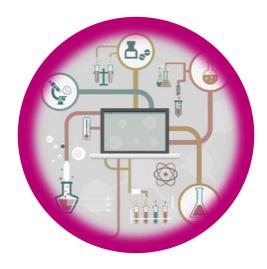
Future

Data-Driven Method Development & Analysis



Prediction-driven experiments

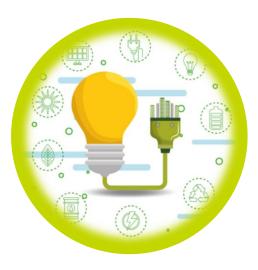
Continuous Innovation



Automation integrated across all workflow stages

Keep the human in the loop

Commitment to Sustainability



Optimize resource use, minimize waste, and make green choices



Thank You

Questions?





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