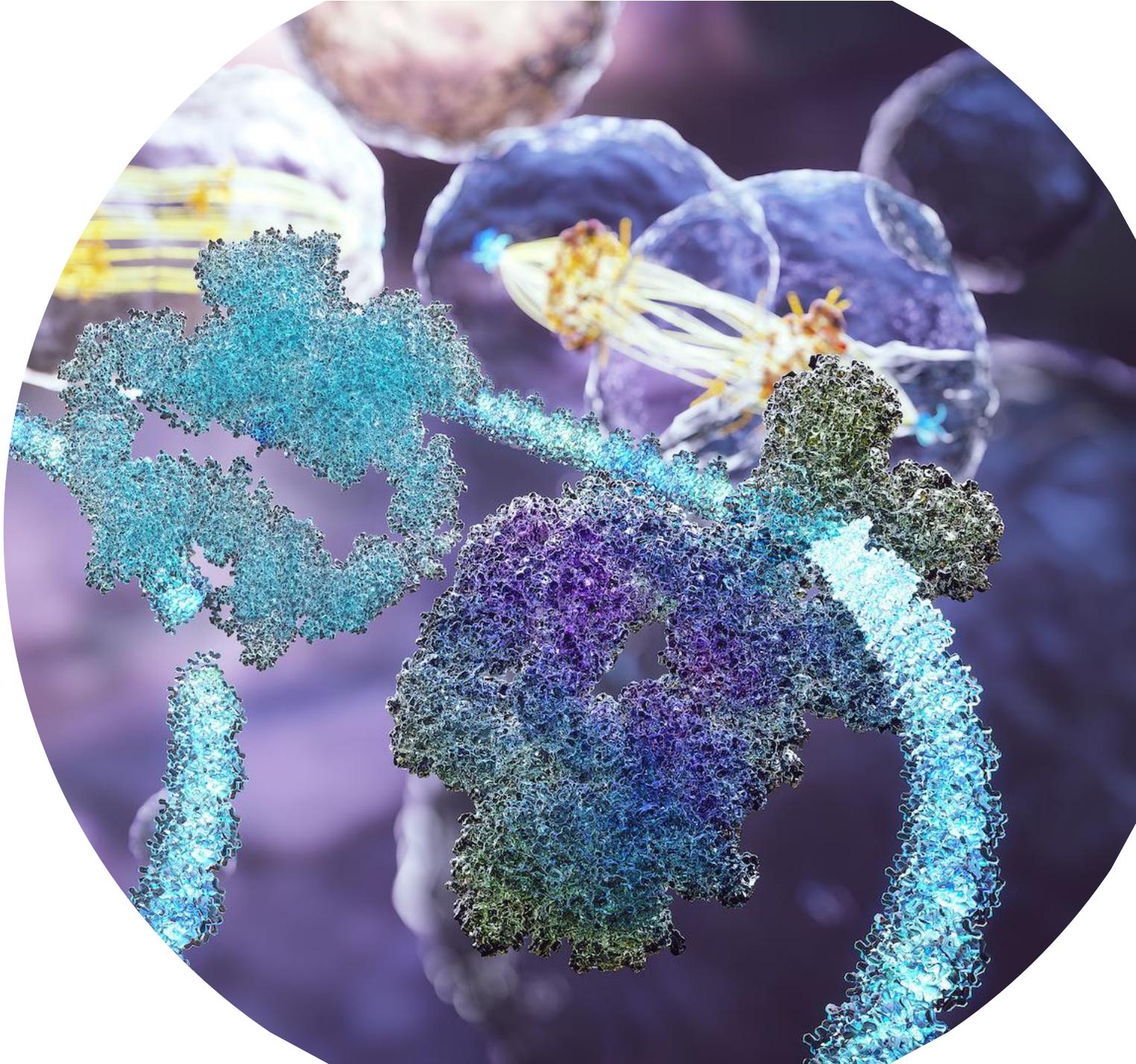




H1 and Q2 2023 Results

Conference call and webcast
for investors and analysts

28 July 2023



Forward-looking statements

In order, among other things, to utilise the 'safe harbour' provisions of the US Private Securities Litigation Reform Act of 1995, AstraZeneca (hereafter 'the Group') provides the following cautionary statement: This document contains certain forward-looking statements with respect to the operations, performance and financial condition of the Group, including, among other things, statements about expected revenues, margins, earnings per share or other financial or other measures. Although the Group believes its expectations are based on reasonable assumptions, any forward-looking statements, by their very nature, involve risks and uncertainties and may be influenced by factors that could cause actual outcomes and results to be materially different from those predicted. The forward-looking statements reflect knowledge and information available at the date of preparation of this document and the Group undertakes no obligation to update these forward-looking statements. The Group identifies the forward-looking statements by using the words 'anticipates', 'believes', 'expects', 'intends' and similar expressions in such statements. Important factors that could cause actual results to differ materially from those contained in forward-looking statements, certain of which are beyond the Group's control, include, among other things: the risk of failure or delay in delivery of pipeline or launch of new medicines; the risk of failure to meet regulatory or ethical requirements for medicine development or approval; the risk of failures or delays in the quality or execution of the Group's commercial strategies; the risk of pricing, affordability, access and competitive pressures; the risk of failure to maintain supply of compliant, quality medicines; the risk of illegal trade in the Group's medicines; the impact of reliance on third-party goods and services; the risk of failure in information technology or cybersecurity; the risk of failure of critical processes; the risk of failure to collect and manage data in line with legal and regulatory requirements and strategic objectives; the risk of failure to attract, develop, engage and retain a diverse, talented and capable workforce; the risk of failure to meet regulatory or ethical expectations on environmental impact, including climate change; the risk of the safety and efficacy of marketed medicines being questioned; the risk of adverse outcome of litigation and/or governmental investigations; intellectual property-related risks to our products; the risk of failure to achieve strategic plans or meet targets or expectations; the risk of failure in financial control or the occurrence of fraud; the risk of unexpected deterioration in the Group's financial position; the impact that global and/or geopolitical events such as the COVID-19 pandemic and the Russia-Ukraine war may have or continue to have on these risks, on the Group's ability to continue to mitigate these risks, and on the Group's operations, financial results or financial condition. Nothing in this document, or any related presentation/webcast, should be construed as a profit forecast.



H1 and Q2 2023 results

Conference call agenda

CEO Opening Remarks

Pascal Soriot
Chief Executive Officer

Financial Results

Aradhana Sarin
Chief Financial Officer

Oncology

Dave Fredrickson
EVP, Oncology Business

Susan Galbraith
EVP, Oncology R&D

BioPharmaceuticals

Ruud Dobber
EVP, BioPharmaceuticals Business

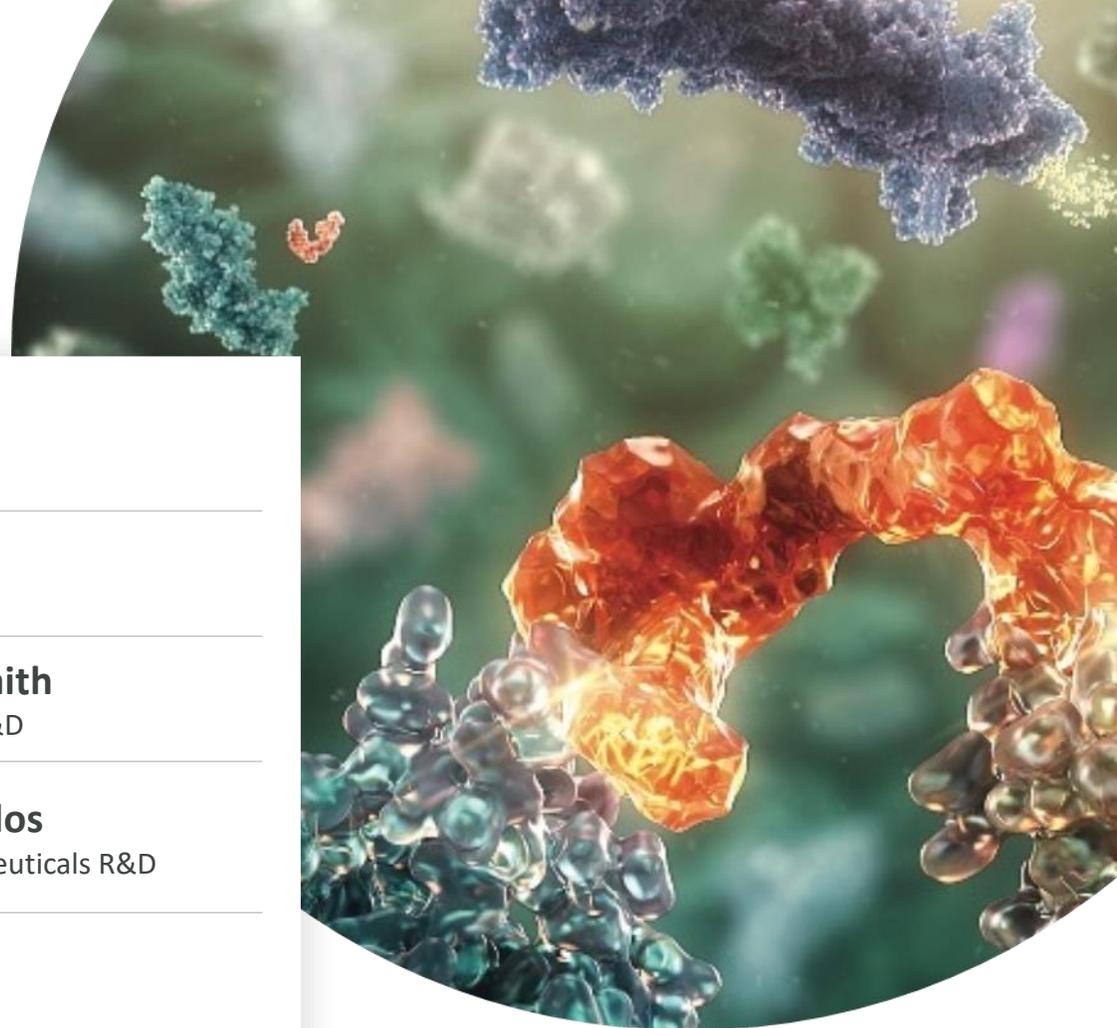
Mene Pangalos
EVP, BioPharmaceuticals R&D

Rare Disease

Marc Dunoyer
Chief Executive Officer, Alexion

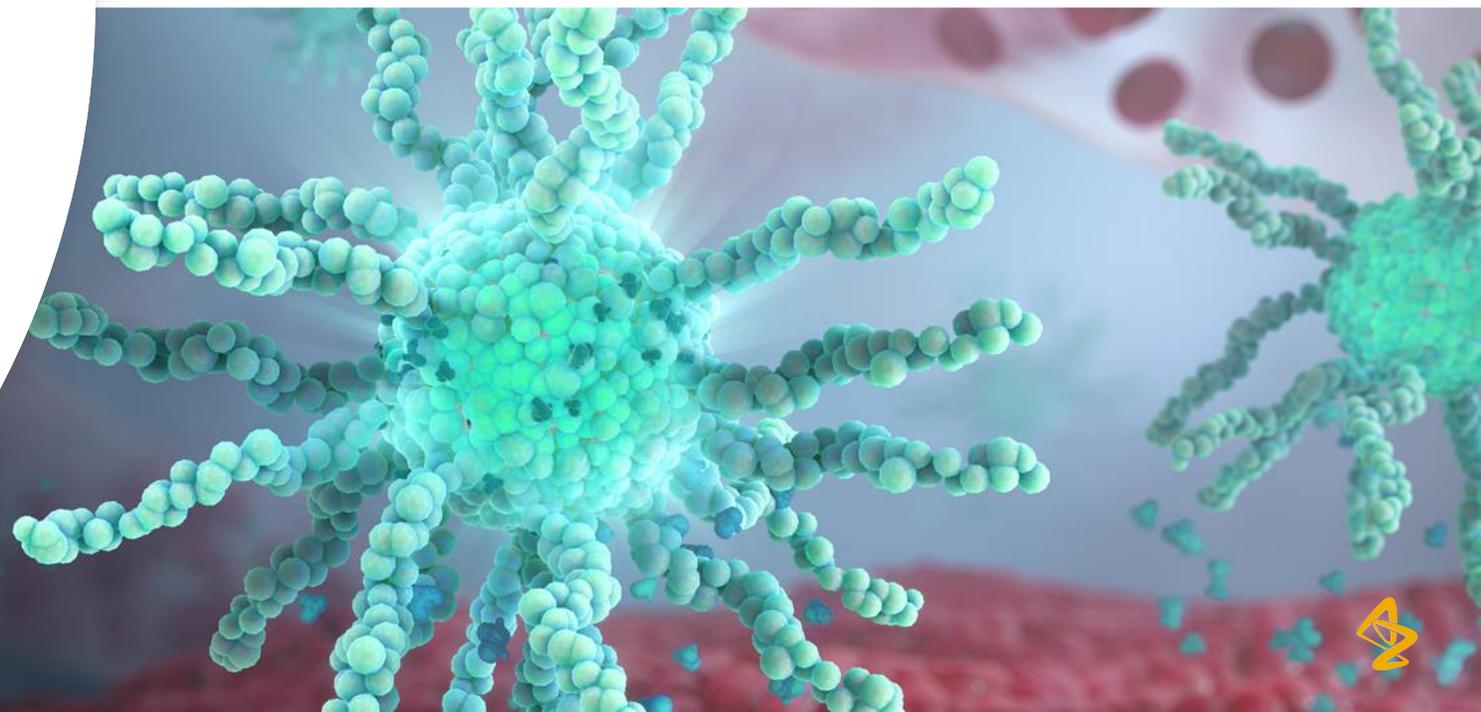
CEO Closing Remarks, Q&A

Pascal Soriot
Chief Executive Officer



CEO Opening Remarks

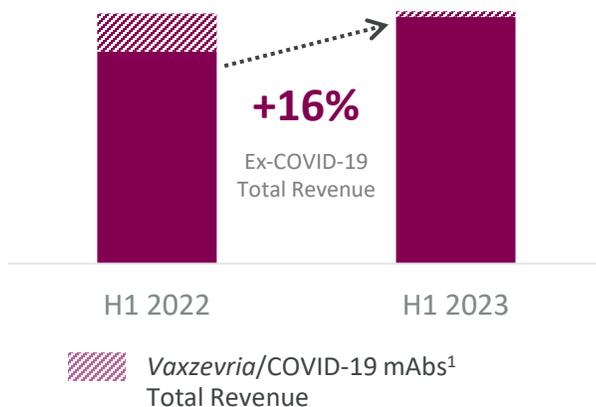
Pascal Soriot
CHIEF EXECUTIVE OFFICER



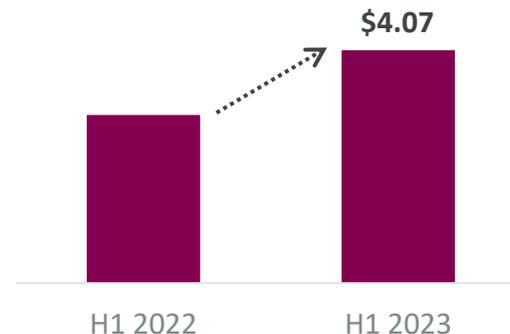
Benefitting from broad-based, diverse source of revenue

Strong commercial performance and financial execution in H1 2023

Total Revenue | +4%

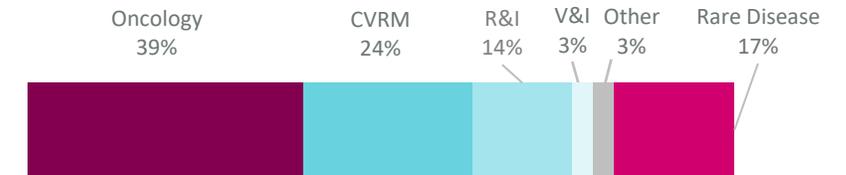


Core EPS | +21%

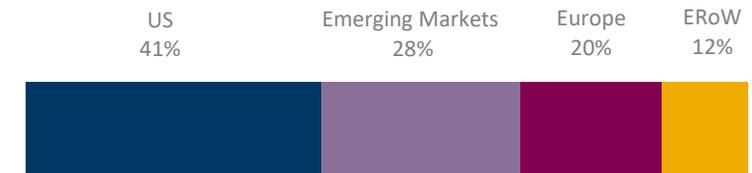


Broad-based, diverse source of revenue

H1 2023 | % Total Revenue by therapy area



H1 2023 | % Total Revenue by geography



Reiterating 2023 guidance: Core EPS to increase by a high single-digit to low double-digit %

All growth rates at CER. Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.

1. COVID-19 mAbs = *Evusheld* and AZD3152, the antibody currently in development.

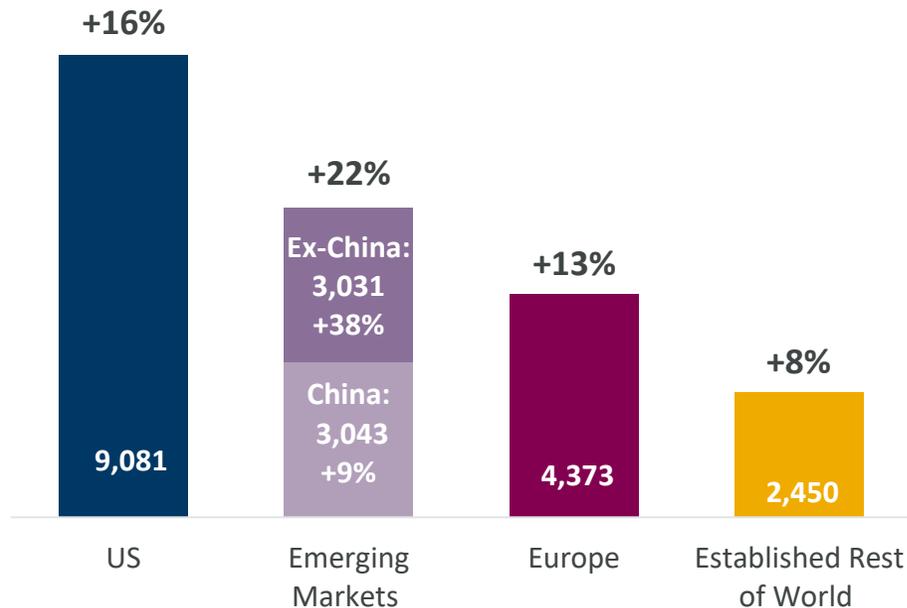
CER = constant exchange rates; EPS = earnings per share; CVRM = Cardiovascular, Renal & Metabolism; R&I = Respiratory & Immunology; V&I = Vaccines & Immune Therapies; ERoW = Established Rest of World.



Strong ex-COVID-19 growth across regions, disease areas

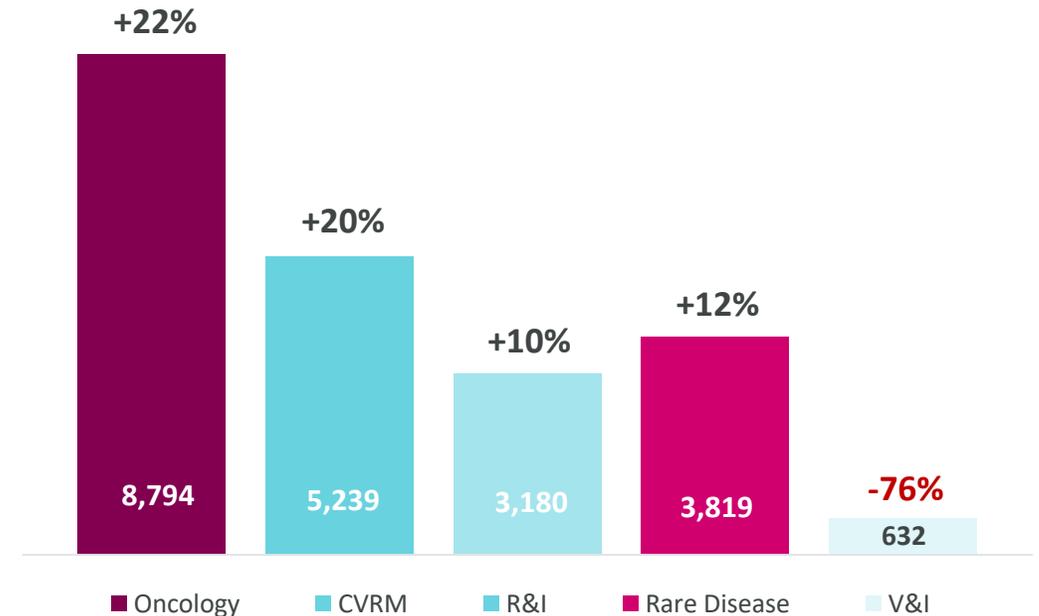
Double-digit growth in US, Emerging Market, EU

H1 Total Revenues (USD millions) and Growth vs PY



Double-digit growth across Oncology, CVRM, R&I and Rare disease

H1 Total Revenues (USD millions) and Growth vs PY



Demand for multiple medicines driving H1 growth

H1 2023 Performance

	Total Revenue USD million	Growth vs. PY USD million	Growth vs. PY CER%
<i>Farxiga</i>	2,834	729	40%
<i>Imfinzi</i>	1,976	682	57%
<i>Ultomiris</i>	1,364	511	64%
<i>Enhertu</i>	580	376	>2x
<i>Calquence</i>	1,185	282	33%
<i>Tagrisso</i>	2,915	211	12%
<i>Breztri</i>	307	128	76%
<i>Tezspire</i>	135	119	>8x
<i>Strensiq</i>	562	112	26%
<i>Fasenra</i>	744	82	14%
<i>Saphnelo</i>	115	79	>3x
<i>Lokelma</i>	198	69	59%
<i>Koselugo</i>	159	58	57%

LCM strategy sustaining growth momentum, new launches accelerating globally



Leading late-stage pipeline, proven R&D execution

Robust late-stage pipeline comprised of both LCM and NME opportunities

>120

ongoing late-stage clinical trials across our pipeline

14

unique NMEs in late-stage development

10 potential blockbuster opportunities
from 30 potential Phase III trials planned in 2023¹

Continued pipeline momentum with 8 positive pivotal trials in H1 2023 and catalyst-rich H2 2023



Tagrisso ADAURA OS (Adj EGFRm NSCLC)



Imfinzi AEGEAN (Neoadj-adj NSCLC)



Enhertu DESTINY-Pantumor02 (HER2 expressing tumours)



Lynparza + *Imfinzi* + anti-VEGF + CTx DUO-O (ovarian)



Tagrisso + CTx FLAURA2 (EGFRm NSCLC)



Lynparza + *Imfinzi* + CTx DUO-E (endometrial)



Imfinzi + FLOT MATTERHORN (Neoadj-adj gastric and GEJ)



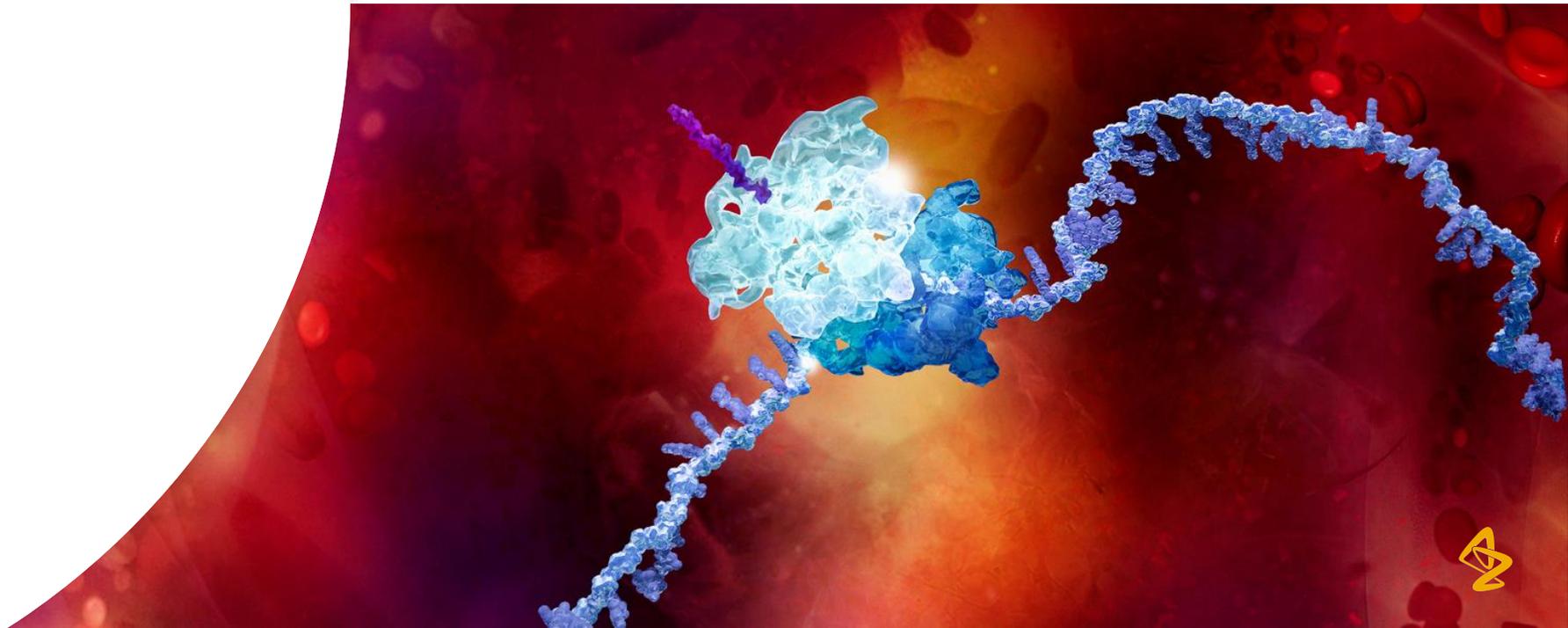
Dato-DXd TROPION-Lung01 (NSCLC)

1. Phase III trial start defined as first patient clinically dosed. R&D = Research & Development; siRNA = small interfering ribonucleic acid; NME = new molecular entity; OS = overall survival; EGFRm = epidermal growth factor receptor mutant; NSCLC = non-small cell lung cancer; Neoadj-adj = neoadjuvant-adjuvant; HER2 = human epidermal growth factor receptor 2; anti-VEGF = anti-vascular endothelial growth factor; CTx = chemotherapy; FLOT = fluorouracil, leucovorin, oxaliplatin and docetaxel; GEJ = gastroesophageal junction.

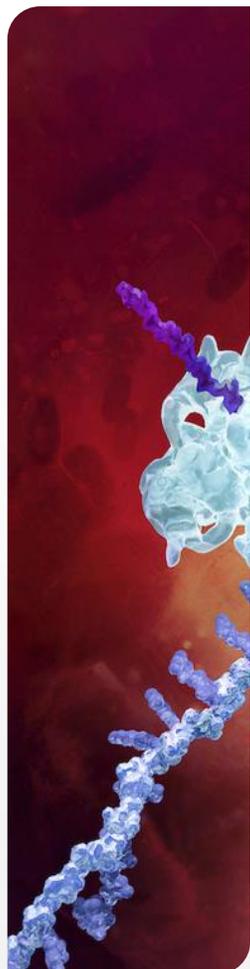


Financial Results

Aradhana Sarin
CHIEF FINANCIAL OFFICER



H1 2023 – Reported profit and loss



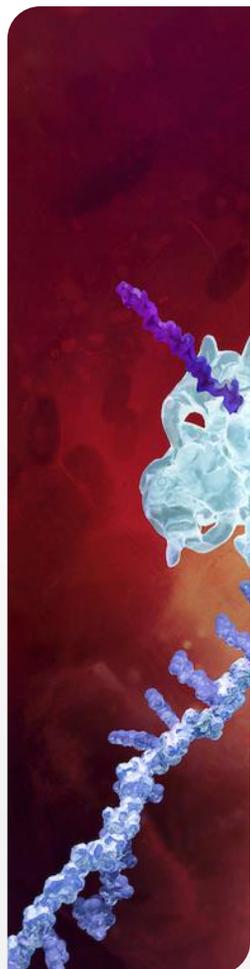
	H1 2023 \$m	CER change %	% Total Revenue	Q2 2023 \$m	CER change %	% Total Revenue
Total Revenue	22,295	4	100	11,416	9	100
- Product Sales	21,448	3	96	10,882	5	95
- Alliance Revenue	627	>2x	3	341	>2x	3
- Collaboration Revenue	220	(15)	1	193	n/m	2
Product Sales Gross margin	82.0%	+13 pp		82.0%	+12 pp	
Total operating expense¹	(14,588)	4	65	(7,784)	8	68
- R&D expense	(5,278)	16	24	(2,667)	7	23
- SG&A expense	(9,045)	(2)	41	(4,986)	8	44
Other operating income and expense	1,163	>5x	5	784	>6x	7
Operating profit	5,005	>4x	22	2,456	>6x	22
Tax rate	17%			13%		
Reported EPS	\$2.34	>6x		\$1.17	>9x	

Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.

Absolute values at actual exchange rates; changes at CER. Product Sales Gross margin excludes the impact of Alliance and Collaboration Revenue and any associated costs, thereby reflecting the underlying performance of Product Sales. 1. Total operating expenses include distribution, R&D and SG&A expenses. R&D = Research & Development; SG&A = Sales, General & Administrative; EPS = earnings per share; pp = percentage points; CER = constant exchange rates.



H1 2023 – Core profit and loss



	H1 2023 \$m	CER change %	% Total Revenue	Q2 2023 \$m	CER change %	% Total Revenue
Total Revenue	22,295	4	100	11,416	9	100
- Product Sales	21,448	3	96	10,882	5	95
- Alliance Revenue	627	>2x	3	341	>2x	3
- Collaboration Revenue	220	(15)	1	193	n/m	2
Product Sales Gross margin	82.9%	+3 pp		82.4%	+2 pp	
Total operating expense ¹	(11,483)	8	52	(5,995)	8	53
- R&D expense	(4,868)	9	22	(2,568)	8	22
- SG&A expense	(6,350)	8	28	(3,296)	8	29
Other operating income and expense	1,102	>5x	5	784	>6x	7
Operating profit	8,237	20	37	4,291	39	38
Tax rate	18%			17%		
Core EPS	\$4.07	21		\$2.15	38	

Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.

11 Absolute values at actual exchange rates; changes at CER. Product Sales Gross margin excludes the impact of Alliance and Collaboration Revenue and any associated costs, thereby reflecting the underlying performance of Product Sales.

1. Total operating expenses include distribution, R&D and SG&A expenses. R&D = Research & Development; SG&A = Sales, General & Administrative; EPS = earnings per share; pp = percentage points; CER = constant exchange rates.



Cash Flow, Net Debt and 2023 Financial Guidance

Continued EBITDA improvement

Net Debt bridge



Net Debt/EBITDA: 1.9x

Net Debt/EBITDA adjusted for Alexion inventory fair value uplift: 1.7x

Reiterating 2023 Guidance (CER)

Total Revenue

- *Excluding COVID-19 medicines*: low double-digit % growth
- *Including COVID-19 medicines*: low-to-mid single-digit % growth

Core EPS

- High single-digit to low double-digit %

Expected FX-impact: Total Revenue: A low single-digit adverse impact
Core EPS: Low to mid single-digit adverse impact³

Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.

1. Comprises disposal of intangible assets, movement in profit participation liability, purchase of intangible assets, payment of contingent consideration on business combinations, purchase and disposal of non-current asset investments, payment of Acerta Pharma share purchase liability and acquisitions of subsidiaries, net of cash acquired. 2. EBITDA adding back the impact of \$1,221m 12-month rolling period (FY 2022: \$3,484m) unwind of inventory fair value uplift recognised on acquisition of Alexion. AstraZeneca credit ratings: Moody's: short-term rating P-2, long-term rating A3, outlook stable. S&P Global Ratings: short-term rating A-1, long-term rating A, outlook stable. 3. Assuming average June 2023 foreign exchange rates for July to December 2023. EBITDA = earnings before interest, tax, depreciation and amortisation; CFO = net cash inflow from operating activities; EPS = earnings per share.



AI in Global Operations

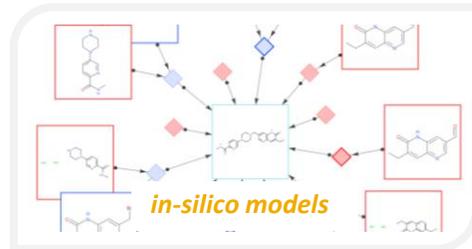
Leveraging the power of AI throughout our end-to-end supply chain

GLOBAL OPERATIONS | *drug development, manufacturing and supply chain powered by AI*



Drug development

digital synthetic route design



50% reduction in route design lead time with AI-enabled predictive tools

Solving for optimised routes with:

- > Fewest synthetic steps
- > Lowest potential COGS
- > Lowest carbon footprint

Manufacturing

advanced analytics and optimisation



>20% increased yield with AI-enabled analytic tools



Identify new parameters



Target process adjustments



Define new process ranges

Supply chain

digital twins for raw material planning



90% reduction in dispensing planning time with AI-enabled digital twins

1
2

- 1 | balance material, asset, resource availability
- 2 | balance for various operations and products

while ensuring "just-in-time" process

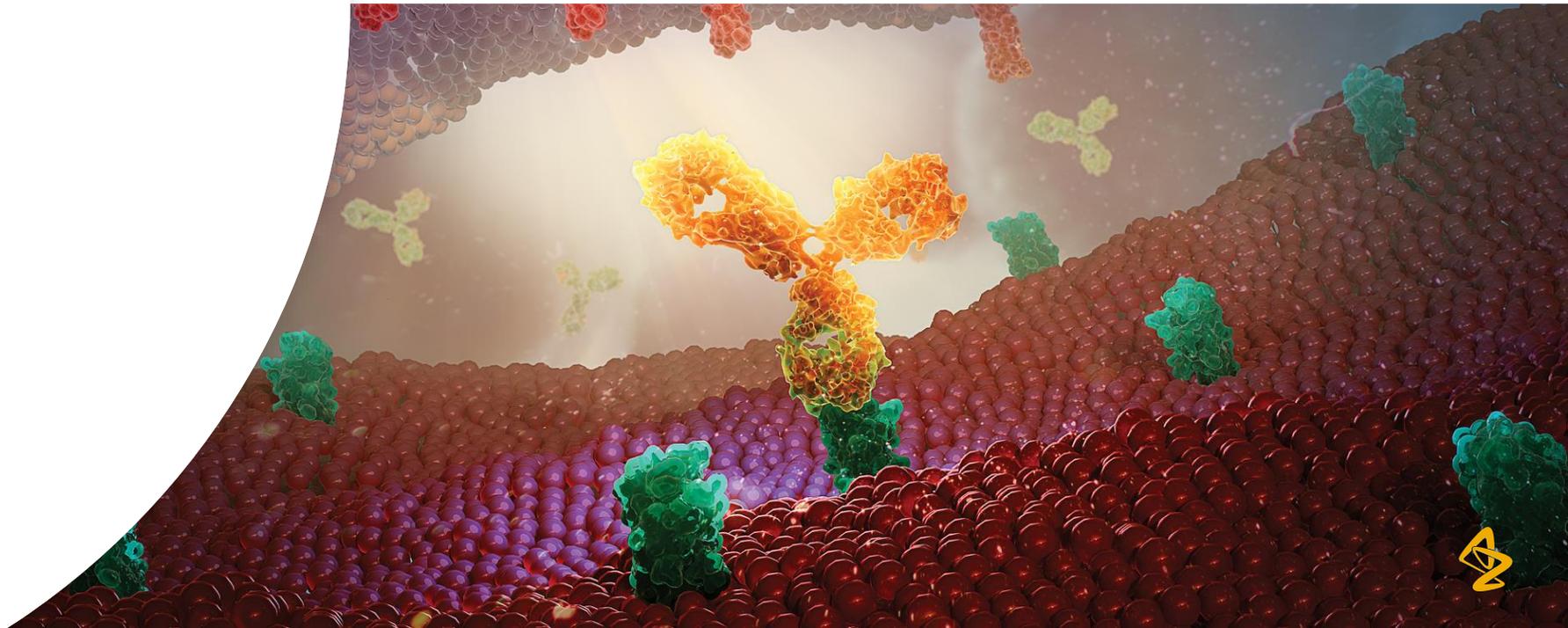


Oncology



Dave Fredrickson
ONCOLOGY BUSINESS

Susan Galbraith
ONCOLOGY R&D

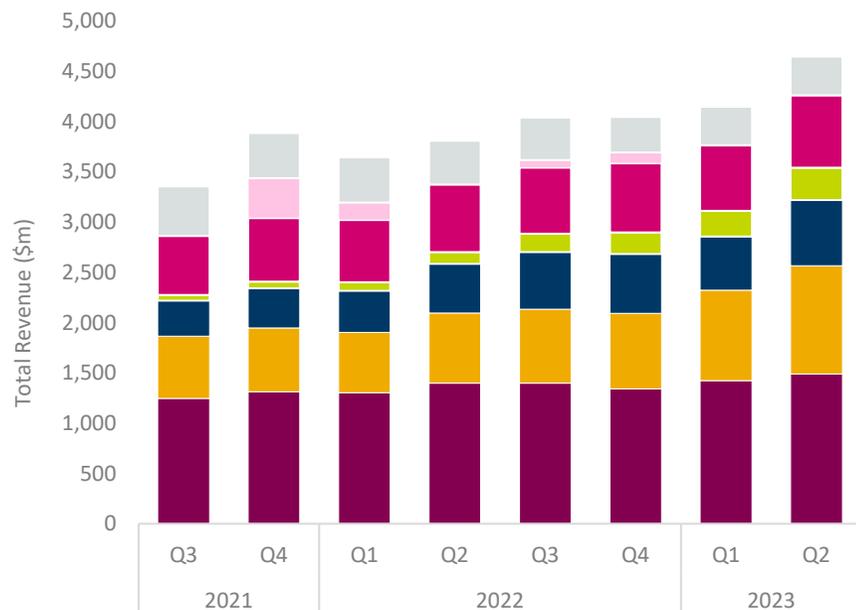


Oncology – H1 and Q2 2023

Total Revenue +22% in H1 2023 fuelled by demand growth and launch momentum

Oncology

H1 2023 \$8.8bn, +22%



Tagrisso Imfinzi/Imjudo Calquence Enhertu Lynparza (PS) Lynparza milestones Others

Q2 2023: key dynamics

- **Tagrisso** +10%, strong global demand growth underpinned by ADAURA, slightly offset by NRDL price impact
 - **Lynparza PS** +9%, double-digit growth in EU, ERoW and EM, offset by flattening demand in 2L ovarian cancer in US
 - **Imfinzi** +58%, driven by global launch acceleration (TOPAZ-1, HIMALAYA, POSEIDON)
 - **Calquence** +34%, sustained BTKi class leadership
 - **Enhertu** >2x, DB03/DB04 launch momentum, expanded reimbursement
-
- New indications: US (*Lynparza* PROpel BRCAm), CN (*Enhertu* DB04)
 - capivasertib CAPItello-291 granted Priority Review in US

All growth rates at CER.

Collaboration partners: Daiichi Sankyo (*Enhertu*), Merck & Co., Inc. (*Lynparza*).

CER = constant exchange rates; NRDL = National Reimbursement Drug List; PS = Product Sales; EU = Europe; ERoW = Established Rest of World; EM = Emerging Markets; BTKi = bruton tyrosine kinase inhibitor; BRCAm = breast cancer gene mutation; CN = China; DB03 = DESTINY-Breast03; DB04 = DESTINY-Breast04.



Oncology – key *Imfinzi* opportunities in focus

Strong launch uptake, robust LCM pipeline supporting continued IO leadership

Imfinzi new launches continue to drive near-term growth

HIMALAYA unresectable HCC

- First dual IO regimen
- Unprecedented 4-year OS data presented at ESMO World GI 2023

TOPAZ-1 1L BTC

- First, innovative IO regimen in BTC
- Established SoC in US within months
- EU, JP launch trajectory outpacing US

POSEIDON 1L NSCLC

- Encouraging launch uptake in crowded competitive landscape

New launch momentum (POSEIDON, HIMALAYA, TOPAZ) builds on top of sustained leadership in PACIFIC, CASPIAN

LCM HLRs in 2023 contribute to sustained, mid-term *Imfinzi* growth

-  **AEGEAN** (early resec. NSCLC)
 - Significant improvement in EFS vs neoadjuvant CTx
-  **DUO-O** (1L ovarian)
 - Further improvement with *Lynparza* + *Imfinzi*
-  **DUO-E** (1L endometrial)
 - Greater benefit in *Imfinzi* + *Lynparza* + CTx arm
-  **MATTERHORN** (Gastric and GEJ)
 - First IO + FLOT to demonstrate clinical benefit
-  **PACIFIC-2** (Stg. III unresec. NSCLC)
 - Potential to move IO upfront + cCRT

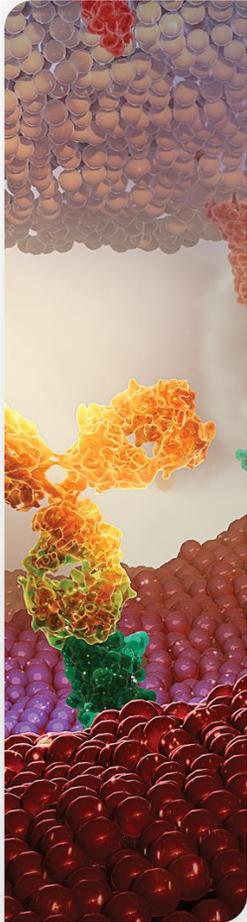
H2 '23
-  **EMERALD-1** (locoregional HCC)
 - Potential to improve PFS vs TACE therapy

H2 '23



Oncology – R&D highlights

Eight positive pivotal trial readouts so far this year



ASCO highlights 2023



ADAURA final overall survival

unprecedented survival in EGFRm NSCLC with 88% of patients alive on *Tagrisso* at 5 years



DUO-O interim analysis

37% reduction in risk of progression or death with *Lynparza* and *Imfinzi* added to chemotherapy and bevacizumab



DESTINY-PanTumor02 interim analysis

37% ORR and 11.8 month mDoR with *Enhertu* across range of HER2-expressing solid tumours

FLAURA2 | 1L EGFRm advanced NSCLC

Presidential plenary



R
1:1

Tagrisso

Tagrisso + carbo/cis + pemetrexed

Tagrisso

Tagrisso + pemetrexed

DUO-E | advanced endometrial cancer

R
1:1:1

Platinum CTx + *Imfinzi* placebo

Platinum CTx + *Imfinzi*

Platinum CTx + *Imfinzi*

Imfinzi placebo + *Lynparza* placebo

Imfinzi + *Lynparza* placebo

Imfinzi + *Lynparza*

MATTERHORN | resectable gastric and GEJ cancers

R
1:1

Placebo + FLOT

Imfinzi + FLOT

SURGERY

Placebo + FLOT

Imfinzi + FLOT

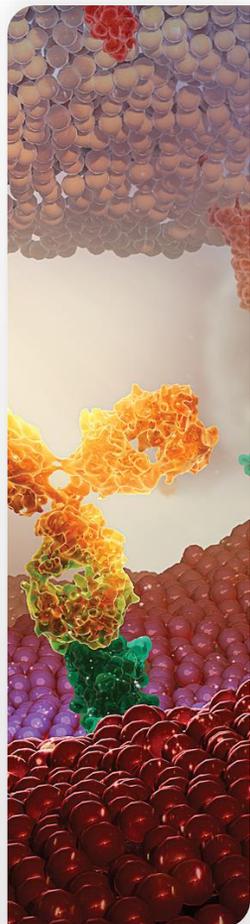
Placebo

Imfinzi



Oncology – R&D highlights

TROPION-Lung01 reinforces importance of Dato-DXd in NSCLC and beyond



TROPION-Lung01

Dato-DXd in 2-3L advanced NSCLC

Statistically significant PFS benefit

Favourable early OS trend

All grade ILD consistent with previous Dato-DXd trials

Some Grade 5 ILD cases observed

No new safety signals

Proceeding to file data with FDA

	Replace CTx as monotherapy	Further outcomes with novel combinations	Assess predictive value of TROP2 biomarker
Lung cancer	TROPION-Lung01 2-3L NSCLC w/ and w/o AGAs ✓	TROPION-Lung02 + pembro ± platinum CTx 1-3L NSCLC w/o AGAs ✓	
	TROPION-Lung05 2L+ NSCLC w/ AGA ✓	TROPION-Lung04 + <i>Imfinzi</i> / <i>rilvegostomig</i> / <i>volrustomig</i> ± carbo 1-2L NSCLC w/o AGAs ✓	
		TROPION-Lung07 + pembro ± platinum CTx 1L PD-L1<50% NSCLC w/o AGAs	
		TROPION-Lung08 + pembro 1L PD-L1≥50% NSCLC w/o AGAs	
		AVANZAR + <i>Imfinzi</i> + carbo 1L NSCLC w/o AGAs	
Breast cancer	TROPION-Breast01 2L+ HR+ HER2- mBC H2	BEGONIA + <i>Imfinzi</i> 1L aTNBC ✓	
	TROPION-Breast02 1L PD-L1- aTNBC ¹	TROPION-Breast03 ± <i>Imfinzi</i> adjuvant eTNBC	
Other tumours	TROPION-PanTumor01 advanced solid tumours ✓		
		TROPION-PanTumor03 +/- combinations advanced solid tumours	

Solid colour indicates Phase III trial; outline only indicates Phase I or II trial. Lung studies in the locally advanced / metastatic setting.

1. or not candidate for PD-(L)1 therapy for another reason. Dato-DXd = datopotamab deruxtecan; NSCLC = non-small cell lung cancer; PFS = progression free survival; OS = overall survival; ILD = interstitial lung disease; FDA = US Food and Drug Administration; AGA = actionable genomic alterations; pembro = pembrolizumab; CTx = chemotherapy; carbo = carboplatin; PD-L1 = programmed cell death ligand 1; EGFRm = epidermal growth factor receptor mutation; mBC = metastatic breast cancer; HR+ = hormone receptor positive; HER2- = human epidermal growth factor receptor negative; aTNBC = advanced triple negative breast cancer; eTNBC = early triple negative breast cancer.

Collaboration partners: Daiichi Sankyo (Dato-DXd)



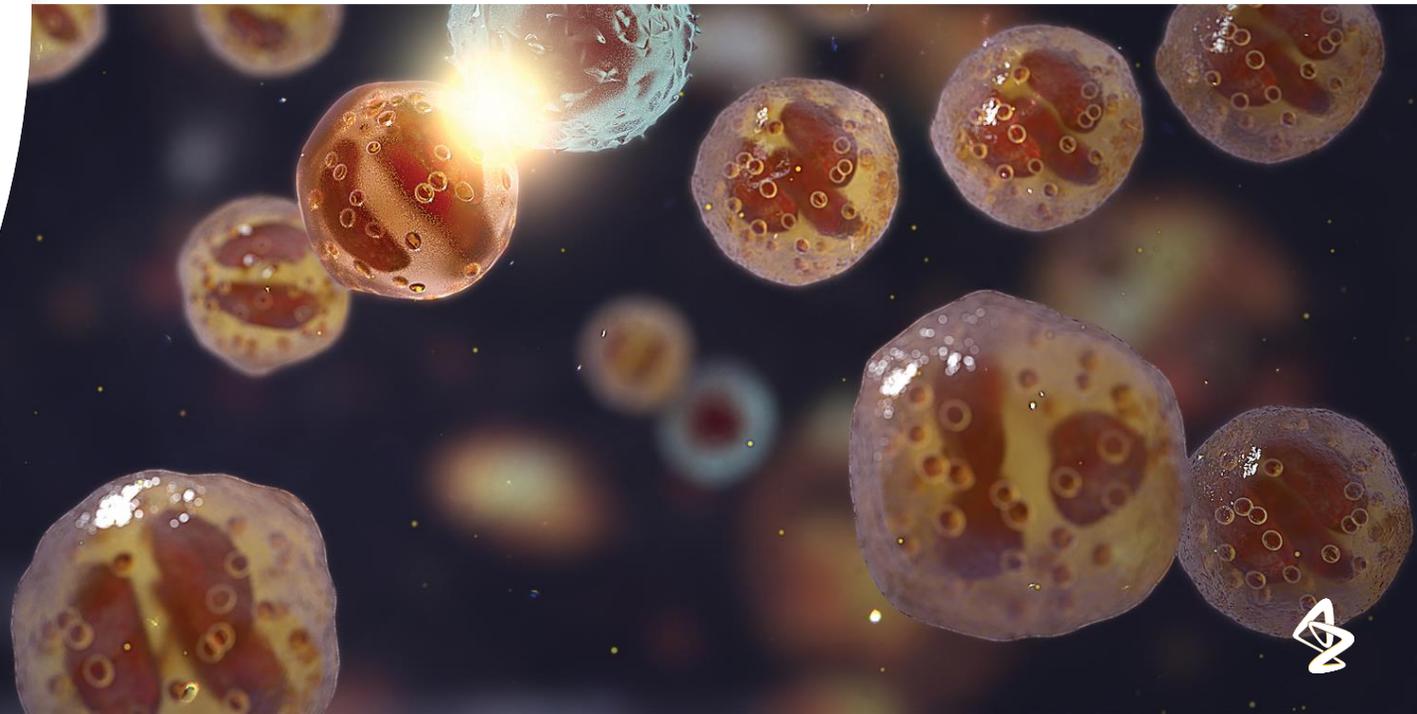
BioPharmaceuticals

Ruud Dobber

BIOPHARMACEUTICALS BUSINESS

Mene Pangalos

BIOPHARMACEUTICALS R&D

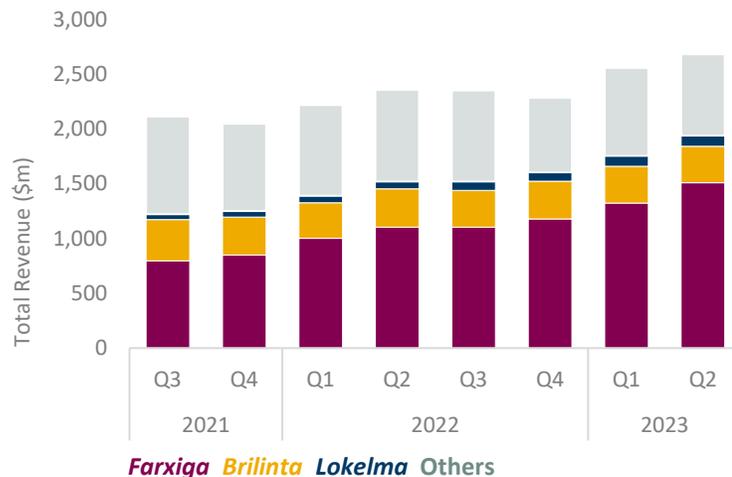


BioPharmaceuticals – H1 and Q2 2023

Double digit growth from CVRM and R&I

CVRM

H1 2023 \$5.2bn, +20%

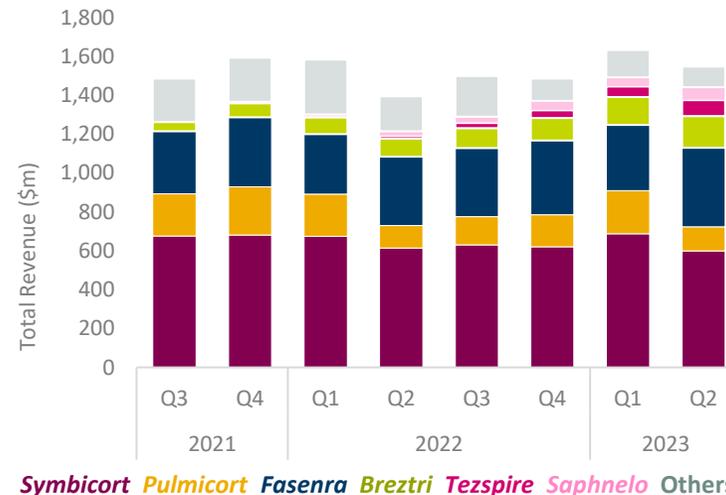


Q2 dynamics

- **Farxiga** +41%, to \$1.5bn
- **Lokelma, roxadustat and Andexxa**, strong double-digit growth

R&I

H1 2023 \$3.2bn, +10%

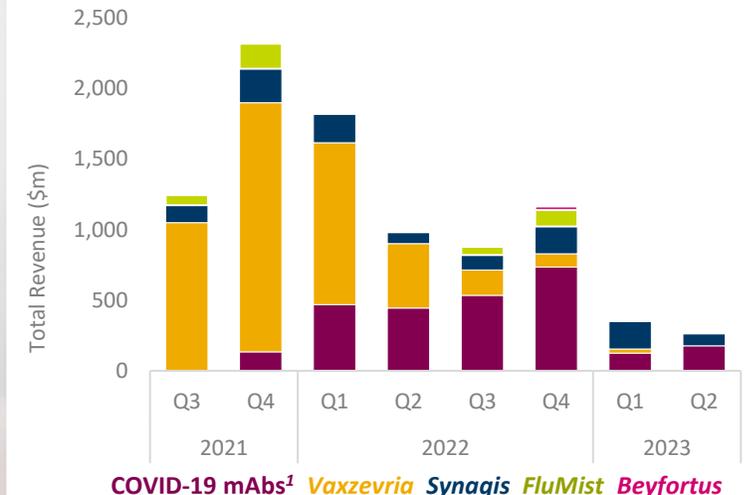


Q2 dynamics

- **Fasenra** +16%, to \$406m
- **Breztri** +79%, to \$163m
- **Tezspire** +50%, sequential QoQ growth

V&I

H1 2023 \$632m, -76%



Q2 dynamics

- **COVID-19 medicines** declined by \$0.7bn
- **FluMist** \$10m, milestone for Japan approval
- **Beyfortus**, first Product Sales

All growth rates at CER.

1. COVID-19 mAbs = Evusheld and AZD3152, the antibody currently in development

CER = constant exchange rates; CVRM = Cardiovascular, Renal and Metabolism; R&I = Respiratory and Immunology; QoQ = quarter on quarter; V&I = Vaccine and Immune Therapies; mAbs = monoclonal antibodies.

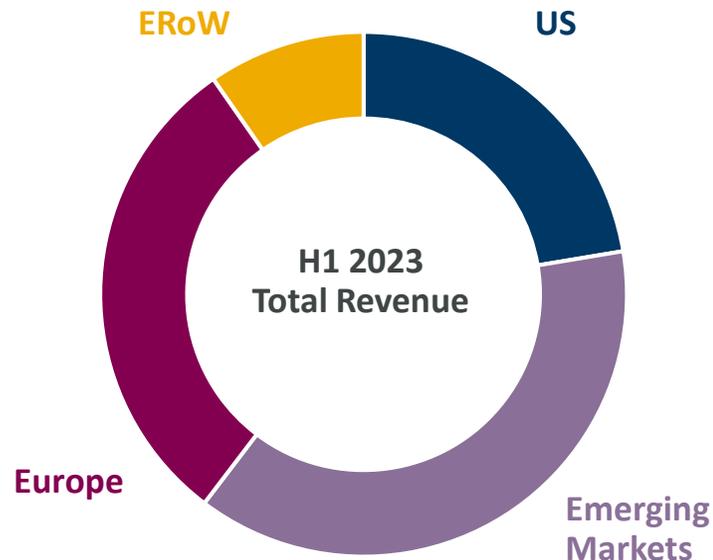
Collaboration partners: Amgen (Tezspire); Sanofi (Beyfortus).



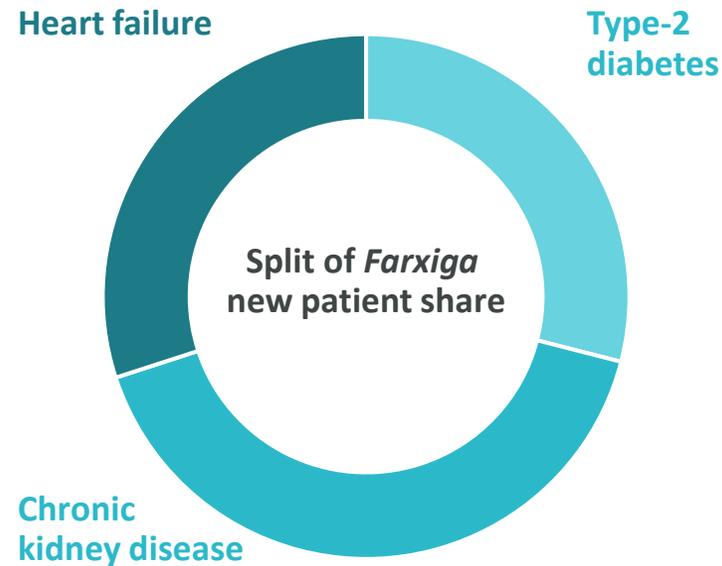
Farxiga – strategic development

Redefined outcomes for patients, clear evidence of AstraZeneca's core strength in CVRM

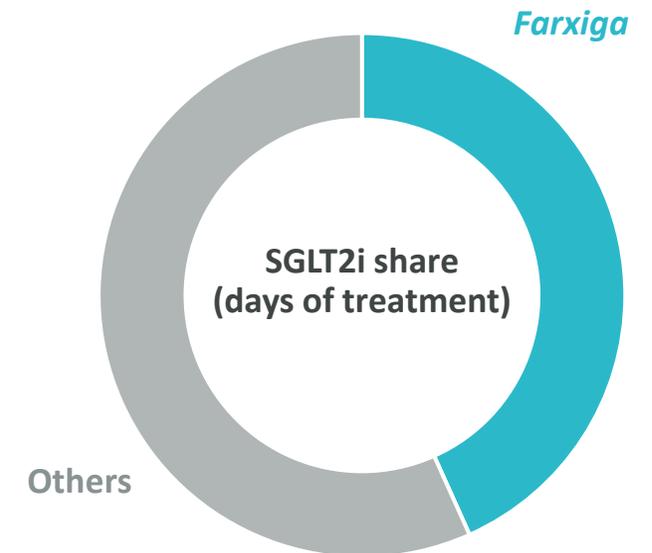
Diverse source of Total Revenues



Growth driven by HF and CKD¹



Farxiga SGLT2i class leadership²



Phased LOE with a late-stage pipeline to sustain therapy area leadership



BioPharmaceuticals – R&D highlights

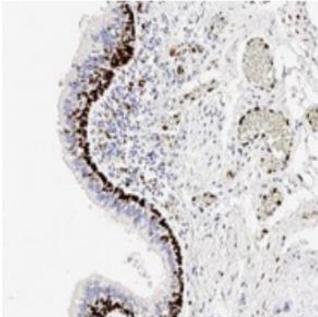
ATS and ERA data highlights improvement in patient outcomes across R&I and CVRM

tozorakimab

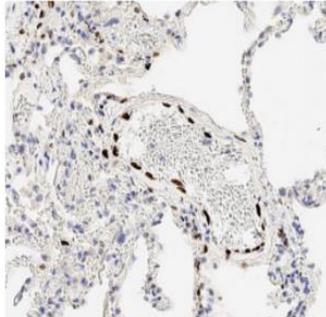


Severe viral lung infection

Airway epithelium



Airway endothelium



- Staining showed localised IL-33 in airway
- IL-33red/sST2 levels associated with poor clinical outcomes
- ST2 expressed in alveolar capillary cells

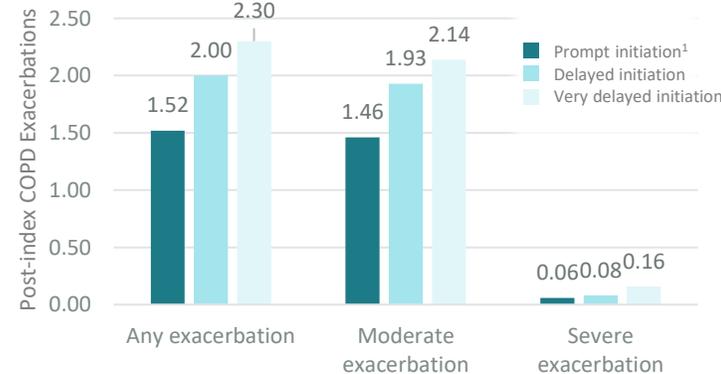
Clear rationale for targeting IL-33 in severe viral lung infection

Breztri



Real-world evidence

- **EROS:** relationship between timing of *Breztri* initiation and exacerbation rate
- Initiating <30 days resulted in 24% reduction in the risk of future exacerbations vs delaying treatment up to six months



Exacerbation risk increased by 5% every month *Breztri* treatment was delayed

Farxiga and Lokelma



Real-world evidence and guideline updates

- **REVEAL-CKD:** 85-97% Stage 3 CKD undiagnosed²
- Delaying diagnosis of CKD by one year:
 - ↑ risk of deterioration by 40%
 - ↑ risk of kidney transplant/long-term dialysis treatment by 63%
- **ZORA** highlighted role of *Lokelma* for patients with HK whilst on RAASi
- CKD and HF guidelines³ recommend and support use of K⁺ binders

Highlights importance of early diagnosis and concomitant K⁺ use in RAASi therapy

1. Prompt initiation = initiated therapy ≤30 days, Delayed initiation = initiated therapy >30 - ≤180 days, Very delayed initiation = initiated therapy 180 days. 2. REVEAL-CKD multinational study, data referenced includes Spain, Australia, Canada and Brazil.

3. Guidelines include: KDIGO (HTN in CKD and Diabetic Kidney), ESC (HF), AHA/ACC/HFSA (HF)



BioPharmaceuticals – novel platforms

Disruptive biology to access next-generation therapeutics



Oligonucleotides

eplontersen (TTR ASO),
ATTRv-PN/CM – Phase III

AZD2693 (PNPLA3 ASO),
NASH – Phase IIb

AZD7503 (HSD17B13 LICA),
NASH – Phase I

AZD2373 (APOL1 ASO),
CKD – Phase I



Advanced biologics

MEDI7352 (NGF/TNF),
pain – Phase II

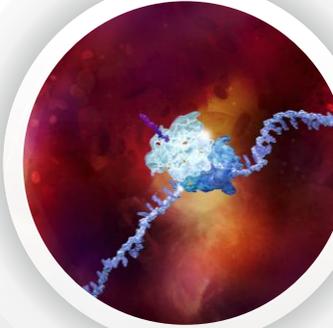
AZD8630 (iTSLP),
asthma – Phase I



Cell therapy

Procella – HF¹
THERAPEUTICS

Quell^{TX} – CAR-Treg
(T1D and IBD)

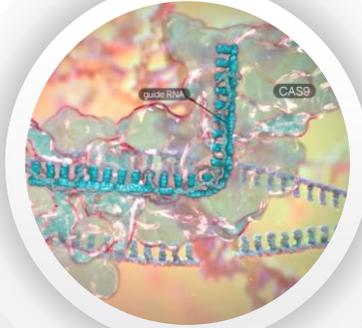


RNA therapies

mRNA

saRNA

cRNA



Gene therapy

Familial hyper-
cholesterolaemia (LDLR)

Duchenne musculo-
dystrophy (dystrophin)

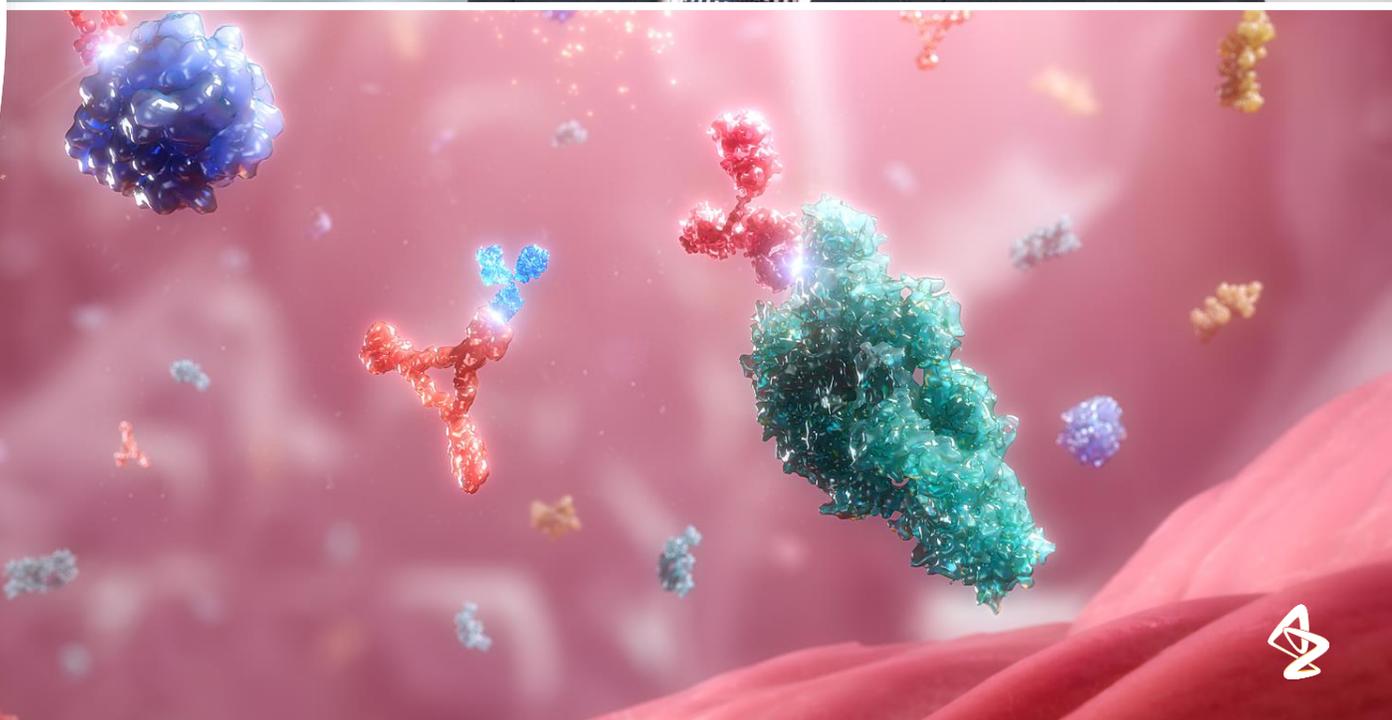
Huntington's disease (HTT)



Rare Disease

Marc Dunoyer

CHIEF EXECUTIVE OFFICER,
ALEXION

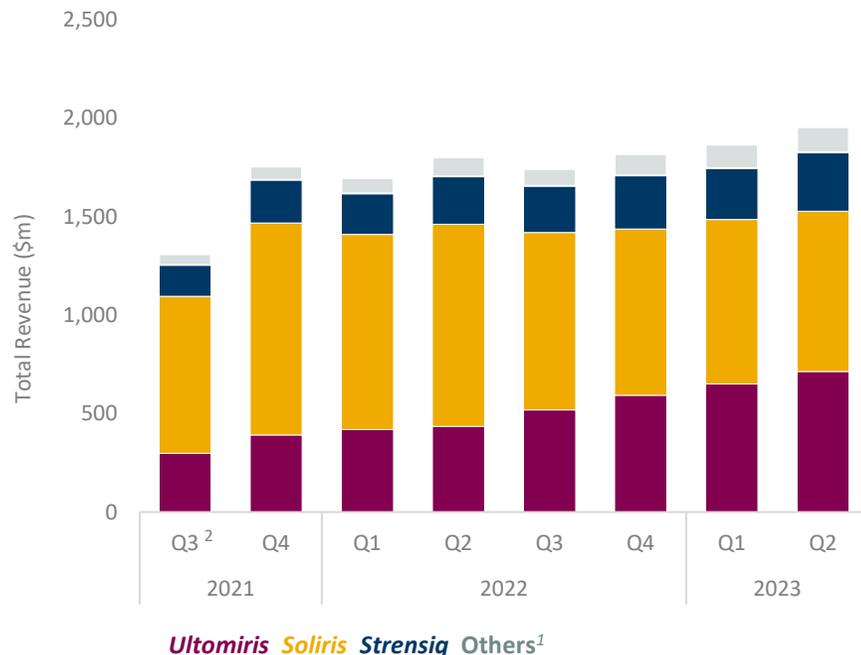


Rare Disease – H1 and Q2 2023

Continued expansion in neurology, growth beyond complement

Rare Disease

H1 2023 \$3.8bn, +12%



Q2 2023: key dynamics

Continued strength of C5 Franchise

- **Ultomiris** +66%, driven by growth in neurology, expansion into new markets and successful conversion from *Soliris*
- **Soliris** (19%), decline reflecting conversion partially offset by NMOSD growth

Strensiq, +25% and **Koselugo**, +30%

- Reflecting continued strength of patient demand and expansion into new markets

All growth rates at CER.

1. Includes *Kanuma* and *Koselugo*. 2. Q3 2021 Total Revenues reported only comprise of those booked by AstraZeneca following completion of the acquisition of Alexion on 21 July 2021.

25 C5 = C5 inhibitors *Ultomiris* and *Soliris*; CER = constant exchange rates; NMOSD = neuromyelitis optica spectrum disorder.

Collaboration partners: Merck & Co., Inc. (*Koselugo*).



Rare Disease – R&D highlights

Accelerating in ALXN2220 to Phase III for ATTR-CM

ALXN2220 (NI006) | mAb IgG1

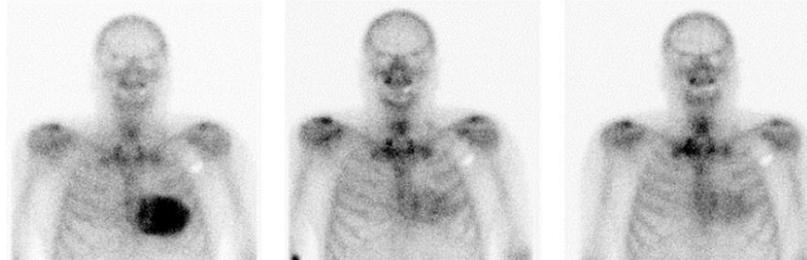
Phase Ib¹ in ATTR-CM



Representative images from scintigraphy after treatment with ALXN2220

A Patient 1, 75yo male, ATTRwt, 60 mg/kg

Baseline → NI006 → 4 months → NI006 → 12 months



H/WB ratio
Total NI006

6.9 %

2.5 %

20.7 g (46.1 d*mg/mL)

2.5 %

61.7 g (162.7 d*mg/mL)

✓ clearance of cardiac amyloid shown at 4 and 12 months

✓ improvement in cardiac function (NT-proBNP)

✓ potential monthly i.v. dosing

Selectively binds and removes misfolded amyloid fibrils

Complimentary mechanisms

transforming ATTR-CM

pathophysiology

medicine and modality

1

TTR production in the liver



Silencer

eplontersen blocks TTR synthesis

2

Tetramer formation

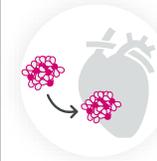


Stabiliser

acoramidis² stabilises TTR tetramers

3

Organ deposition



Depleter

ALXN2220 binds to misfolded TTR, removes toxic fibrils

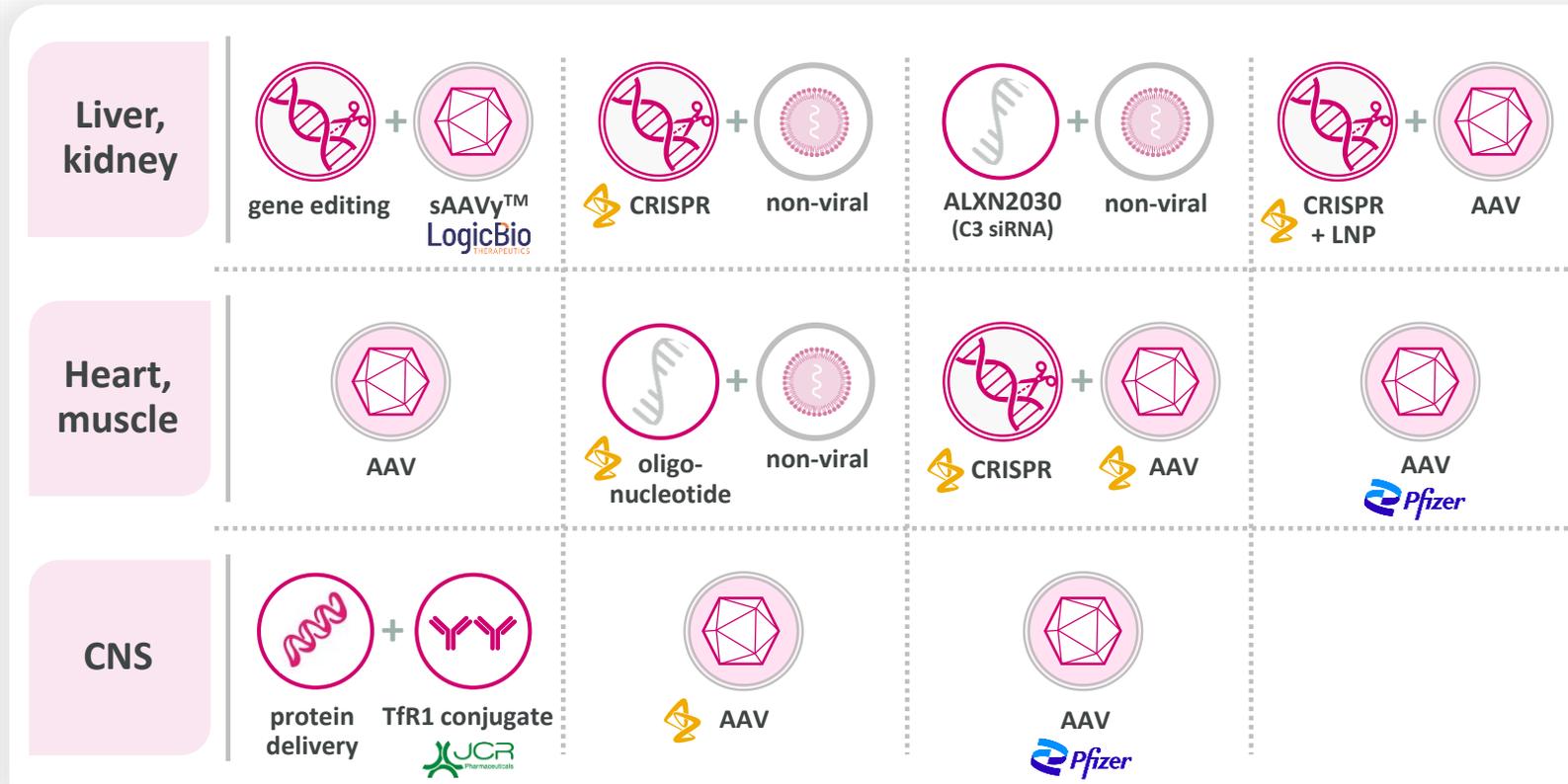
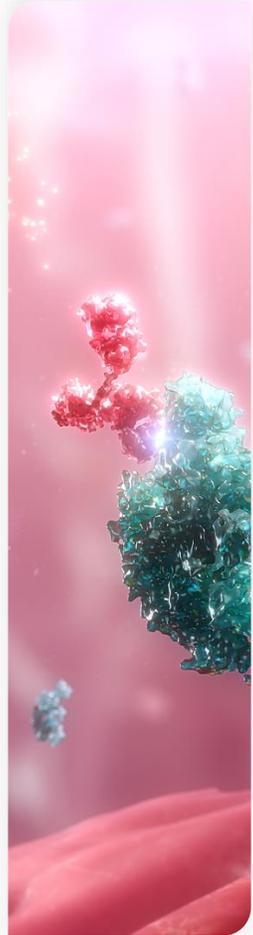
1. Garcia-Pavia et al. NEJM Phase 1 Trial of Antibody ALXN2220 (NI006) for Depletion of Cardiac Transthyretin Amyloid. Representative images from serial bisphosphonate scintigraphy from one patient randomly assigned to receive ALXN2220 (NI006). 2. Alexion, AstraZeneca Rare Disease has the Japanese rights to acoramidis. ATTR-CM = transthyretin amyloid cardiomyopathy; mAb = monoclonal antibody; IgG1 = immunoglobulin G1; yo = year-old; wt = wild-type; H/WB = heart-to-whole body ratio; TTR = transthyretin.

Collaboration partner: Neurimmune; Ionis; BridgeBio.



Rare Disease – R&D highlights

Innovative delivery platforms expanding genomic research



Furthering our genomic capabilities

- > accelerating genomic portfolio ambitions with acquisition of Pfizer AAV capsids
- > combining innovative technologies with LogicBio and AstraZeneca
- > partnering to access novel gene delivery technology

Acquisitions and collaborations complement existing AstraZeneca technologies and increases genomic medicine portfolio >4x

sAAVy™ = proprietary adeno-associated virus capsid engineering platform; CRISPR = clustered regularly interspaced short palindromic repeats; C3 = complement protein 3; siRNA = short interfering RNA; LNP = lipid nanoparticle;

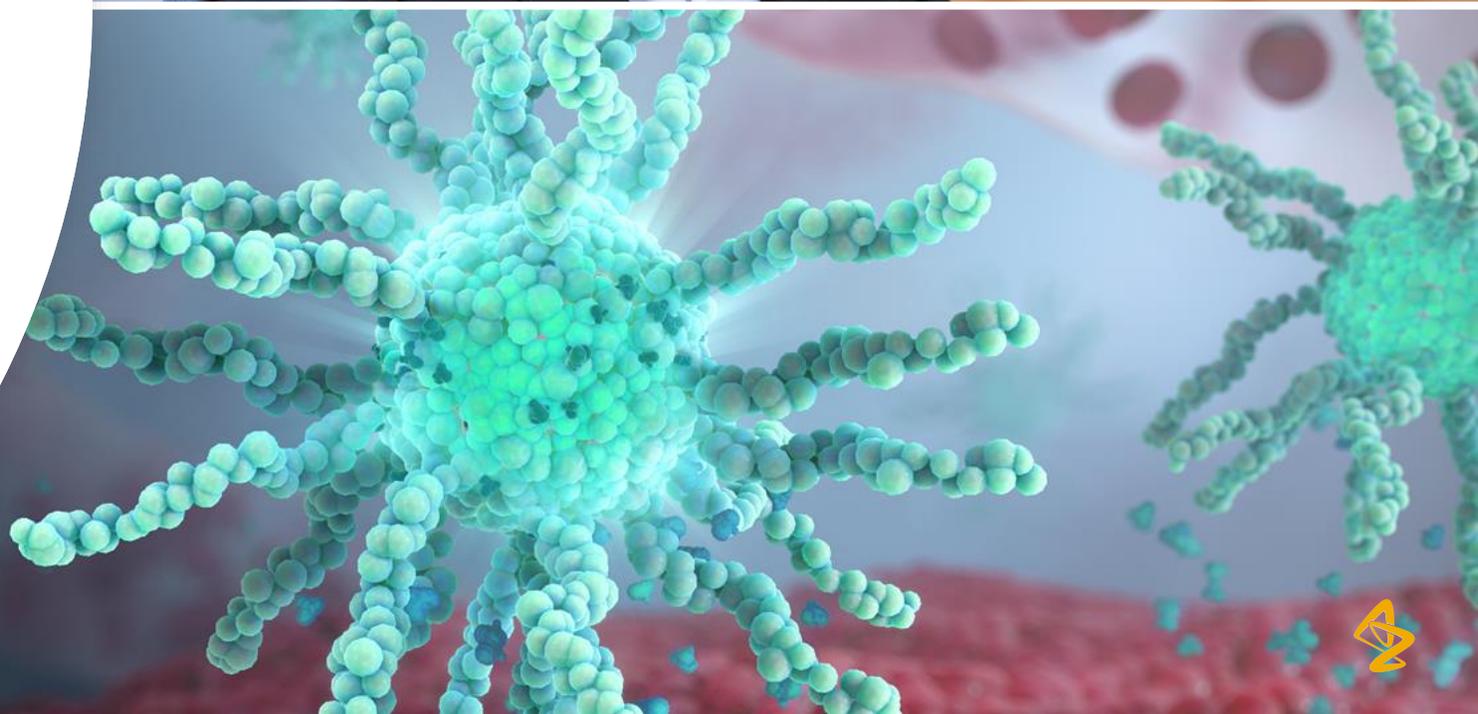
27 AAV = adeno-associated virus; CNS = central nervous system; TfR1 = transferrin receptor protein 1.

Note: Acquisition of Pfizer preclinical gene therapy programmes subject to customary regulatory clearances and other closing conditions.



CEO Closing Remarks

Pascal Soriot
CHIEF EXECUTIVE OFFICER



Progressing Ambition Zero Carbon

Taking bold action to mitigate climate change through new and expanded initiatives

AMBITION ZERO CARBON | on track to deliver absolute reductions in GHG emissions across our value chain

**Scope
1 & 2**

Decarbonisation

through pioneering partnership with Vanguard Renewables

Enabling the delivery of **renewable natural gas** to all US sites **by 2026**¹

to save:

650,000m

British thermal units
per year

35kt

GHG emissions
per year

AstraZeneca

Vanguard
RENEWABLES

Scope 3

AZ Forest

expanding our global reforestation and biodiversity initiative

Investing **\$400m** to plant and maintain **200 million trees** by 2030

to:

- restore biodiversity
- build on existing forest projects
- maximise environmental and community co-benefits
- sequester carbon

**Scopes
1, 2, 3**

Transition plan

focused on major decarbonisation initiatives and residual emissions removal

Making progress towards our **SBTi approved targets**²

	Scope 1 & 2	Scope 3
Major initiatives	<ul style="list-style-type: none"> — site efficiency — renewable power/heat — electric vehicle — management of F-gas 	<ul style="list-style-type: none"> — supplier engagement — NGP — sustainable product design
Residual emissions removal	<ul style="list-style-type: none"> — <2% using bioenergy with carbon capture and storage 	<ul style="list-style-type: none"> — nature-based solutions³, including AZ Forest



Robust Phase III catalysts in H2 2023

Key trial readouts reinforce transformative pipeline potential



capivasertib
CAPitello-290
(1L mTNBC)

*Extend beyond
PIK3/AKT/PTEN alterations
in unselected population*



Fasenra
MANDARA
(EGPA)

*Reinforcing first-choice
biologic in eosinophil
driven diseases*



Dato-DXd
TROPION-Breast01
(2L+ HR+/HER2- mBC)

*Building on TNBC efficacy,
expanding into HR+/HER2-
mBC (c.70% mBC subtypes)*



Imfinzi
PACIFIC-2
(Stg. III unresec. NSCLC)

*Potential to move IO
upfront + cCRT*



Imfinzi
EMERALD-1
(loco-regional HCC)

*Potential to improve PFS vs
TACE therapy*



AZD3152
SUPERNOVA
immuno-bridging sub-study
(COVID-19 prevention)

*Next-gen prophylactic LAAB
for immunocompromised
(c.2% population)*



Total Revenue ambition¹:
low double-digit % 2021-2025
Industry-leading growth 2025+



**Remain focused on Operating
Margin expansion**



**At least 15 NMEs
approved by 2030**



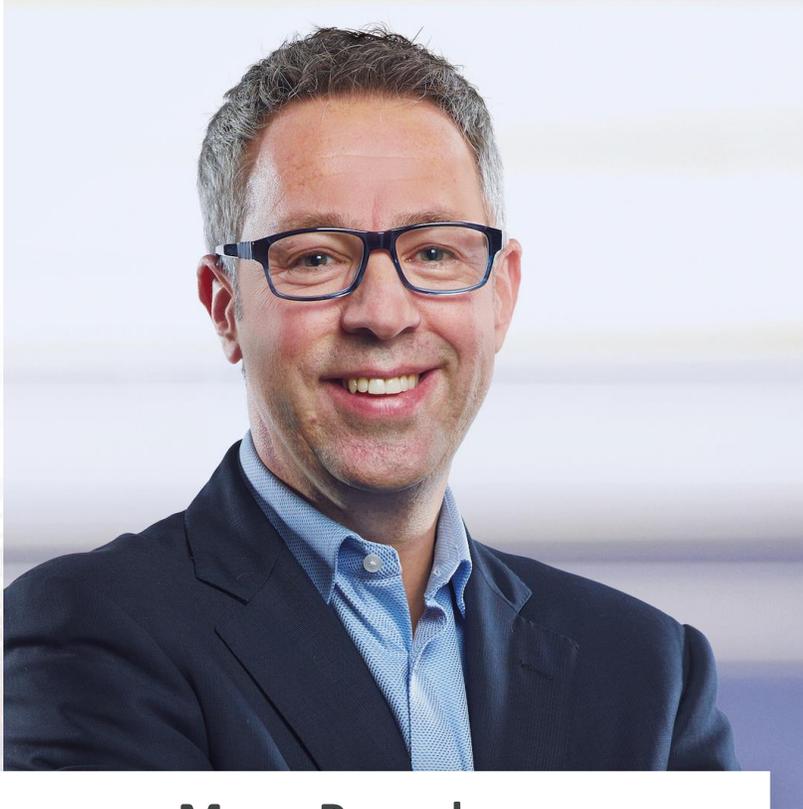
Emissions reduction:
98% by end 2025 – Scope 1 & 2
50% by 2030 – Scope 3

Confident in leading growth profile: base business strength with innovative late-stage pipeline

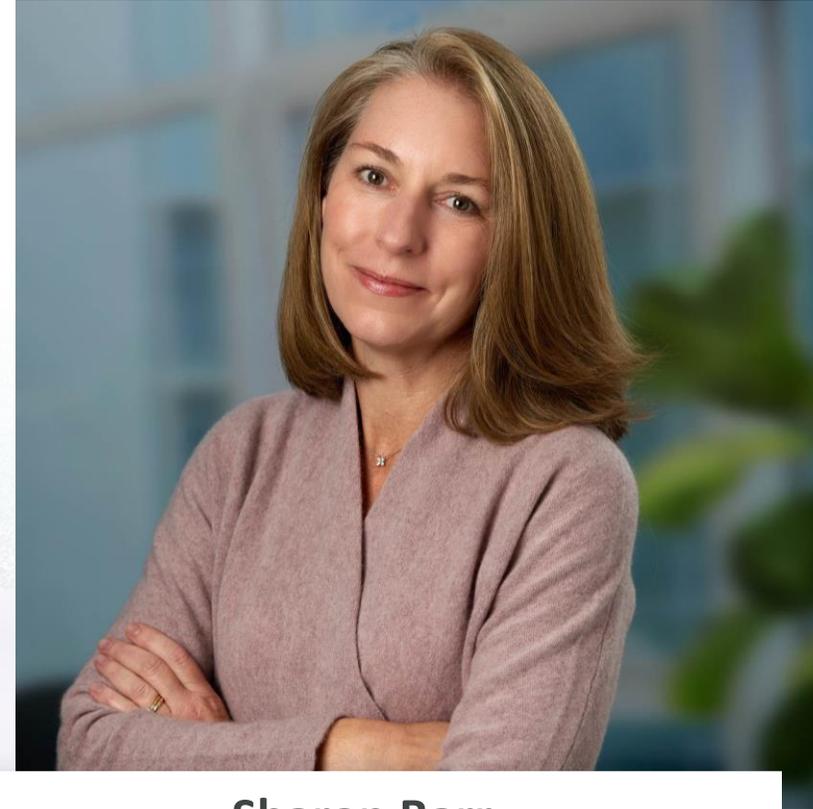
1. Indicates Company ambition to achieve Total Revenue low double-digit CAGR through 2025 (2021 base year, Alexion pro-forma) and industry-leading Total Revenue beyond 2025; this is not formal guidance. 1L = 1st-line; mTNBC = metastatic triple negative breast cancer; PIK3/AKT/PTEN = phosphatidylinositol-3 kinase/protein kinase B/phosphatase and tensin homolog; EGPA = eosinophilic granulomatosis with polyangiitis; 2L+ = 2nd-line plus; HR+ = hormone receptor positive; HER2- = human epidermal growth factor receptor 2 negative; mBC = metastatic breast cancer; Stg. = stage; unresec. = unresectable; NSCLC = non-small cell lung cancer; cCRT = concurrent chemoradiation therapy; HCC = hepatocellular carcinoma; PFS = progression-free survival; TACE = trans arterial chemoembolisation; LAAB = long-acting antibody; NME = new molecular entity.



BioPharmaceuticals R&D leadership change



Mene Pangalos
to retire after almost 14 years
with AstraZeneca



Sharon Barr
to assume role of EVP,
BioPharmaceuticals R&D

Question & Answer Session



Pascal Soriot
EXECUTIVE DIRECTOR &
CHIEF EXECUTIVE OFFICER



Aradhana Sarin
EXECUTIVE DIRECTOR &
CHIEF FINANCIAL OFFICER



Marc Dunoyer
CHIEF EXECUTIVE OFFICER,
ALEXION



Susan Galbraith
EXECUTIVE VICE PRESIDENT,
ONCOLOGY R&D



Dave Fredrickson
EXECUTIVE VICE PRESIDENT,
ONCOLOGY BUSINESS



Mene Pangalos
EXECUTIVE VICE PRESIDENT,
BIOPHARMACEUTICALS R&D



Ruud Dobber
EXECUTIVE VICE PRESIDENT,
BIOPHARMACEUTICALS
BUSINESS



Iskra Reic
EXECUTIVE VICE PRESIDENT,
VACCINES AND IMMUNE
THERAPIES



Leon Wang
EXECUTIVE VICE PRESIDENT,
INTERNATIONAL



Appendix

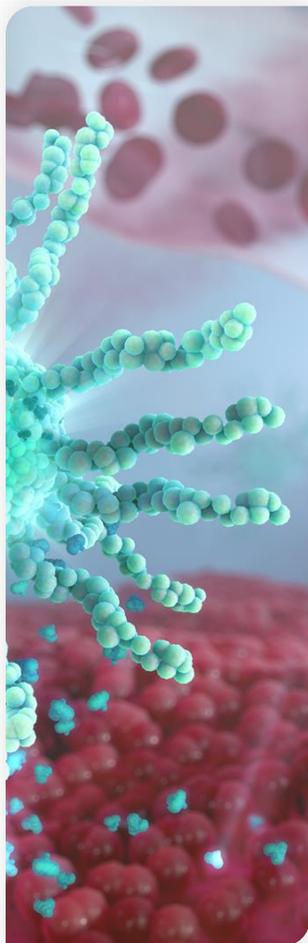
- Pipeline Highlights
- Oncology tumour maps
- Emerging Markets performance
- Key medicines performance by therapy area



Delivering on science-led innovation

Selected key pipeline highlights since Q1 2023 results

Oncology BioPharmaceuticals Rare Disease



10 regulatory approvals in major markets, including:

Lynparza (US)

prostate cancer (1st-line) (PROpel)

Enhertu (CN)

HER2-low breast cancer (3rd-line)

Farxiga (US)

HFpEF (DELIVER)

Beyfortus (US)

RSV (MELODY/MEDLEY)

Xigduo (CN)

type-2 diabetes (XR formulation)

Soliris (EU)

generalised myasthenia gravis
(refractory, children and adolescents)

Soliris (CN)

generalised myasthenia gravis

Ultomiris (EU, JP)

neuromyelitis optica spectrum disorder
(CHAMPION-NMO)

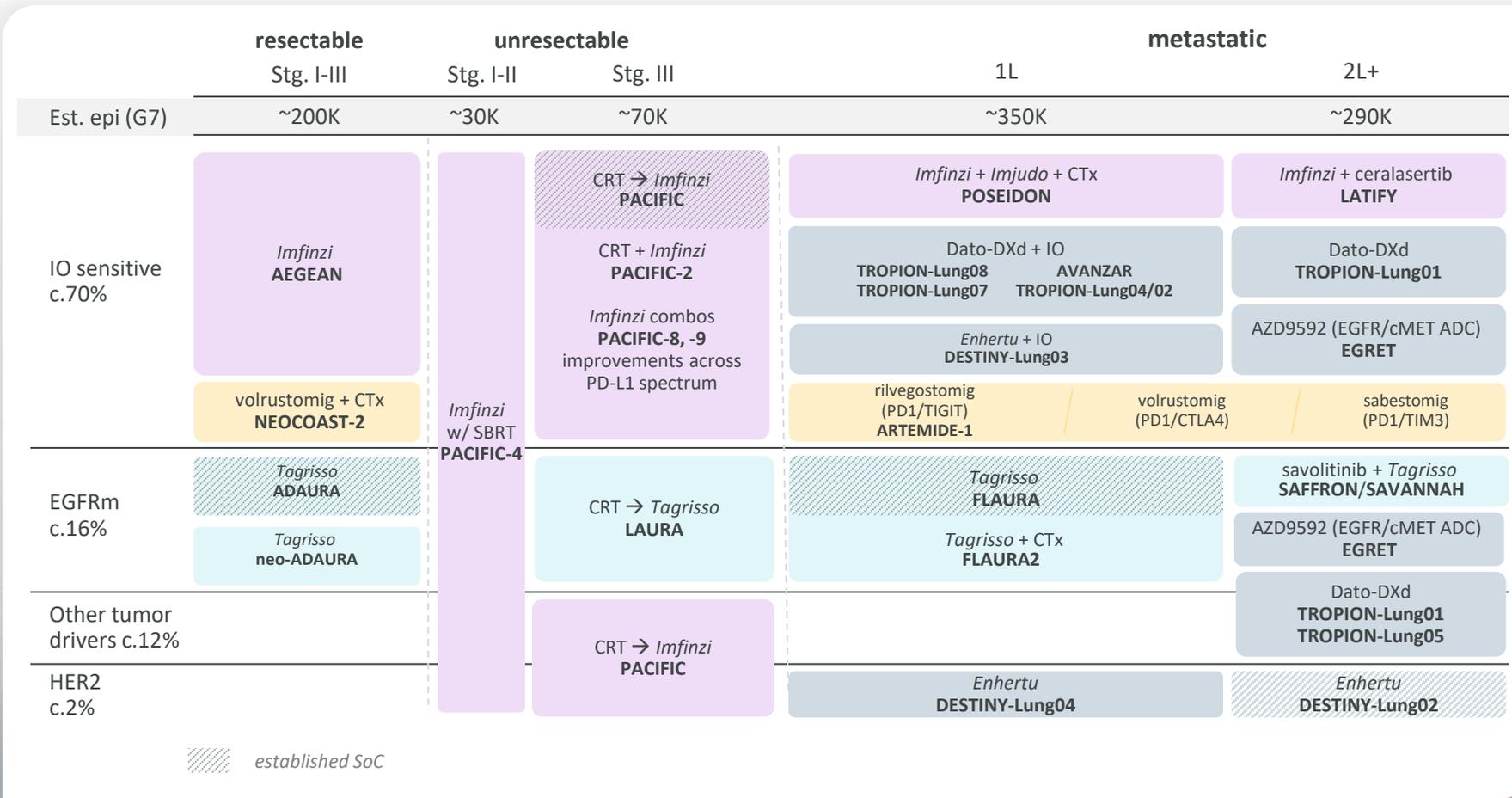
Koselugo (CN)

neurofibromatosis type-1 with plexiform
neurofibromas (SPRINT)



AstraZeneca in Lung Cancer

Ambition for >50% of lung cancer patients to be eligible for AZN medicine by 2030



Leading the future of lung cancer treatment

- *Tagrisso* established TKI backbone in EGFRm
- *Imfinzi* leading IO in unresectable
- Advancing best-in-class ADCs to replace systemic chemotherapy
- Delivering next-wave bispecifics to improve on PD1/PD-L1
- Developing novel combinations, including IO + ADC
- Investing behind new technologies and platforms, including cell therapy, testing/screening

Est epi (G7) = estimated epidemiology across G7 (US, EU5, JP); Stg. = stage; CTx = chemotherapy; SBRT = stereotactic body radiation therapy; CRT = chemoradiotherapy; pembro = pembrolizumab; IO = immunotherapy; ADC = antibody-drug conjugate; PD1 = programmed cell death protein 1; EGFR = epidermal growth factor receptor; c-MET = mesenchymal-epithelial transition factor; TIGIT = T-cell immunoreceptor with immunoglobulin and ITIM domains; CTLA4 = cytotoxic T-lymphocyte associated protein 4; TIM3 = T-cell immunoglobulin and mucin domain-containing protein 3; SoC = standard of care; TKI = tyrosine kinase inhibitor.
 Collaboration partners: Daiichi Sankyo (*Enhertu*, Dato-DXd), CompuGen (rilvegostomig).



AstraZeneca in Breast Cancer

Ambition to eliminate breast cancer as a cause of death

established SoC	Early		RECURRENT	Metastatic			
	Noadjuvant	Adjuvant		1st line	2nd line	3rd line	4th line +
Est. epi (G7)	540k			125k	90k	65k	55k
HER2-positive 15-20%	<i>Enhertu</i> +/- THP DESTINY-Breast11	NST → residual disease → <i>Enhertu</i> DESTINY-Breast05		<i>Enhertu</i> DESTINY-Breast09	<i>Enhertu</i> DESTINY-Breast03	<i>Enhertu</i> DESTINY-Breast02	
HR-positive 65-75% --- <i>HER2-low</i> 60%		Low risk Current SoC drives good outcomes for patients with low risk HR-positive eBC CTx → AI (+/- CDK4/6i) → camizestrant CAMBRIA-1		camizestrant + CDK4/6i SERENA-4 ESR1m CDK4/6i + AI → CDK4/6i + camizestrant SERENA-6 capiasertib + <i>Faslodex</i> + CDK4/6i CAPitello292	capiasertib + <i>Faslodex</i> CAPitello291 <i>HER2-low</i> <i>Enhertu</i> DESTINY-Breast06	Dato-DXd TROPION-Breast01 <i>HER2-low</i> <i>Enhertu</i> DESTINY-Breast04	
TNBC 10-15% --- <i>HER2-low</i> 35%		NST → residual disease → Dato-DXd +/- <i>Imfinzi</i> TROPION-Breast03		capiasertib + paclitaxel CAPitello290 PD-L1- 60% Dato-DXd TROPION-Breast02	<i>HER2-low</i>		
gBRCAm 5% of HR-positive 15% of TNBC		CTx → <i>Lynparza</i> OlympiA			<i>Lynparza</i> OlympiAD		

All numbers are approximate. Illustrative settings and populations, not to scale.

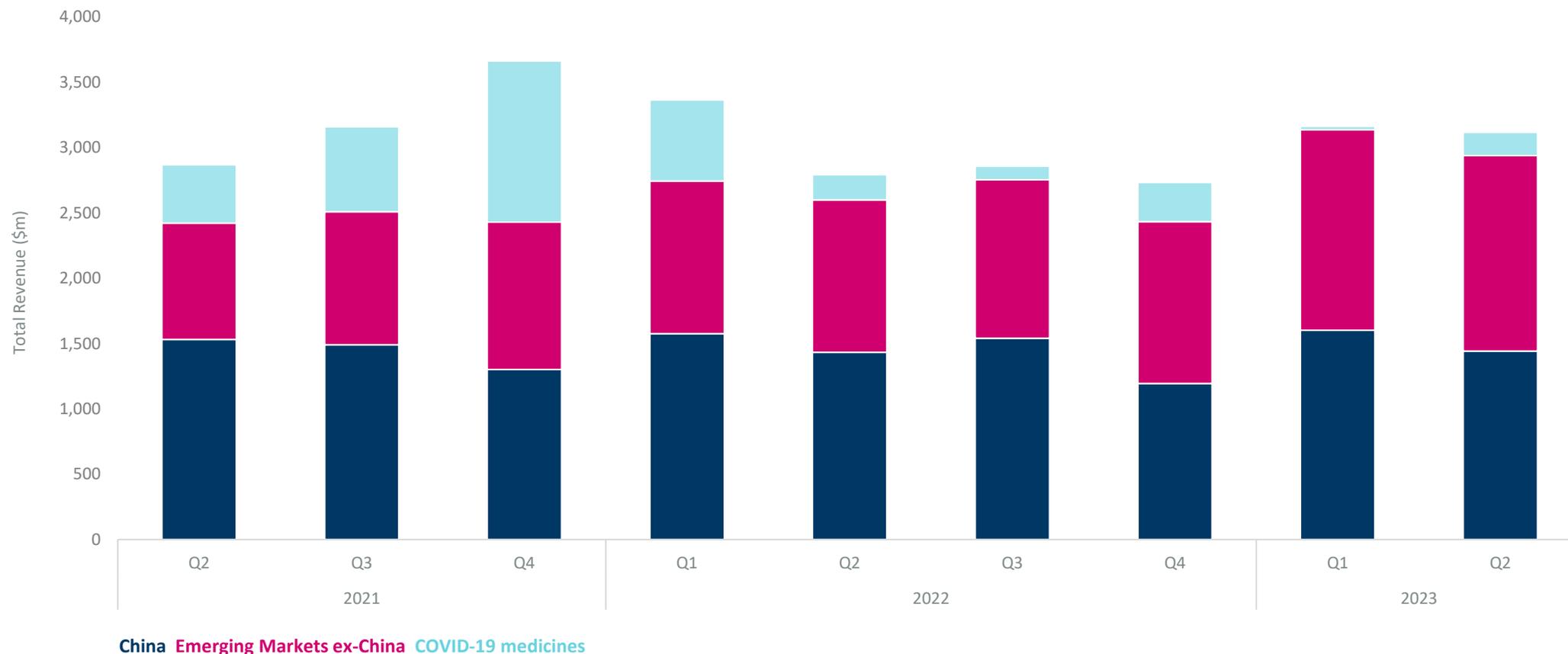
1/2/3/4L = 1st/2nd/3rd/ 4th-line; est epi (G7) = estimated epidemiology across G7 (US, EU5, JP for drug treated patients). HER2 = human epidermal growth factor receptor 2; THP = docetaxel, trastuzumab, and pertuzumab; NST = neoadjuvant systemic treatment; HR = hormone receptor; SoC = standard of care; CTx = chemotherapy; AI = aromatase inhibitor; CDK4/6i = cyclin-dependent kinase 4 and 6 inhibitor; ESR1m = oestrogen receptor 1 gene mutation; Dato-DXd = datopotamab deruxetecan; TNBC = triple negative breast cancer; PD-L1 = programmed cell death ligand 1; gBRCAm = germline BRCA-mutated.

Collaboration partners: Daiichi Sankyo (*Enhertu*, Dato-DXd), Merck & Co., Inc. (*Lynparza*).



Emerging Markets – H1 2023

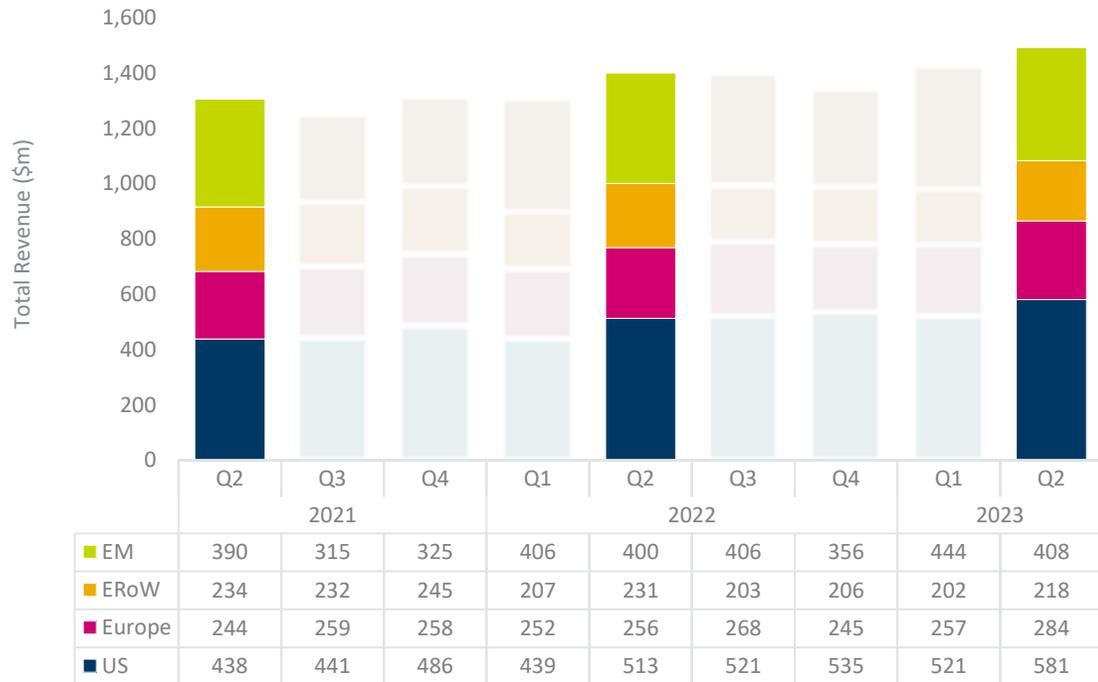
Total Revenue +9% at CER to \$6.3bn, +22% at CER ex-COVID-19 medicines



Oncology

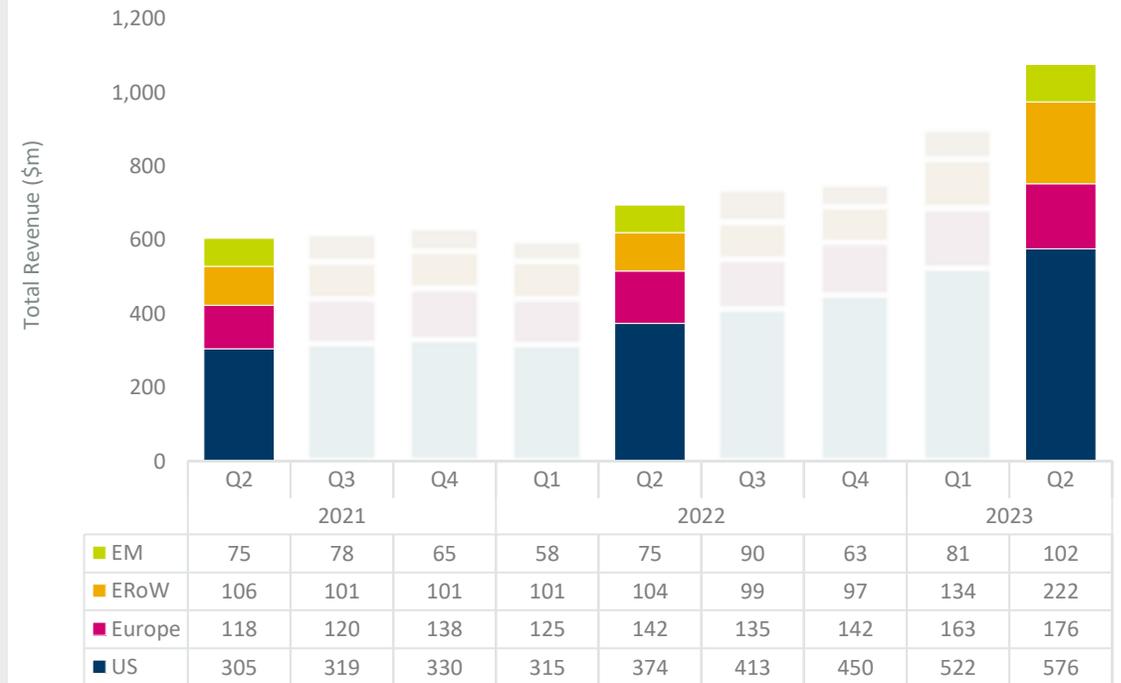
Tagrisso

12% growth at CER to \$2,915m in H1 2023



Imfinzi

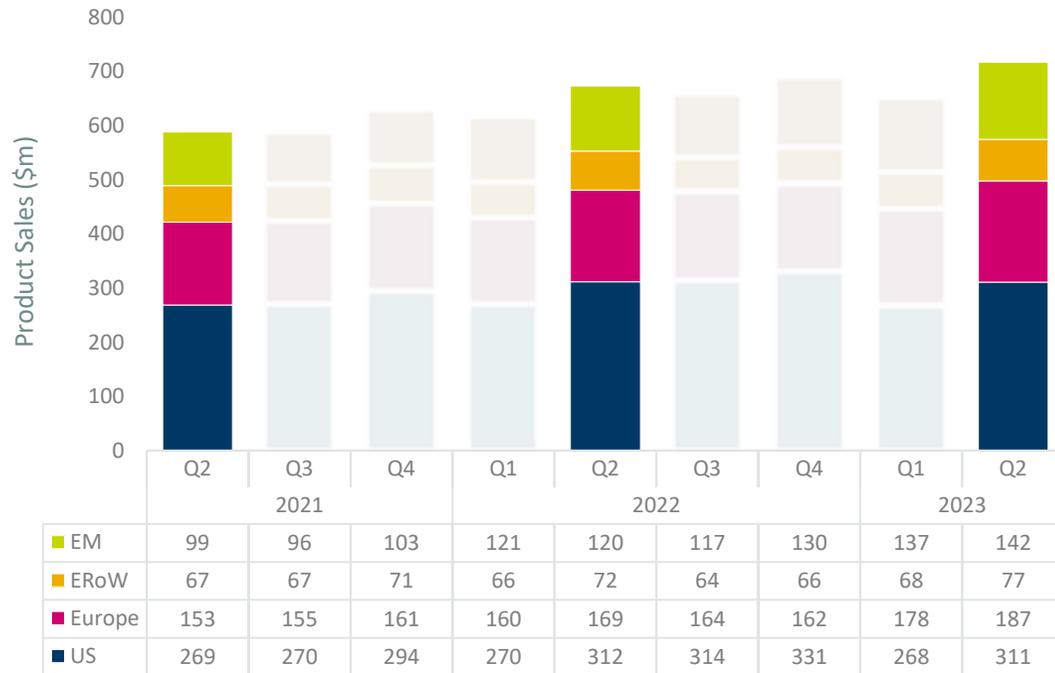
57% growth at CER to \$1,976m in H1 2023



Oncology

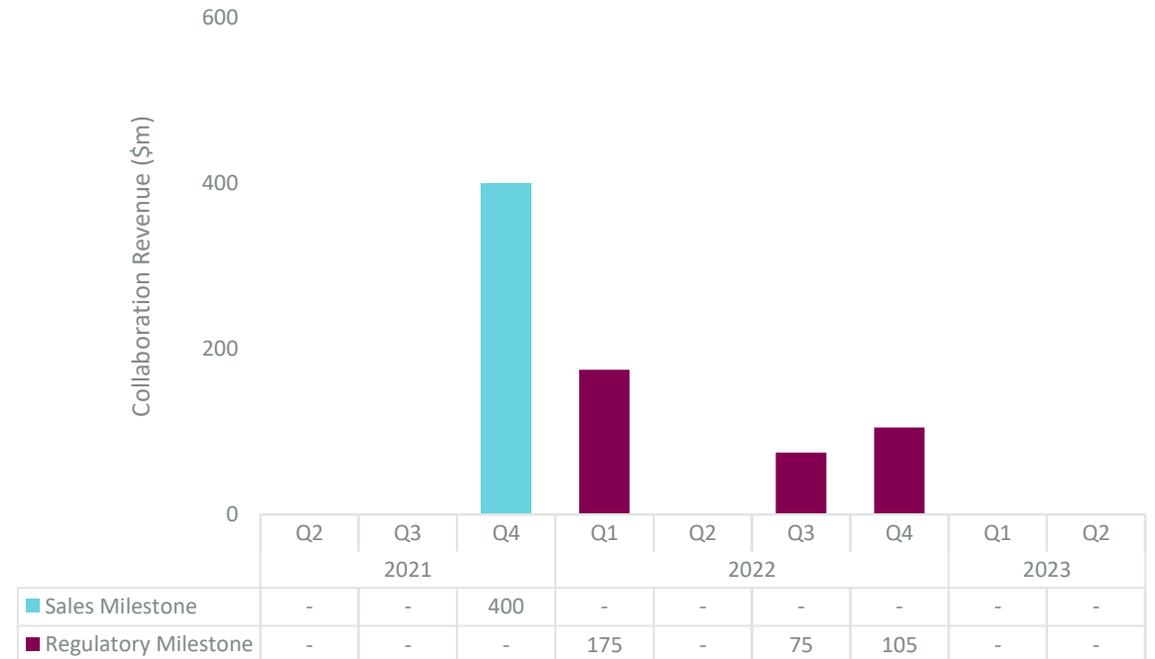
Lynparza

10% growth at CER to \$1,368m in H1 2023



Lynparza

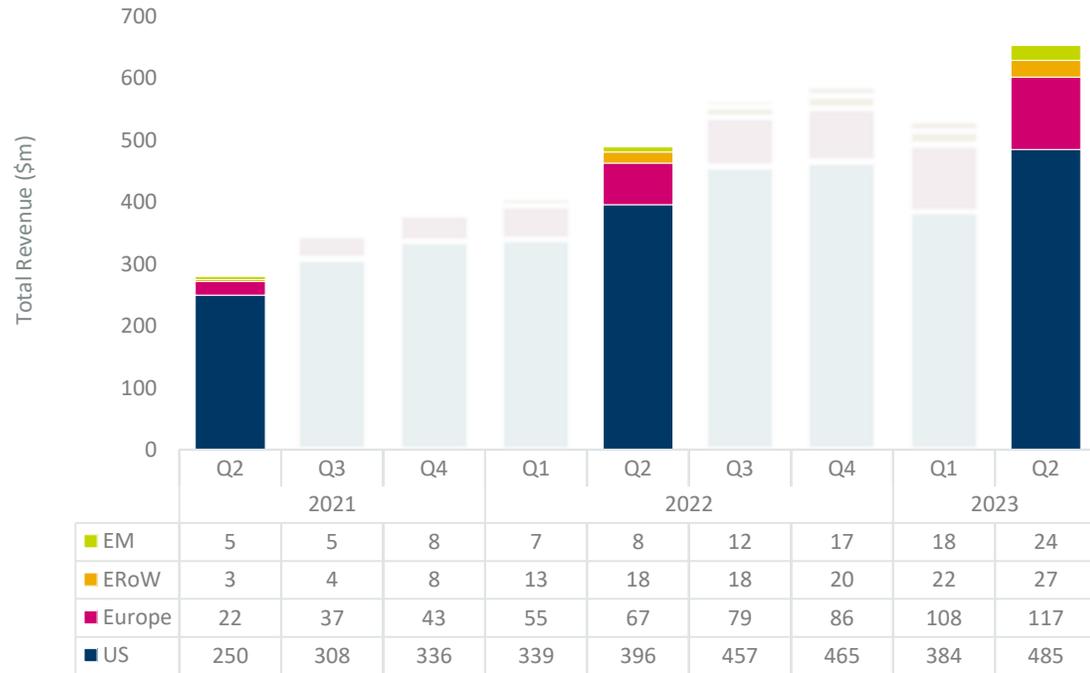
Collaboration Revenue: \$3.8bn recorded cumulative, \$3.9bn future potential



Oncology

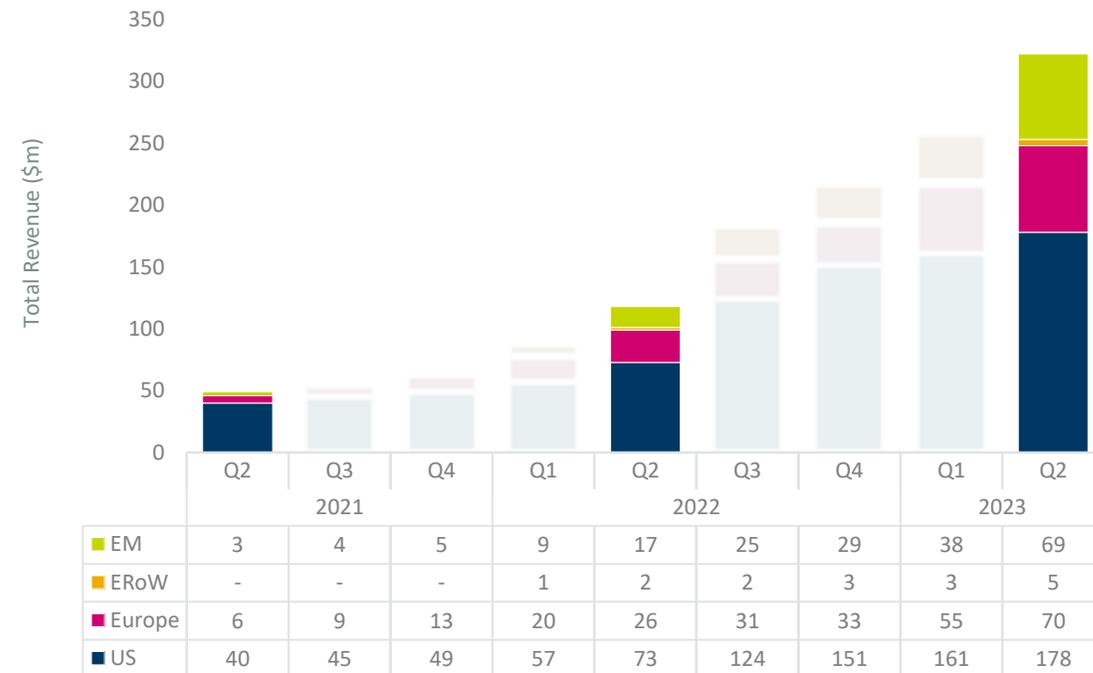
Calquence

33% growth at CER to \$1,185m in H1 2023



Enhertu

>2x growth at CER to \$580m in H1 2023



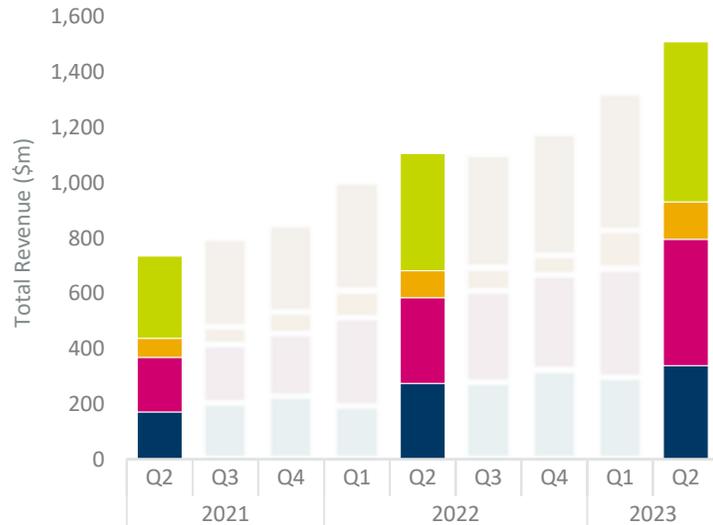
Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.



BioPharmaceuticals: Cardiovascular, Renal & Metabolism

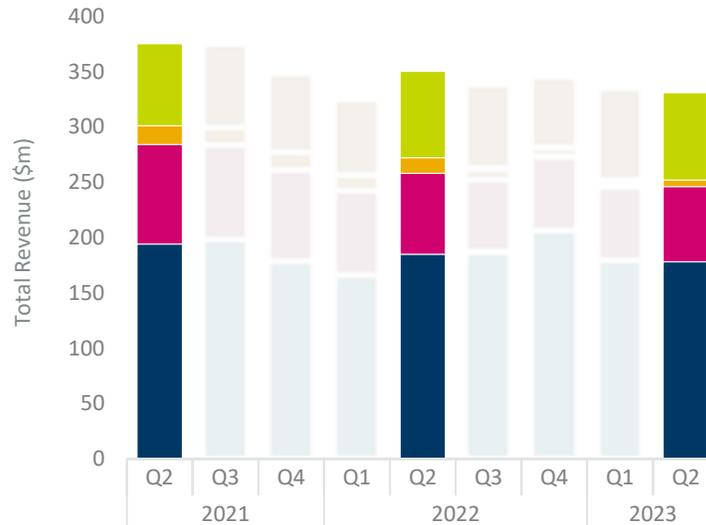
Farxiga

40% growth at CER to \$2,834m in H1 2023



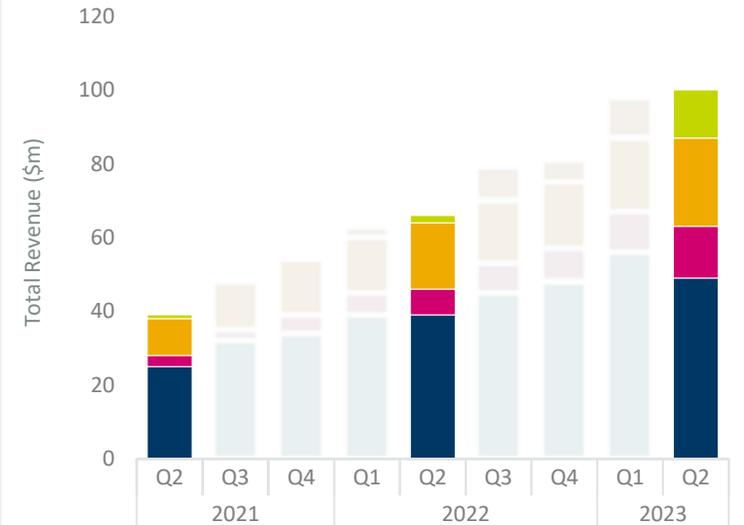
Brilinta

1% growth at CER to \$665m in H1 2023



Lokelma

59% growth at CER to \$198m in H1 2023



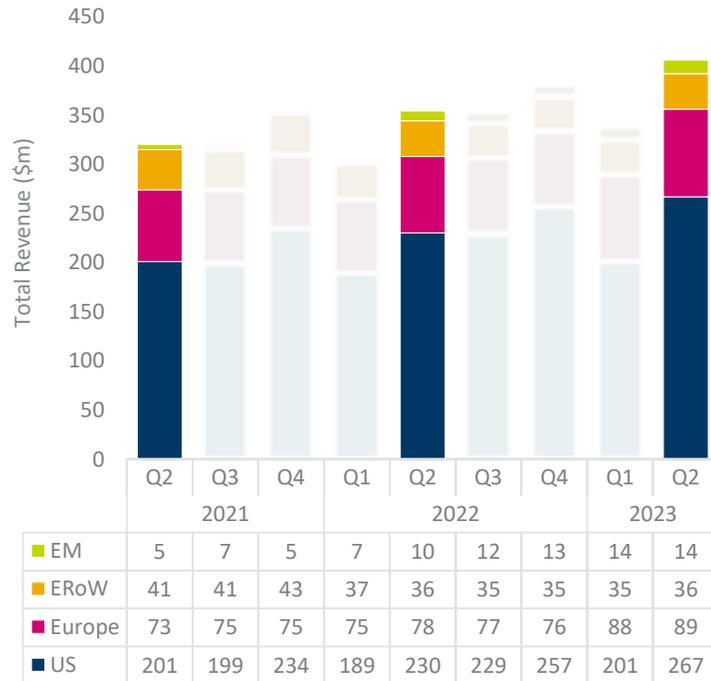
41 Due to rounding, the sum of a number of dollar values and percentages may not agree to totals. CER = constant exchange rates; EM = Emerging Markets; ERoW = Established Rest of World.



BioPharmaceuticals: Respiratory & Immunology

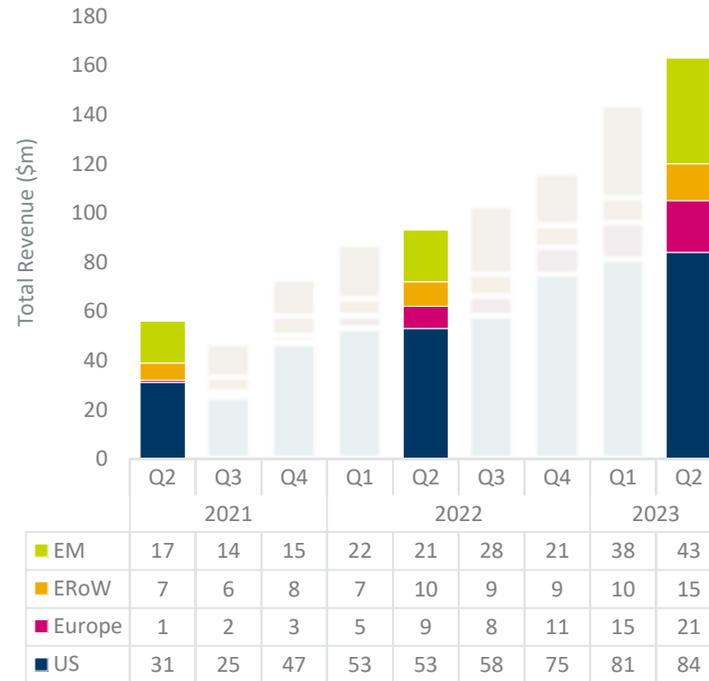
Fasenra

14% growth at CER to \$744m in H1 2023



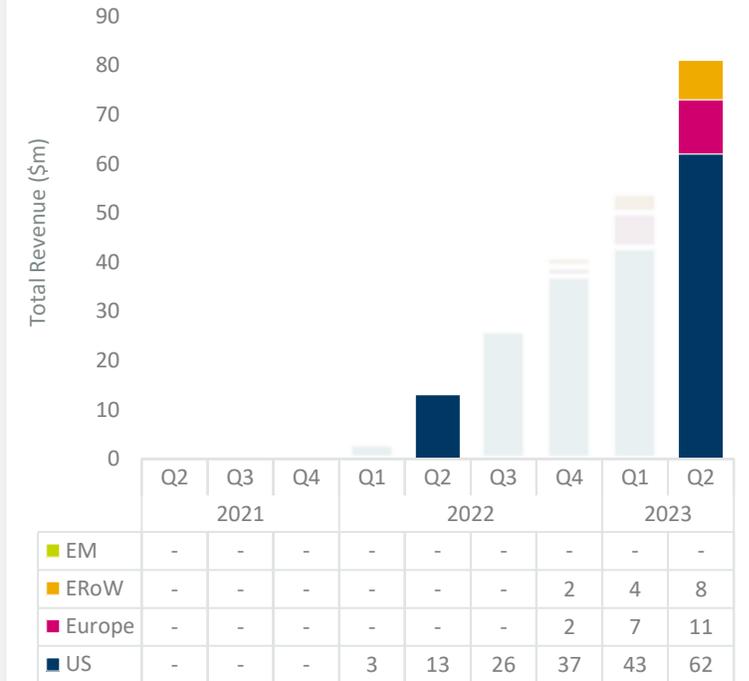
Breztri

76% growth at CER to \$307m in H1 2023



Tezspire

>8x growth at CER to \$135m in H1 2023



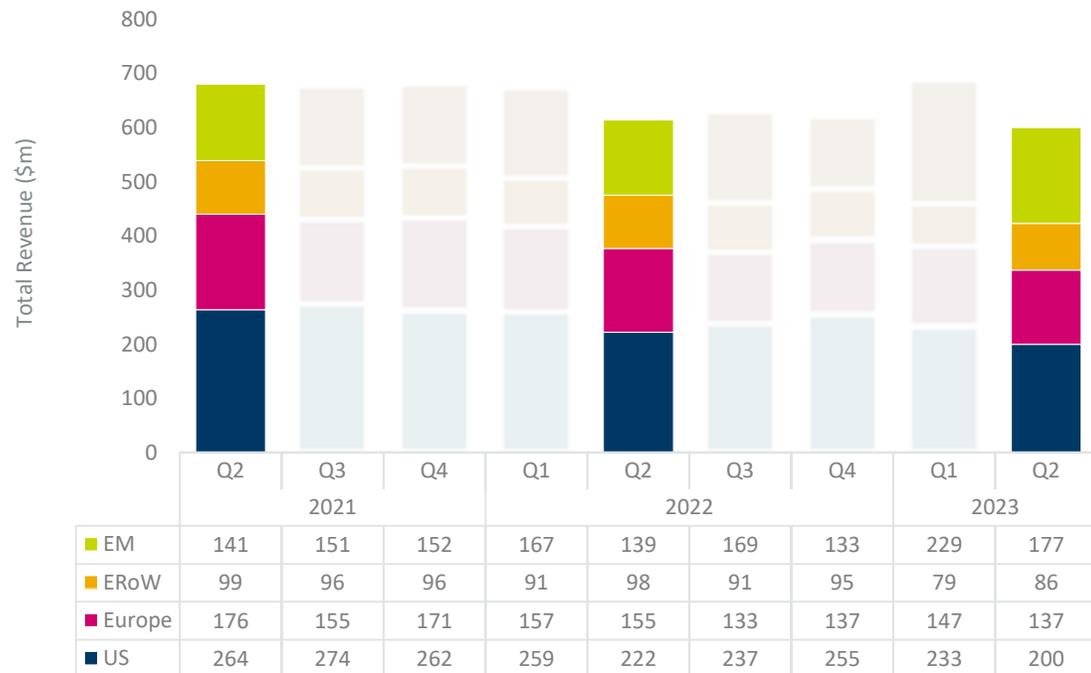
Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.



BioPharmaceuticals: Respiratory & Immunology

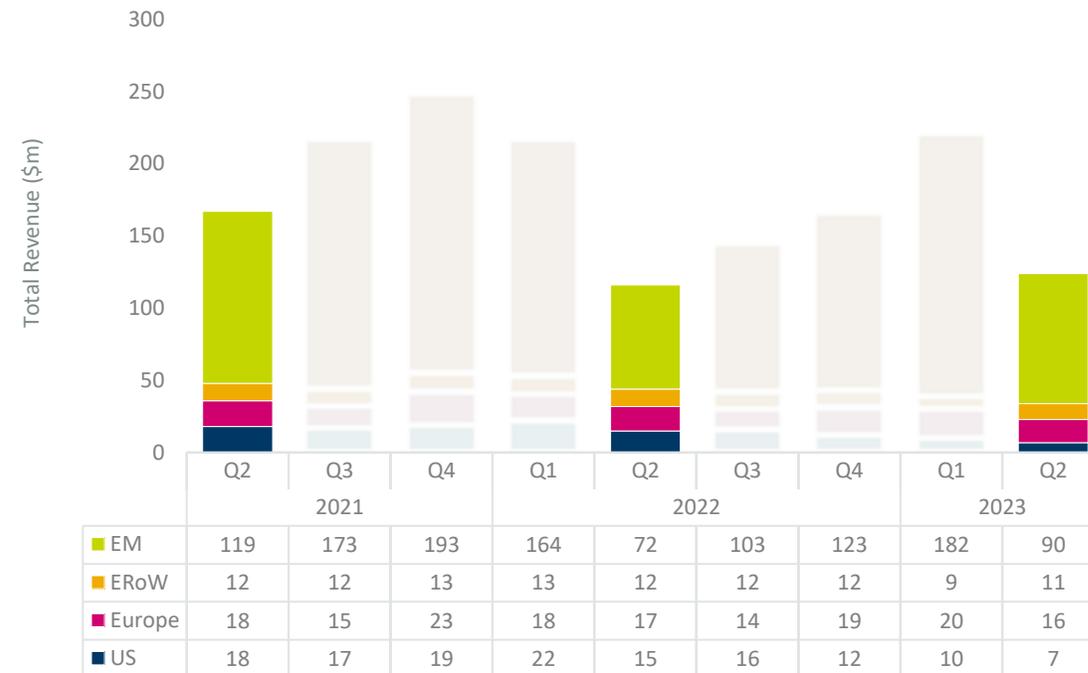
Symbicort

4% growth at CER to \$1,288m in H1 2023



Pulmicort

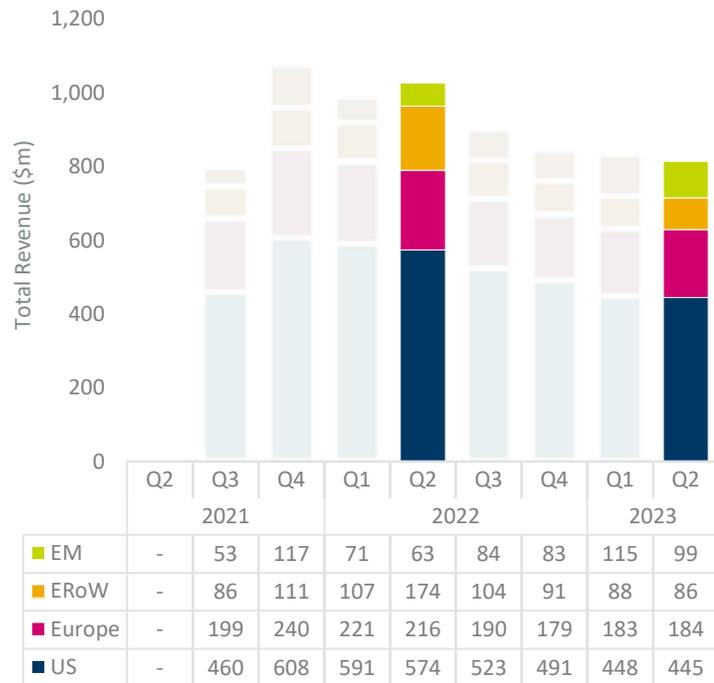
11% growth at CER to \$346m in H1 2023



Rare Disease

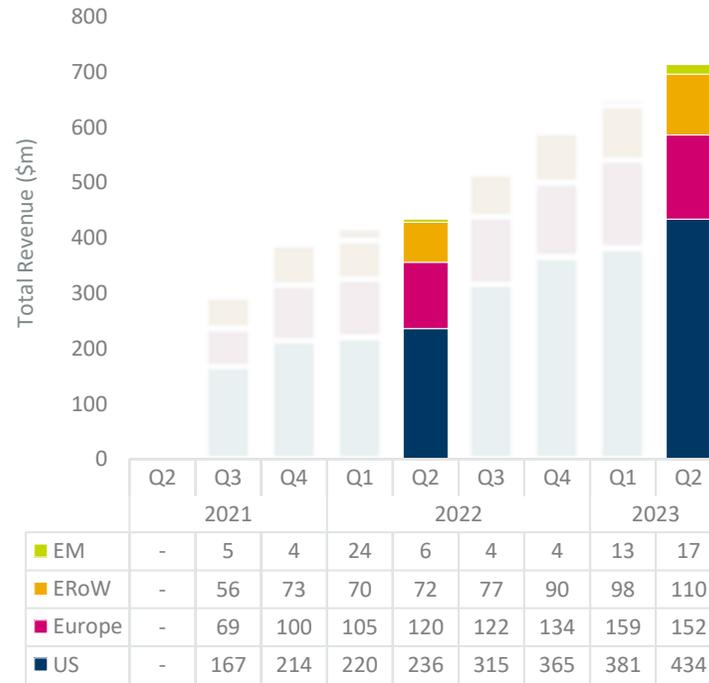
Soliris

16% decrease at CER to \$1,648m in H1 2023



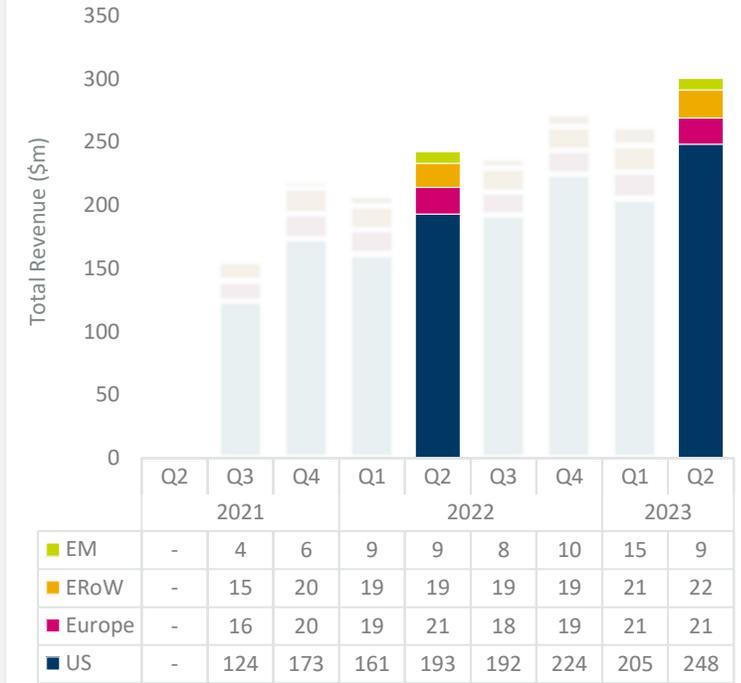
Ultomiris

64% growth at CER to \$1,364m in H1 2023



Strensiq

26% growth at CER to \$562m in H1 2023



Due to rounding, the sum of a number of dollar values and percentages may not agree to totals.

