# APELOA Pharmaceutical CDMO

Dec 2024

**Technology for Health** 

#### **APELOA Overview**



At Apeloa, part of the Hengdian Group, one of China's largest private enterprises, we are committed to delivering high-quality APIs, Intermediates and Regulatory Starting Materials (RSMs) from R&D to GMP commercial-scale production



\$1.7 B<sub>revenue</sub>

2024 milestone

7,000+

**Employees** 

36 years

Founded

manufacturing sites

cGMP, ISO9001, ISO14001, OSHAS18001

sites passed



sites passed



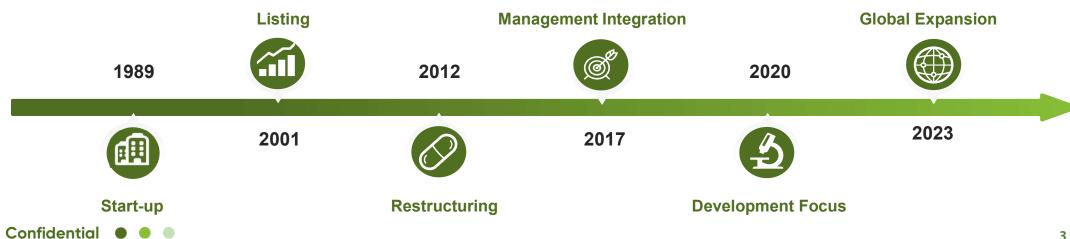
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# What APELOA is doing



API & Intermediates



CDMO



**FDF** 



Aesthetic & Care Ingredients



# **CDMO Manufacturing sites**







Shandong Tospo (ISO)

RSM, Int. Area: 300,000 m<sup>2</sup> Capacity: 642 m<sup>3</sup>



# **Shandong Hanxing** (ISO)

RSM

Area: 533,330 m<sup>2</sup> Capacity: 5,378 m<sup>3</sup>



# Anhui Apeloa Biotech (cGMP)

RSM, Int., API Area: 583,000 m<sup>2</sup> Capacity: 1920 m<sup>3</sup>



# Zhejiang Biotech (cGMP)

RSM, Int., API Area: 141,000 m<sup>2</sup> Capacity: 3232 m<sup>3</sup>



#### Zhejiang Apeloa Jiayuan (cGMP)

RSM, Int., API Area: 250,533 m<sup>2</sup> Capacity: 4364 m<sup>3</sup>



#### Zhejiang Apeloa Tospo (cGMP)

RSM, Int., API Area: 166,500 m² Capacity: 632 m³



#### Zhejiang Apeloa Kangyu (cGMP)

RSM, Int., API Area: 408,700 m<sup>2</sup> Capacity: 1,449 m<sup>3</sup>

## **CDMO R&D Centers**





#### Hengdian Zhejiang - CHINA

80,000 sq ft 250+ chemists Lead optimization / Process / CMC



# Pudong Shanghai - CHINA

64,000 sq ft 300+ chemists **Screening / Library / Process** 



#### **Boston - USA**

17,000 sq ft 30+ chemists **Tech Development Platform** 



Commercial

Phase III

Phase II

Phase I

Preclinical

**Discovery** 

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# Capacity



# **Chemical Synthesis**

Total Capacity 11,000 m<sup>3</sup> Reactor Size 50 - 45,000 L

APIs 4,000 MT / year Intermediates 38,000 MT / year

## **Bio-Production**

Total Capacity 6,570 m<sup>3</sup>

Reactor Size 50 - 120,000 L

APIs 2,000 MT / year

Intermediates 2,000 MT / year

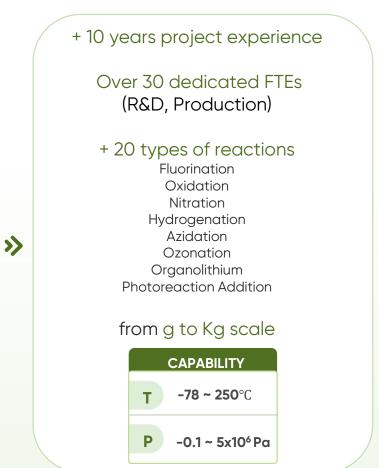
# **Technology Platform: FLOW CHEMISTRY**

CORNING

# Full life-cycle services from lab to production











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# Technology Platform: SYNTHETIC BIOLOGY & ENZYME CATALYSIS APFIC



#### Strain platform

Strain improvement and optimization

#### Fermentation

Fermentation process optimization

#### Separation

DSP optimization

#### Biotransformation

Enzyme screening, evolution and process optimization

**Process development** (1-3 months)

Capabilities

Pilot test (1-2 months) Manufacture (1 month)





- Gene knock-in/knock-out, gene overexpression
- > Transcription factor, RBS & promoter engineering
- Various microbial chassis, including Escherichia coli. Streptomyces spp., Pichia pastoris & Saccharomyces cerevisiae
- > Strain acclimatization & rejuvenation
- > Strain mutagenesis & screening
- ➤ High-throughput screening

#### Recent case studies

- Figure Gene replacement in a streptomyces to eliminate a prominent impurity in the fermentation process;
- Screen and engineer a strong promoter for a PKS gene cluster in a fermentation strain to enhance the production titre by 20%;
- Overexpression of certain limiting genes to enhance the metabolic flux to a desired product, which increased fermentation titre by 10%;
- Development of CRISPR-based approaches for the construction of plasmid-free recombinant strains.



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# **Technology Platform: HIGH POTENCY APIs**



- Auto temperature and pressure control
  - DCS and dedicated HEPA
- Bay segregation:
  - Suitable containment
  - 6 bays for RSM/Intermediates and APIs
  - Dedicated waste treatment
- 400 m<sup>2</sup> HP lab for trial & testing

## OEB 4

Large reactor area (Exposure value:  $< 0.5 \text{ ug/m}^3$ )

**2,000 L ~ 10,000 L** GL reactors



- Total reactor volume: **75 m³**
- 35 reactors, 3 sets distillation units
- Filter drier: 6 Conical drier: 1
- Prep-HPLC: DAC 100, DAC 150, DAC 300
- Lyophilizer: 0.5 m<sup>2</sup>, 1 m<sup>2</sup>, 5 m<sup>2</sup>
- Jet mill, Hammer mill and Wet mill

#### OEB 5

Small reactor area

(Exposure value:  $< 0.05 \text{ ug/m}^3$ )

20 L ~ 1000 L Glass/GL reactors







# Technology Platforms: TPD/PROTAC – Peptides – ADCs



# TPD/PROTAC

80+ PROTAC focused FTEs

Strong analytical and ADME capability

One-step conjugation with POI Binders

Flexibility with US-China sites

# **ADCs – Payloads & Linkers**

Dedicated team with 30 scientists

Linkers/toxins/payloads preparation under cGMP condition

Flow chemistry process for hazardous reaction

## **Peptides**



**Analysis** 

Dedicated team with 20 scientists

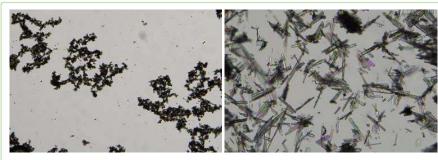
LPPS & SPPS (up to 100 kg scale)

Amino acid analysis and enantiomeric purity

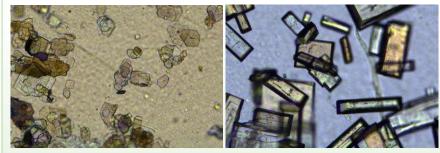


# Technology Platform: CRYSTALLIZATION & SOLID-STATE SCREENING APELOA

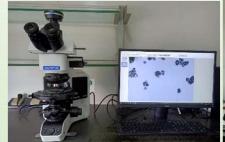


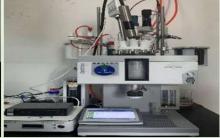


Morphology and impurity control



Polymorph control





# **Platform Function**

Early solid form screening: salt, cocrystal, polymorph, solvate

Pre-formulation study

Fast fit-for-purpose crystallization development, crystallization optimization and scale-up

Chiral resolution via crystallization

Solid-state attributes control

# **ESG System at APELOA Sustainability & Decarbonization**

**Laws & Regulation** 

Under ISO14000, ISO45001, OSHAS18001

**EHS Facility** 

Waste water treatment capability: **50,000** MT/day

RTO for waste gas treatment: **30,000** m<sup>3</sup>/h

Waste-to-energy: **75** MT/day Expense in EHS: >\$80 Million/year

**Audits** 

**12** FSG audits from client in 2023

**248** supplier EHS & quality audits in 2023

**Empowering our community** 

>\$2 Million donation during Covid-19 pandemic

Yearly sponsorships for **>600** elderly people & **>60** juveniles

## Tackling climate change

Exploring green technology and renewable energy adoption Measuring GHG emission and energy consumption





"Reaction Basis" EIA/SIA permit for CDMO manufacturing platform

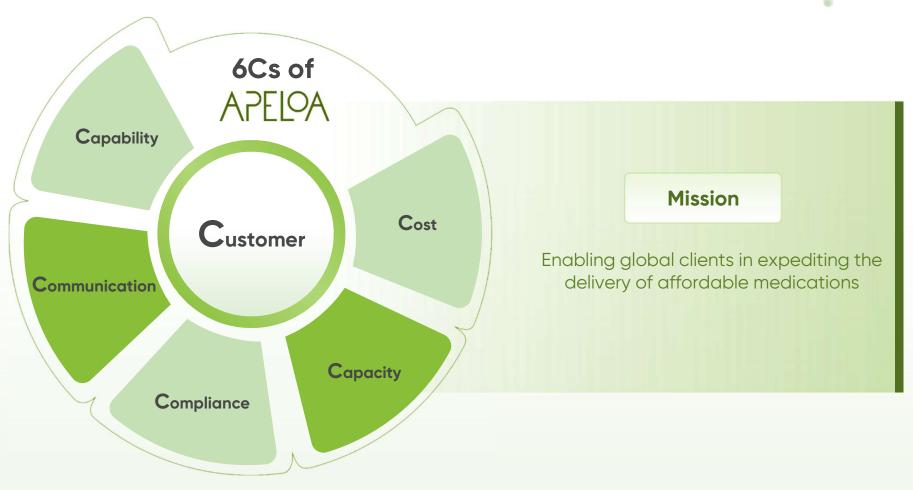
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# TECHNOLOGY FOR HEALTH





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