Roche Group development pipeline

Marketed products development programmes

Roche Pharma global development programmes

Roche Pharma research and early development (pRED)

Genentech research and early development (gRED)

Spark

Changes to the development pipeline

HY 2022 update

New to phase I

2 NMEs:
RG6351 NME – retinal disease
RG6526 camonsertib – solid tumors

1 AI:
RG6264 Phesgo OBI - HER2+ BC

New to phase II

1 NME:
RG6237 latent myostatin + Evrysdi – SMA

New to phase III

4 AIs:
RG1594 Ocrevus SC - PPMS & RMS
RG6171 giredestrant + Phesgo – 1L ER+/HER2+ BC
RG1450 gantenerumab – early Alzheimer's
RG7828 Lunsumio (mosunetuzumab) + Polivy - 2L+ SCT ineligible DLBCL

New to registration

Removed from phase I

1 NME:
RG6338 NME – metabolic diseases

2 AIs:
RG7440 ipatasertib + rucaparib - mCRPC, solid tumors
RG7440 ipatasertib - prostate cancer, pretreated

Removed from phase II

1 NME:
RG6173 anti-tryptase - asthma

1 AI:
RG6171 giredestrant – 2/3L ER+/HER2- mBC

Removed from phase III

Approvals

1 NME (EU):
RG7828 Lunsumio (mosunetuzumab) - 3L FL

1 AI (US):
RG7916 Evrysdi SMA presymptomatic pediatric <2mo

2 AIs (EU):
RG7596 Polivy – 1L DLBCL
RG7446 Tecentriq - NSCLC adj

Status as of July 21, 2022

Roche Group development pipeline

Phase I (49 NMEs + 11 AIs)

RG6007	HLA-A2-WT1 x CD3	AML
RG6026	glofitamab monotherapy + combos	heme tumors
RG6058	tiragolumab combos	heme & solid tumors
RG6076	CD19-4-1BBL combos	heme tumors
RG6129	HLA-A2-MAGE-A4 x CD3	solid tumors
RG6160	cevostamab (FcRH5 x CD3)	r/r multiple myeloma
RG6171	giredestrant (SERD)	solid tumors
RG6114	inavolisib (mPI3K alpha inh)	solid tumors
RG6156	EGFRvIII x CD3	glioblastoma
RG6180	autogene cevumeran ± T	solid tumors
RG6185	belvarafenib (pan-RAF inh) + Cotellic ± T	solid tumors
RG6189	FAP-CD40 ± T	solid tumors
RG6194	runimotamab (HER2 x CD3)	BC
RG6234	GPRC5D x CD3	multiple myeloma
RG6264	Phesgo OBI	HER2+ BC
RG6279	PD1-IL2v ± T	solid tumors
RG6286	-	colorectal cancer
RG6290	MAGE-A4 ImmTAC ± T	solid tumors
RG6292	CD25 MAb ± T	solid tumors
RG6323	IL15/IL15Ra-Fc ± T	solid tumors
RG6330	KRAS G12C	solid tumors
RG6333	CD19 x CD28 + glofitamab	r/r NHL
RG6344	BRAF inhibitor (3)	solid tumors
RG6392	-	oncology
RG6433	SHP2i	solid tumors
RG6440	TGFβ (SOF10)	solid tumors
RG6526**	camonsertib	solid tumors
RG7446	Morpheus platform	solid tumors
RG7601	Venclexta ± azacitidine	r/r MDS
RG7802	cibisatamab ± T	solid tumors
RG7827	FAP-4-1BBL monotherapy + combos	solid tumors
RG7828	Lunsumio (mosunetuzumab) monotheraphy + combos	heme tumors
CHU	FIXa x FX	hemophilia
CHU	glypican-3 x CD3	solid tumors
CHU	codrituzumab	HCC
CHU	CD137 switch antibody	solid tumors
CHU	LUNA18	solid tumors
CHU	SPYK04	solid tumors
SQZ	PBMC vaccine	solid tumors
RG6287	-	IBD
RG6341	-	asthma
RG6418	selnoflast (NLRP3 inh)	inflammation
RG6315	-	immunologic disorders
RG7828	Lunsumio (mosunetuzumab)	SLE
RG7880	efmarodocokin alfa	aGVHD
RG6006	Abx MCP	bacterial infections
RG6319	LepB inhibitor	complicated urinary tract infection
RG6035	BS-CD20 MAb	multiple sclerosis
RG6091	rugonersen (UBE3A LNA)	Angelman syndrome
RG6163	-	psychiatric disorders
RG6182	-	neurodegenerative diseases
RG6237	latent myostatin	neuromuscular disorders
RG6289	-	Alzheimer´s
RG7637	-	neurodevelopmental disorders
RG6120	VEGF-Ang2 DutaFab	nAMD
RG6312	-	geographic atrophy
RG6351	NME	retinal disease
RG6501*	OpRegen	geographic atrophy
RG7921	-	nAMD
CHU	AMY109	endometriosis

Phase II (22 NMEs + 11 AIs)

RG6026	glofitamab + chemo	1L ctDNA high risk DLBCL
RG6058	tiragolumab + T	NSCLC
	tiragolumab + T + chemo	1L non-squamous NSCLC
	tiragolumab + T + chemo	NSCLC neoadj-adj
	tiragolumab + T	cervical cancer
	tiragolumab + T	1L PD-L1+ mSCCHN
RG6107	crovalimab	sickle cell disease
RG6139	PD1 x LAG3	solid tumors
RG6180	autogene cevumeran + pembrolizumab	1L melanoma
RG6354	zinpentraxin alfa (PRM-151)	myelofibrosis
RG6357	SPK-8011	hemophilia A
RG6358	SPK-8016	hemophilia A with inhibitors to factor VIII
RG7601	Venclexta + carfilzomib	r/r MM t(11;14)
CHU	Oncolytic Type 5 adenovirus	esophageal cancer
RG6149	astegolimab (Anti-ST2)	COPD
RG6299†	ASO factor B	IgA nephropathy
RG7854/RG7907/RG6346/RG6084[1]	TLR7 ago(3)/CpAM (2)/ siRNA/PDL1 LNA	HBV
RG6359	SPK-3006	Pompe disease
RG6100	semorinemab	Alzheimer's
RG6102	BS-gantenerumab	Alzheimer's
RG6237	latent myostatin + Evrysdi	SMA
RG6416	bepranemab	Alzheimer's
RG7412	crenezumab	familial Alzheimer's healthy pts
RG7816	alogabat (GABA Aa5 PAM)	ASD
RG7906	ralmitaront	schizophrenia
RG7935	prasinezumab	Parkinson's
RG6147	galegenimab (HtrA1)	geographic atrophy
RG6179	-	DME
RG7774	-	retinal disease
RG6299†	ASO factor B	geographic atrophy

New Molecular Entity (NME)
Additional Indication (AI)
Oncology / Hematology
Immunology
Infectious Diseases

Metabolism
Neuroscience
Ophthalmology
Other

CHU - Chugai managed
†IONIS managed
SQZ - SQZ Biotechnology managed
*Lineage Cell Therapeutics managed
**Repare Therapeutics managed

[1]combination platform
RG-No - Roche/Genentech
T=Tecentriq
BS=Brain Shuttle
OBI=On-Body Delivery System

Status as of July 21, 2022

Roche Group development pipeline

Phase III (10 NMEs + 43 AIs)

RG3502	Kadcyla + T	2L+ HER-2+ PD-L1+ mBC
	Kadcyla + T	HER-2+ eBC high-risk
RG6026	glofitamab + chemo	2L+ DLBCL
RG6058	tiragolumab + T	1L esophageal cancer
	tiragolumab + T	1L PD-L1+ NSCLC
	tiragolumab + T	locally advanced esophageal cancer
	tiragolumab + T	stage III unresectable 1L NSCLC
RG6107	crovalimab	PNH
	crovalimab	aHUS
RG6114	inavolisib (mPI3K alpha inh)	1L HR+ mBC
RG6171	giredestrant (SERD)	1L ER+/HER2- mBC
	giredestrant (SERD)	ER+ BC adj
	giredestrant (SERD) + Phesgo	1L ER+/HER2+ BC
RG7440	ipatasertib + abiraterone	1L CRPC
RG7446	Tecentriq + platinum chemo	NSCLC neoadj
	Tecentriq	NMIBC, high risk
	Tecentriq	RCC adj
	Tecentriq + cabozantinib	RCC adv
	Tecentriq + cabozantinib	2L NSCLC
	T ± chemo	SCCHN adj
	T + capecitabine or carbo/gem	1L TNBC
	T + paclitaxel	TNBC adj
	T + Avastin	HCC adj
	T ± chemo	1L mUC
	Tecentriq SC	2L NSCLC
	Tecentriq	ctDNA+ high-risk MIBC
	T+ lurbinectedin	1L maintenance SCLC

RG7601	Venclexta	r/r MM t(11:14)
	Venclexta + azacitidine	1L MDS
RG7828	Lunsumio (mosunetuzumab) + lenalidomide	2L+ FL
	Lunsumio (mosunetuzumab) + Polivy	2L+ DLBCL
RG7853	Alecensa	ALK+ NSCLC adj
RG3648	Xolair	food allergy
RG6354	zinpentraxin alfa (PRM-151)	IPF
RG7159	Gazyva	lupus nephritis
	Gazyva	membranous nephropathy
	Gazyva	systemic lupus erythematosus
RG6152	Xofluza	influenza, pediatric (0-1 year)
	Xofluza	influenza direct transmission
RG1450	gantenerumab	prodromal to mild Alzheimer's
	gantenerumab	early Alzheimer's
RG1594	Ocrevus higher dose	RMS & PPMS
	Ocrevus SC	RMS & PPMS
RG6042	tominersen	Huntington's
RG6168	Enspryng	myasthenia gravis
RG6356	delandistrogene moxeparvovec (SRP-9001)	DMD
RG7845	fenebrutinib	RMS
RG7845	fenebrutinib	PPMS
RG6321	Susvimo (PDS)	DME
	Susvimo (PDS)	DR
	Susvimo (PDS)	wAMD, 36-week
RG7716	Vabysmo (faricimab)	BRVO
	Vabysmo (faricimab)	CRVO

Registration US & EU (4 NMEs + 8 AIs)

RG6013	Hemlibra[1]	mild to moderate hemophilia A
RG6026	glofitamab[2]	3L+ DLBCL
RG6396	Gavreto[1]	RET+ MTC, TC
RG7596	Polivy[3]	1L DLBCL
RG7828	Lunsumio (mosunetuzumab)[4]	3 L+ FL
RG6321	Susvimo (PDS) [1]	wAMD
RG7716	Vabysmo (faricimab) [1]	DME
	Vabysmo (faricimab) [1]	wAMD
RG6152	Xofluza	influenza, pediatric
RG56413+ RG6412	Ronapreve[2]	SARS-CoV-2 hospitalised
RG1569	Actemra[4]	COVID-19 pneumonia
RG7916	Evrysdi[1]	SMA pediatric <2months

[1] Approved in US, filed in EU
[2] Filed in the EU
[3] Approved in EU
[4] Approved in EU, filed in US

T=Tecentriq
PDS=Port Delivery System with ranibizumab

New Molecular Entity (NME)
Additional Indication (AI)
Oncology / Hematology
Immunology
Infectious Diseases
Metabolism
Neuroscience
Ophthalmology
Other

Status as of July 21, 2022

NME submissions and their additional indications

Projects in phase II and III

New Molecular Entity (NME)
Additional Indication (AI)
Oncology / Hematology
Immunology
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Ophthalmology
Other

✓ Indicates submission to health authorities has occurred
Unless stated otherwise submissions are planned to occur in US and EU
PDS=Port Delivery System with ranibizumab
Mosun=mosunetuzumab
†IONIS managed

2022

RG6026	**glofitamab** 3L+ DLBCL ✓
RG6058	**tiragolumab + T** 1L esophageal cancer (CN)
RG6107	**crovalimab** PNH (CN)
RG1450	**gantenerumab** prodromal to mild Alzheimer‘s

2023

RG6058	**tiragolumab + T** 1L PD-L1+ NSCLC
RG6321	**Susvimo (PDS)** DME
RG6321	**Susvimo (PDS)** DR (US)
RG7716	**Vabysmo (faricimab)** BRVO/CRVO

2024

RG6058	**tiragolumab + T** Stage III unresectable 1L NSCLC
RG6107	**crovalimab** aHUS
RG6114	**inavolisib (mPI3K alpha inh)** 1L HR+ BC
RG6354	**zinpentraxin alfa (PRM-151)** IPF
RG6356	**delandistrogene moxeparvovec (SRP-9001)** DMD

2025 and beyond

RG6026	**glofitamab + chemo** 2L DLBCL
RG6026	**glofitamab + chemo** 1L ctDNA+ high risk DLBCL
RG6058	**tiragolumab + T** 1L PD-L1+ cervical cancer
RG6058	**tiragolumab + T** locally adv esophageal cancer
RG6058	**tiragolumab + T** 1L non-sq NSCLC
RG6058	**tiragolumab + T** 1L PD-L1+ mSCCHN
RG6058	**tiragolumab+T+/- chemo** NSCLC neoadj/adj
RG6107	**crovalimab** sickle cell disease
RG6139	**PD1xLAG3** solid tumors
RG6171	**giredestrant (SERD)** 1L ER+/HER2- mBC
RG6171	**giredestrant (SERD)** ER+ BC adj
RG6171	**giredestrant (SERD) + Phesgo** 1L ER+/HER2+ BC
RG6180	**autogene cevumeran** 1L melanoma
RG6354	**zinpentraxin alfa (PRM-151)** myelofibrosis
RG7828	**Lunsumio (mosun) + lenalidomide** 2L FL
RG7828	**Lunsumio (mosun) + Polivy** 2L+ DLBCL (US)
RG6149	**astegolimab (anti-ST2)** COPD
RG6299†	**ASO factor B** IgA nephropathy
RG7907/ RG7854/ RG6346/ RG6084	**TLR7 ago (3)/CpAM (2) /siRNA/ PDL1 LNA** HBV
RG1450	**gantenerumab** early Alzheimer‘s
RG6100	**semorinemab** Alzheimer’s
RG6102	**brain shuttle gantenerumab** Alzheimer’s
RG6237	**latent myostatin + Evrysdi** SMA
RG6416	**bepranemab** Alzheimer's
RG7816	**alogabat (GABA Aa5 PAM)** ASD
RG7845	**fenebrutinib** RMS
RG7845	**fenebrutinib** PPMS
RG7906	**ralmitaront** schizophrenia
RG7935	**prasinezumab** Parkinson’s
RG6321	**Susvimo (PDS)** wAMD, 36-week refill
RG6147	**galegenimab (HtrA1)** geographic atrophy
RG6179	**NME** DME
RG6299†	**ASO factor B** geographic atrophy
RG7774	**NME** retinal disease

Status as of July 21, 2022

AI submissions for existing products

Projects in phase II and III

New Molecular Entity (NME)
Additional Indication (AI)
Oncology / Hematology
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Infectious Diseases

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Neuroscience
Ophthalmology
Other

2022		2023		2024		2025 and beyond	
RG6413+ RG6412	**Ronapreve**** SARS-CoV-2 hospitalized (EU) ✓	RG6264	**Phesgo OBI** HER2+ BC	RG1594	**Ocrevus SC** RMS & PPMS	RG7446	**Tecentriq** ctDNA+ high-risk MIBC	RG3502	**Kadcyla + Tecentriq** 2L+ HER-2+ PD-L1+ mBC
RG1569	**Actemra** COVID-19 pneumonia[1] ✓	RG6396	**Gavreto** Tumor agnostic	RG3648	**Xolair** food allergy	RG7601	**Venclexta + azacitidine** 1L MDS	RG3502	**Kadcyla + Tecentriq** HER-2+ eBC high-risk
RG7446	**Tecentriq ± chemo** 1L mUC	RG7446	**Tecentriq SC** 2L NSCLC	RG6152	**Xofluza** direct transmission	RG7159	**Gazyva** lupus nephritis	RG7446	**Tecentriq + paclitaxel** TNBC adj
RG7596	**Polivy** 1L DLBCL (US)	RG7446	**Tecentriq + cabozantinib** 2L NSCLC	RG6152	**Xofluza** influenza, pediatric (0-1 year)	RG6168	**Enspryng** myasthenia gravis	RG7446	**Tecentriq** High risk NMIBC
		RG7446	**Tecentriq + cabozantinib** RCC adv					RG7446	**Tecentriq+ lurbinectedin** 1l maintenance SCLC
		RG7446	**Tecentriq + Avastin** HCC adj					RG7159	**Gazyva** membranous nephropathy
		RG7446	**Tecentriq[2]** NSCLC neoadj					RG7159	**Gazyva** systemic lupus erythematosus
		RG7446	**Tecentriq** SCCHN adj					RG1594	**Ocrevus higher dose** RMS & PPMS
		RG7601	**Venclexta** r/r MM t(11:14)						
		RG7446	**Tecentriq + capecitabine or carbo/gem** TNBC						
		RG7853	**Alecensa** ALK+ NSCLC adj						

Status as of July 21, 2022

✓ Indicates submission to health authorities has occurred
Unless stated otherwise submissions are planned to occur in US and EU
[1]Approved in EU, filed in US
[2]filing timeline based on data from interim analysis

PDS=Port Delivery System with ranibizumab
OBI=On-Body Delivery System
**Ronapreve (casirivimab+imdevimab also known as REGEN-COV in the US) developed in collaboration with Regeneron Pharmaceuticals

Major pending approvals 2022

US	
RG6152	**Xofluza** influenza pediatric Filed March 2020
RG7828	**Lunsumio (mosunetuzumab)** 3L+ FL Filed Dec 2021
RG1569	**Actemra** COVID-19 pneumonia Filed Jan 2022

EU	
RG6321	**Susvimo (PDS)** wAMD Filed April 2021
RG7716	**Vabysmo (faricimab)** DME Filed May 2021
RG7716	**Vabysmo (faricimab)** wAMD Filed May 2021
RG6013	**Hemlibra** mild to moderate hemophilia A Filed Oct 2021
RG6396	**Gavreto** RET+ MTC, TC Filed Nov 2021
RG6152	**Xofluza** influenza pediatric Filed Nov 2021
RG7916	**Evrysdi** SMA presymptomatic pediatric <2mo Filed Nov 2021
RG6413+ RG6412	**Ronapreve**** SARS-CoV-2 hospitalized Filed Jan 2022
RG6026	**glofitamab** 3L+ DLBCL Filed April 2022

China	
RG6268	**Rozlytrek** ROS1+ NSCLC Filed Oct 2021
RG6268	**Rozlytrek** NTRK+ solid tumors Filed Nov 2021
RG7596	**Polivy** 1L DLBCL Filed Nov 2021
RG7596	**Polivy** r/r DLBCL Filed Dec 2021

Japan-Chugai	
RG7596	**Polivy** 1L DLBCL Filed Dec 2021
RG7159	**Gazyva** 1L CLL Filed March 2022

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PDS=Port Delivery System with ranibizumab
**Ronapreve (casirivimab+imdevimab also known as REGEN-COV in the US) developed in collaboration with Regeneron Pharmaceuticals

Status as of July 21, 2022

Major granted approvals 2022

US		EU		China		Japan-Chugai	
RG7716	**Vabysmo (faricimab)** DME Jan 2022	**RG7596**	**Polivy** 1L DLBCL May 2022	**RG7446**	**Tecentriq** NSCLC adj March 2022	**RG1569**	**Actemra** COVID-19 pneumonia Jan 2022
RG7716	**Vabysmo (faricimab)** wAMD Jan 2022	**RG7446**	**Tecentriq** NSCLC adj June 2022	**RG1569**	**Actemra** RA SC April 2022	**RG7716**	**Vabysmo (faricimab)** DME March 2022
RG1569	**Actemra** GCA IV Feb 2022	**RG7828**	**Lunsumio (mosunetuzumab)** 3L+ FL June 2022			**RG7716**	**Vabysmo (faricimab)** wAMD March 2022
RG7916	**Evrysdi** SMA presymptomatic pediatric <2mo May 2022					**RG1273**	**Perjeta + Herceptin** HER-2+ CRC March 2022
						RG7446	**Tecentriq** NSCLC adj May 2022
						RG6013	**Hemlibra** acquired Hemophilia A June 2022
						RG105	**Rituxan** NMOSD June 2022

- New Molecular Entity (NME)
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Roche Pharma research and early development (pRED)

Genentech research and early development (gRED)

Spark

Hemlibra

Factor VIII mimetic for treatment of hemophilia A

Indication	Hemophilia A patients without inhibitors to factor VIII	Hemophilia A patients with and without inhibitors to Factor VIII, dosing every 4 weeks
Phase/study	**Phase III HAVEN 3**	**Phase III HAVEN 4**
# of patients	N=135	N=46
Design	Patients on FVIII episodic treatment prior to study entry: ▪ **ARM A:** Hemlibra prophylaxis qw ▪ **ARM B:** Hemlibra prophylaxis q2w ▪ **ARM C:** Episodic FVIII treatment; switch to Hemlibra prophylaxis possible after 24 weeks Patients on FVIII prophylaxis prior to study entry: ▪ **ARM D:** Hemlibra prophylaxis qw	Multicenter, open-label, non-randomized study to assess the efficacy, safety, pharmacokinetics, and pharmacodynamics of Hemlibra administered every 4 weeks. ▪ **Part 1:** Pharmacokinetic run-in part (N=6) ▪ **Part 2:** Expansion part (N=40)
Primary endpoint	▪ Number of bleeds over 24 weeks	▪ Number of bleeds over 24 weeks
Status	▪ FPI Q3 2016, recruitment completed Q2 2017 ▪ Study met primary and key secondary endpoints Q4 2017 ▪ FDA granted Breakthrough Therapy Designation April 2018 ▪ Data presented at WFH 2018 ▪ Filed in US (priority review) and EU in Q2 2018 ▪ Data published in *NEJM* 2018; 379: 811-822	▪ FPI Q1 2017, recruitment completed Q2 2017 ▪ Pharmacokinetic run-in data at ASH 2017 ▪ Positive interim analysis outcome reported Q4 2017 ▪ Data presented at WFH 2018 ▪ Interim data filed in US and EU in Q2 2018 ▪ Data published in Lancet Haematology 2019 Jun;6(6):e295-e305
	•Approved in US Q4 2018 and EU Q1 2019	
CT Identifier	NCT02847637	NCT03020160

In collaboration with Chugai

ASH=American Society of Hematology; WFH=World Federation of Hemophilia; NEJM=New England Journal of Medicine

Hemlibra

Factor VIII mimetic for treatment of hemophilia A

Indication	Hemophilia A patients with and without inhibitors to Factor VIII	Hemophilia A mild to moderate patients without inhibitors to Factor VIII
Phase/study	**Phase III HAVEN 5**	**Phase III HAVEN 6**
# of patients	N=85	N=70
Design	Patients with Hemophilia regardless of FVIII inhibitor status on prophylactic or episodic treatment prior to study entry: • **Arm A**: Hemlibra prophylaxis qw • **Arm B**: Hemlibra prophylaxis q4w • **Arm C**: No prophylaxis (control arm)	Multicenter, open-label study to evaluate the safety, efficacy, pharmacokinetics, and pharmacodynamics of Hemlibra in patients with mild or moderate Hemophilia A without FVIII inhibitors ▪ Hemlibra qw (1.5mg/kg), q2w (3.0mg/kg) or q4w (6.0mg/kg) (patients preference)
Primary endpoint	▪ Number of bleeds over 24 weeks	▪ Safety and efficacy
Status	▪ FPI Q2 2018 ▪ Recruitment completed Q1 2019 ▪ Filed in China Q2 2020 ▪ Approved in China Q2 2021	▪ FPI Q1 2020 ▪ Recruitment completed Q1 2021 ▪ Interim data presented at ASH 2021 and primary data presented at ISTH 2022 ▪ Filed in EU Q4 2021
CT Identifier	NCT03315455	NCT04158648

In collaboration with Chugai
ASH=American Society of Hematology; ISTH=International Society on Thrombosis and Haemostasis

Alecensa

New CNS-active inhibitor of anaplastic lymphoma kinase

Indication	Treatment-naïve ALK+ advanced NSCLC	Adjuvant ALK+ NSCLC
Phase/study	**Phase III ALEX**	**Phase III ALINA**
# of patients	N=286	N=255
Design	▪ **ARM A:** Alecensa 600mg BID ▪ **ARM B:** Crizotinib 250mg BID	▪ **ARM A:** Alecensa 600mg BID ▪ **ARM B:** Platinum-based chemotherapy
Primary endpoint	▪ Progression-free survival	▪ Disease-free survival
Status	▪ Recruitment completed Q3 2015 ▪ Primary endpoint met Q1 2017 ▪ Data presented at ASCO 2017, 2018, ESMO 2017, 2018 ▪ Data published in *NEJM* 2017; 377:829-838 ▪ CNS data presented at ESMO 2017 ▪ Final PFS and updated OS presented at ESMO 2019 ▪ Approved in US Q4 2017 (priority review) and in EU Q4 2017	▪ FPI Q3 2018 ▪ Recruitment completed Q4 2021
CT Identifier	NCT02075840	NCT03456076

In collaboration with Chugai

ALK=anaplastic lymphoma kinase; CNS= Central nervous system; NSCLC=non-small cell lung cancer; OS=Overall survival, PFS=Progression-free survival; ASCO=American Society of Clinical Oncology; *NEJM*=New England Journal of Medicine; ESMO=European Society for Medical Oncology

Kadcyla

First ADC for HER2-positive breast cancer

Indication	HER2-positive early breast cancer (BC) high-risk patients	2L+ HER-2 positive PD-L1 positive metastatic breast cancer (mBC)	HER2-positive early breast cancer (BC) high-risk patients
Phase/study	**Phase III KATHERINE**	**Phase III KATE 3**	**Phase III ASTEFANIA**
# of patients	N=1,484	N=320	N=1700
Design	▪ **ARM A:** Kadcyla 3.6mg/kg q3w ▪ **ARM B:** Herceptin	▪ **ARM A:** Kadcyla plus Tecentriq ▪ **ARM B:** Herceptin plus placebo	▪ **ARM A:** Kadcyla plus Tecentriq ▪ **ARM B:** Kadcyla plus placebo
Primary endpoint	▪ Invasive disease-free survival	▪ Progression-free survival and overall survival	▪ Invasive disease-free survival
Status	▪ Recruitment completed Q4 2015 • Stopped at pre-planned interim data analysis for efficacy Q4 2018 • Data presented at SABCS 2018 • BTD granted by FDA in Q1 2019 • US filling completed under RTOR Q1 2019 and filed in EU Q1 2019 • Approved in US Q2 2019 and in EU Q4 2019 • Data published in *NEJM* 2019; 380:617-628	▪ FPI Q1 2021	▪ FPI Q2 2021
CT Identifier	NCT01772472	NCT04740918	NCT04873362

In collaboration with ImmunoGen, Inc.

ADC=antibody drug conjugate; BTD=Breakthrough therapy designation; HER2=Human Epidermal growth factor Receptor 2; SABCS=San Antonio Breast Cancer Symposium; RTOR=Real time oncology review; NEJM=New England Journal of Medicine

Perjeta

First-in-class HER2 dimerization inhibitor

Indication	**Adjuvant HER2-positive breast cancer (BC)**
Phase/study	**Phase III** **APHINITY**
# of patients	N=4,803
Design	▪ **ARM A:** Perjeta (840mg loading dose, 420mg q3w) plus Herceptin for 52 weeks plus chemotherapy (6-8 cycles) ▪ **ARM B:** Placebo plus Herceptin (52 weeks) plus chemotherapy (6-8 cycles)
Primary endpoint	▪ Invasive disease-free survival (iDFS)
Status	▪ Primary endpoint met Q1 2017 ▪ Data presented at ASCO 2017 and published in *NEJM* 2017; 377:122-131 ▪ Filed in US and EU Q3 2017 ▪ Approved in US Q4 2017 (priority review) and EU Q2 2018 ▪ 6-year iDFS data presented at SABCS 2019 ▪ 8-year iDFS data presented at ESMO virtual 2022
CT Identifier	NCT01358877

HER2=Human Epidermal growth factor Receptor 2, IDFS=Invasive disease-free survival; ASCO=American Society of Clinical Oncology; *NEJM*=New England Journal of Medcine; SABCS=San Antonio Breast Cancer Symposium

Phesgo

FDC of Perjeta and Herceptin for subcutaneous administration

Indication	HER2-positive early breast cancer (BC)		HER2-positive breast cancer (BC)
Phase/study	**Phase III FeDeriCa**	**Phase II PHranceSCa**	**Phase I[1]**
# of patients	N=500	N=160	N=144
Design	FDC of Perjeta and Herceptin for SC administration (Phesgo) in combination with chemotherapy in neoadjuvant/adjuvant setting ▪ **ARM A:** Perjeta IV plus Herceptin IV plus chemotherapy ▪ **ARM B:** Phesgo plus chemotherapy	▪ **ARM A:** Perjeta and Herceptin IV followed by Phesgo ▪ **ARM B:** Phesgo followed by IV	▪ **Arm A:** Phesgo administered using a handheld syringe with hypodermic needle (SC) ▪ **ARM B:** Phesgo administered using the on-body delivery system (OBI)
Primary endpoint	▪ Trough Serum Concentration (Ctrough) of Perjeta during cycle 7	▪ Percentage of patients who preferred Perjeta and Herceptin FDC SC	▪ AUC0-62*, Cmax**
Status	▪ Primary endpoint met Q3 2019 ▪ Data presented at SABCS 2019 ▪ Data published in Lancet Oncology 2021 Jan;22(1):85-97	▪ FPI Q4 2018 ▪ Final analysis completed, 85% patients preferred FDC SC ▪ Data presented at ESMO 2020 ▪ Data published in *Eur J Cancer* 2021 Jul;152:223-232	▪ FPI Q2 2022
	▪ Filed in US Dec 2019 & in EU Jan 2020; Approved in US Q2 2020 and EU Q4 2020		
CT Identifier	NCT03493854	NCT03674112	NCT05275010

[1]In collaboration with West Pharmaceuticals

*AUC0-62=comparability of area under the time-concentration curve from the start of dosing to 63 days; **Cmax=maximum serum concentration for pertuzumab and trastuzumab within Phesgo; FDC=Fixed-dose combination; Phesgo=FDC of Perjeta and Herceptin for SC administration;HER2=Human Epidermal growth factor Receptor 2, IV=intravenous; SC=Subcutaneous; ASCO=American Society of Clinical Oncology; NEJM=New England Journal of Medcine; SABCS=San Antonio Breast Cancer Symposium; Eur J Cancer=European Journal of Cancer; ESMO=European Society for Medical Oncology

Tecentriq

Anti-PD-L1 cancer immunotherapy – lung cancer

Indication	Adjuvant NSCLC	Neoadjuvant NSCLC
Phase/study	**Phase III** **IMpower010**	**Phase III** **IMpower030**
# of patients	N=1,280	N=450
Design	Following adjuvant cisplatin-based chemotherapy ▪ **ARM A:** Tecentriq ▪ **ARM B:** Best supportive care	▪ **ARM A:** Tecentriq plus platinum-based chemotherapy ▪ **ARM B:** Platinum-based chemotherapy
Primary endpoint	▪ Disease-free survival	▪ Event-free survival
Status	▪ Trial amended from PD-L1+ selected patients to all-comers ▪ FPI for all-comer population Q4 2016 ▪ Recruitment completed Q3 2018 ▪ Study met primary endpoint Q1 2021 ▪ Data presented at ASCO, WCLC and ESMO 2021 ▪ Filed in US (priority review) and EU Q2 2021 ▪ Approved in US Q4 2021 and EU Q2 2022	▪ FPI Q2 2018 ▪ Recruitment completed Q3 2021
CT Identifier	NCT02486718	NCT03456063

NSCLC=non-small cell lung cancer; PD-L1=Programmed cell death-ligand 1; ASCO=American Society of Clinical Oncology; ESMO==European Society for Medical Oncology; WCLC=World Conference on Lung Cancer

Tecentriq

Anti-PD-L1 cancer immunotherapy – lung cancer

Indication	1L maintenance extensive-stage SCLC	2L NSCLC previously treated with an immune checkpoint inhibitor
Phase/study	**Phase III** **IMforte**[1]	**Phase III** **CONTACT-01**
# of patients	N=450	N=366
Design	▪ **ARM A:** Platinum-etoposide + Tecentriq followed by maintenance Tecentriq plus lurbinectedin ▪ **ARM B:** Platinum-etoposide + Tecentriq followed by maintenance Tecentriq	▪ **ARM A:** Tecentriq plus cabozantinib ▪ **ARM B:** Docetaxel
Primary endpoint	▪ Progression-free survival and overall survival	▪ Overall survival
Status	▪ FPI Q4 2021	▪ FPI Q3 2020 ▪ Recruitment completed Q4 2021
CT Identifier	NCT05091567	NCT04471428

[1]In collaboration with Jazz Pharma

NSCLC=non-small cell lung cancer; PD-L1=Programmed cell death-ligand 1; SCLC=small cell lung cancer;

Tecentriq

Anti-PD-L1 cancer immunotherapy – lung cancer

Indication	1L NSCLC	Stage IV NSCLC
Phase/study	**Phase II/III** **B-FAST**	**Phase Ib/III** **IMscin001**[1]
# of patients	Modular design	N=371
Design	▪ **Cohort A:** ALK+ (Alecensa) ▪ **Cohort B:** RET+ (Alecensa) ▪ **Cohort C:** bTMB-high (Tecentriq) ▪ **Cohort D:** ROS1+ (Rozlytrek) ▪ **Cohort E:** BRAF+ (Zelboraf plus Cotellic plus Tecentriq) ▪ **Cohort F:** EGFR Exon 20+ (Tecentriq, Avastin, carboplatin, pemetrexed) ▪ **Cohort G:** GDC-6036 or Docetaxel	**Phase Ib** ▪ Dose finding, Tecentriq SC followed by Tecentriq IV **Phase III** ▪ 2L NSCLC non inferiority of Tecentriq SC vs Tecentriq IV
Primary endpoint	▪ Cohort A/B/D: Objective response rate ▪ Cohort C/D: Progression-free survival ▪ Cohort E: Time in response ▪ Cohort F: Investigator-assessed objective response rate	▪ Observed concentration of Tecentriq in serum at cycle 1
Status	▪ FPI Q3 2017 ▪ Recruitment completed for cohort A Q3 2018 and cohort C Q3 2019 ▪ Cohort A: primary endpoint met Q3 2019; approved in US Q1 2021 ▪ Cohort C: did not show statistical significance for primary endpoint, data presented at ESMO 2021 ▪ Cohort F: FPI Q2 2021	▪ FPI Q4 2018 ▪ FPI in phase III part Q4 2020 ▪ Recruitment completed Q1 2022
CT Identifier	NCT03178552	NCT03735121

[1]SC with Halozyme's rHuPH20/ Halozyme's human hyaluronidase

ALK=Anaplastic lymphoma kinase; BRAF=V-raf murine sarcoma viral oncogene homolog B; bTMB=Blood-based tumor mutational burden; EGFR=Epidermal growth factor receptor; IV=intravenous; NSCLC=non-small cell lung cancer; PD-L1=Programmed cell death-ligand 1; RET=Rearranged during transfection; ROS1=C-ros oncogene 1; SC=Subcutaneous, IV=Intravenous; ESMO=European Society for Medical Oncology

Tecentriq

Anti-PD-L1 cancer immunotherapy – SCCHN and melanoma

Indication	Adjuvant squamous cell carcinoma of the head and neck (SCCHN)
Phase/study	**Phase III** **IMvoke010**
# of patients	N=406
Design	▪ **ARM A:** Tecentriq 1200mg q3w ▪ **ARM B:** Placebo
Primary endpoint	▪ Event-free survival and overall survival
Status	▪ FPI Q1 2018 ▪ Recruitment completed Q1 2020
CT Identifier	NCT03452137

[1]In collaboration with Exelixis; [2]Zelboraf in collaboration with Plexxikon, a member of Daiichi Sankyo Group; [3]Project Orbis=FDA framework for concurrent submission and review of oncology products among international partners
SCCHN=squamous cell carcinoma of the head and neck; PD-L1=Programmed cell death-ligand 1; AACR=American Association for Cancer Research

Tecentriq

Anti-PD-L1 cancer immunotherapy – urothelial carcinoma

Indication	1L metastatic urothelial carcinoma (UC)	High-risk non-muscle-invasive bladder cancer (MIBC)	ctDNA+, high-risk muscle-invasive bladder cancer (MIBC)
Phase/study	**Phase III IMvigor130**	**Phase III ALBAN**	**Phase III IMvigor011**
# of patients	N=1,200	N=516	N=495
Design	▪ **ARM A:** Tecentriq plus gemcitabine and carboplatin or cisplatin ▪ **ARM B:** Tecentriq monotherapy ▪ **ARM C:** Placebo plus gemcitabine and carboplatin or cisplatin	▪ **ARM A**: BCG induction and maintenance ▪ **ARM B:** Tecentriq plus BCG induction and maintenance	▪ **ARM A**: Tecentriq monotherapy ▪ **ARM B:** Placebo
Primary endpoint	▪ Progression-free survival, overall survival and safety	▪ Recurrence-free survival	▪ Recurrence-free survival
Status	▪ FPI Q3 2016 ▪ FPI for arm B (amended study) Q1 2017 ▪ Recruitment completed Q3 2018 ▪ Study met co-primary endpoint of PFS Q3 2019 ▪ Data presented at ESMO 2019 and AACR 2021 ▪ Data published in Lancet 2020 May 16;395(10236):1547-1557	▪ FPI Q4 2018	▪ FPI Q2 2021
CT Identifier	NCT02807636	NCT03799835	NCT04660344

BCG=Bacille Calmette-Guérin; PD-L1=Programmed cell death-ligand 1; PFS=Progression-free survival; AACR=American Association for Cancer Research; ESMO=European Society for Medical Oncology

Tecentriq

Anti-PD-L1 cancer immunotherapy – renal cell cancer

Indication	Adjuvant renal cell carcinoma (RCC)	Advanced renal cell carcinoma (RCC) after immune checkpoint inhibitor treatment
Phase/study	**Phase III** **IMmotion010**	**Phase III** **Contact-03**[1]
# of patients	N=778	N=500
Design	▪ **ARM A:** Tecentriq monotherapy ▪ **ARM B:** Placebo	▪ **ARM A:** Tecentriq plus cabozantinib ▪ **ARM B:** Cabozantinib
Primary endpoint	▪ Investigator-assessed disease-free survival	▪ Progression-free survival and overall survival
Status	▪ FPI Q1 2017 ▪ Recruitment completed Q1 2019 ▪ Study did not meet its primary endpoint of DFS Q2 2022	▪ FPI Q3 2020 ▪ Recruitment completed Q4 2021
CT Identifier	NCT03024996	NCT04338269

[1]In collaboration with Exelixis

PD-L1=Programmed cell death-ligand 1; DFS=Disease-free survival

Tecentriq

Anti-PD-L1 cancer immunotherapy – hepatocellular carcinoma

Indication	1L hepatocellular carcinoma (HCC)	Adjuvant hepatocellular carcinoma (HCC)
Phase/study	**Phase III** **IMbrave150**	**Phase III** **IMbrave050**
# of patients	N=501	N=668
Design	▪ **ARM A:** Tecentriq plus Avastin ▪ **ARM B:** Sorafenib	▪ **ARM A:** Tecentriq plus Avastin ▪ **ARM B:** Active surveillance
Primary endpoint	▪ Overall survival and progression free survival	▪ Recurrence-free survival
Status	▪ FPI Q1 2018 ▪ Recruitment completed Q1 2019 ▪ Data presented at ESMO Asia 2019 ▪ US filing completed under RTOR Q1 2020; filed in EU Q1 2020 ▪ Data published in *NEJM* 2020;382:1894-1905 ▪ Approved in US Q2 2020 and EU Q4 2020	▪ FPI Q4 2019 ▪ Recruitment completed Q4 2021
CT Identifier	NCT03434379	NCT04102098

PD-L1=Programmed cell death-ligand 1; ESMO=European Society for Medical Oncology; NEJM=New England Journal of Medicine; RTOR=Real time oncology review

Tecentriq

Anti-PD-L1 cancer immunotherapy – breast cancer

Indication	Previously untreated metastatic triple negative breast cancer (TNBC)	
Phase/study	**Phase III IMpassion130**	**Phase III IMpassion132**
# of patients	N=902	N=572
Design	▪ **ARM A:** Tecentriq plus nab-paclitaxel ▪ **ARM B:** Placebo plus nab-paclitaxel	▪ **ARM A:** Tecentriq plus capecitabine or carbo/gem ▪ **ARM B:** Placebo plus capecitabine or carbo/gem
Primary endpoint	▪ Progression-free survival and overall survival (co-primary endpoint)	▪ Overall survival
Status	▪ Study met co-primary endpoint of PFS in both PD-L1+ and ITT populations Q3 2018 ▪ Primary PFS and interim OS data presented at ESMO 2018 and ASCO 2019 ▪ Data published in *NEJM* 2018; 379:2108-2121 ▪ US accelerated approval Q1 2019 – US indication voluntarily withdrawn Q3 2021 ▪ Approved in EU Q3 2019 ▪ Final OS presented at ESMO Asia 2020	▪ FPI Q1 2018
CT Identifier	NCT02425891	NCT03371017

Carbo/gem=gemcitabine and carboplatin; ITT=Intention to treat; PD-L1=Programmed cell death-ligand 1; PFS=Progression-free survival; OS=Overall survival; ESMO=European Society for Medical Oncology; ASCO=American Society of Clinical Oncology; NEJM=New England Journal of Medicine

Tecentriq

Anti-PD-L1 cancer immunotherapy – breast cancer

Indication	Neoadjuvant triple negative breast cancer (TNBC)	Adjuvant triple negative breast cancer (TNBC)
Phase/study	**Phase III** **IMpassion031**	**Phase III** **IMpassion030**
# of patients	N=333	N=2,300
Design	▪ **ARM A:** Tecentriq plus nab-paclitaxel ▪ **ARM B:** Placebo plus nab-paclitaxel	▪ **ARM A:** Tecentriq plus paclitaxel followed by AC followed by Tecentriq plus AC, followed by Tecentriq maintenance ▪ **ARM B:** Placebo plus paclitaxel followed by AC followed by placebo
Primary endpoint	▪ Percentage of participants with pathologic complete response	▪ Invasive disease-free survival
Status	▪ FPI Q3 2017 ▪ Recruitment completed Q2 2018 ▪ Study met primary endpoint Q2 2020 ▪ Data presented at ESMO 2020 ▪ Data published in Lancet 2020;396 (10257):1090-1100 ▪ Filed in EU Q4 2020 - application withdrawn Q3 2021	▪ FPI Q3 2018
CT Identifier	NCT03197935	NCT03498716

PD-L1=Programmed cell death-ligand 1; ESMO=European Society for Medical Oncology

Venclexta

Novel small molecule Bcl-2 selective inhibitor – chronic lymphocytic leukemia

Indication	Untreated chronic lymphocytic leukemia (CLL) patients with coexisting medical conditions	Relapsed or refractory chronic lymphocytic leukemia (CLL)	Untreated fit chronic lymphocytic leukemia (CLL) patients
Phase/study	**Phase III CLL14**	**Phase III MURANO**	**Phase III CristaLLo**
# of patients	N=445	N=389	N=165
Design	▪ **ARM A:** Venclexta plus Gazyva ▪ **ARM B:** Chlorambucil plus Gazyva	▪ **ARM A:** Venclexta plus Rituxan ▪ **ARM B:** Rituxan plus bendamustine	▪ **ARM A:** Venclexta plus Gazyva ▪ **ARM B:** Fludarabine plus cyclophosphamide plus Rituxan or bendamustine plus Rituxan
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival	▪ MRD negativity rate in peripheral blood at 15 months
Status	▪ Study met primary endpoint at pre-specified interim analysis Q4 2018 ▪ BTD granted by FDA Q1 2019 ▪ US filing completed under RTOR Q1 2019 ▪ Filed in EU Q2 2019 ▪ Data presented at ASCO 2019, ASH 2019, ASH 2020 and EHA 2021 and EHA 2022 ▪ Data published in *NEJM* 2019; 380:2225-2236 ▪ Approved US Q2 2019 and EU Q1 2020	▪ Study met primary endpoint at interim analysis ▪ Data presented at ASH 2017 ▪ Filed in US Q4 2017 and EU Q1 2018 ▪ Data published in *NEJM* 2018; 378:1107–20 ▪ Updated data presented at ASCO 2018, ASH 2019 and ASH 2020 ▪ Approved in US Q2 2018 (priority review) ▪ EU approval Q4 2018	▪ FPI Q2 2020
CT Identifier	NCT02242942	NCT02005471	NCT04285567

Joint project with AbbVie, in collaboration with The Walter and Eliza Hall Institute

Bcl-2=B-cell lymphoma 2; BTD=Breakthrough therapy designation; CLL=chronic lymphocytic leukemia; MRD=Minimal Residual Disease; ASH=American Society of Hematology; ASCO=American Society of Clinical Oncology; EHA=European Hematology Association; RTOR=Real time oncology review; NEJM=New England Journal of Medicine

Venclexta

Novel small molecule Bcl-2 selective inhibitor – multiple myeloma

Indication	Relapsed or refractory multiple myeloma (MM)		
Phase/study	**Phase I**	**Phase Ib/II**	**Phase III CANOVA**
# of patients	N=117	N=120	N=244
Design	▪ **Dose escalation cohort:** Venclexta dose escalation ▪ **Safety expansion cohort (t11;14):** Venclexta expansion ▪ **Combination:** Venclexta plus dexamethasone	▪ Venclexta plus carfilzomib plus dexamethasone in t(11;14) positive r/r MM	▪ Venclexta plus dexamethazone vs pomalidomide plus dexamethasone in t(11;14) positive r/r MM
Primary endpoint	▪ Safety and maximum tolerated dose	▪ Safety, objective response rate, Pharmacokinetics, Pharmacodynamics	▪ Progression-free survival
Status	▪ FPI Q4 2012 ▪ Data presented at ASCO 2015 ▪ Updated data presented at ASCO 2016 and ASH 2016 ▪ Data published in Blood 2017; 130(22):2401-2409 and Am *J Hematol* 2021 Apr 1;96(4):418-427	▪ FPI Q1 2017 ▪ Data published *Blood Adv* 2021 Oct 12;5(19):3748-3759	▪ FPI Q4 2018
CT Identifier	NCT01794520	NCT02899052	NCT03539744

Joint project with AbbVie, in collaboration with The Walter and Eliza Hall Institute;
Bcl-2=B-cell lymphoma 2; MM=multiple myeloma; r/r=Relapsed or refractory ; ASCO=American Society of Clinical Oncology; ASH=American Society of Hematology

Venclexta

Novel small molecule Bcl-2 selective inhibitor – myelodysplastic syndromes

Indication	Relapsed or refractory myelodysplastic syndromes (MDS)	Treatment-naive myelodysplastic syndromes (MDS)	Newly diagnosed higher-risk myelodysplatic syndrome (MDS)
Phase/study	**Phase Ib**	**Phase Ib**	**Phase III VERONA**
# of patients	N=70	N=129	N=500
Design	Cohort 1: ▪ **ARM A:** Venclexta 400 mg ▪ **ARM B:** Venclexta 800 mg Cohort 2: ▪ **ARM A:** Venclexta plus azacitidine Study expansion: ▪ Venclexta or Venclexta plus azacitidine	▪ **Dose escalation cohort:** Venclexta plus azacitidine dose escalation ▪ **Safety expansion cohort**	▪ **ARM A:** Venclexta plus azacitidine ▪ **ARM B:** Placebo plus azacitidine
Primary endpoint	▪ Safety, efficacy, Pharmacokinetics and Pharmacodynamics	▪ Safety, Pharmacokinetics, RPTD	▪ Complete remission rate and overall survival
Status	▪ FPI Q1 2017 ▪ Recruitment completed Q1 2022	▪ FPI Q1 2017 ▪ Data presented at ASH 2019, ASH 2020 and ASCO 201 ▪ BTD granted by FDA July 2021 ▪ Recruitment completed Q1 2022	▪ FPI Q4 2020
CT Identifier	NCT02966782	NCT02942290	NCT04401748

Joint project with AbbVie, in collaboration with The Walter and Eliza Hall Institute

Bcl-2=B-cell lymphoma 2; BTD=Breakthrough therapy designation; RPTD=Recommended phase II dose ; ASH=American Society of Hematology

Polivy (polatuzumab vedotin)

ADC targeting CD79b to treat B cell malignancies

Indication	1L DLBCL
Phase/study	**Phase III** **POLARIX**
# of patients	N=879
Design	▪ **ARM A:** Polivy plus R-CHP ▪ **ARM B:** R-CHOP
Primary endpoint	▪ Progression-free survival
Status	▪ FPI Q4 2017 ▪ Recruitment completed Q2 2019 ▪ Study met primary endpoint Q3 2021 ▪ Data presented at ASH 2021 ▪ Filed in EU, Japan and China Q4 2021 ▪ Published in *NEJM* 2022 Jan 27;386(4):351-363 ▪ Approved in EU Q2 2022
CT Identifier	NCT03274492

In collaboration with Seagen Inc.

DLBCL=diffuse large B cell lymphoma; R-CHP=Rituxan, cyclophosphamide, hydroxydoxorubicin, prednisone; R-CHOP=Rituxan, cyclophosphamide, doxorubicin, vincristine, and prednisone; ASH=American Society of Hematology, NEJM=New England Journal of Medicine

Rozlytrek (entrectinib)

CNS-active and selective inhibitor of NTRK/ROS1

Indication	Locally advanced or metastatic tumors with ROS1 gene rearrangement	Locally advanced or metastatic tumors with NTRK1/2/3 gene rearrangement	Pediatric tumors with NTRK 1/2/3, ROS-1 or ALK rearrangement
Phase/study	**Phase II STARTRK2**	**Phase II STARTRK2**	**Phase I/Ib STARTRK - NG**
# of patients	N~300 total	N~300 total	N~80
Design	Single arm with Baskets based on tumor type and genomic alteration status	Single arm with Baskets based on tumor type and genomic alteration status	Single arm with Baskets based on tumor type and genomic alteration status
Primary endpoint	▪ Objective response rate	▪ Objective response rate	▪ Maximum tolerated dose and RPTD
Status	▪ FPI Q1 2016 ▪ Data presented at WCLC 2018	▪ FPI Q1 2016 ▪ Data presented at ESMO 2018	▪ FPI Q2 2016 ▪ Initial data presented at ASCO 2019
	▪ Breakthrough Therapy Designation granted by FDA (Q2 2017), PRIME designation granted by EMA (Q1 2018) and Sakigake Designation granted by MHLW (Q4 2017) for NTRK fusion-positive, locally advanced or metastatic solid tumors ▪ Filed in US Q4 2018 and EU Q1 2019 ▪ Approved in US Q3 2019 and EU Q3 2020 ▪ Published in Lancet Oncol. 2020 Feb;21(2):261-271 and 271-282		
CT Identifier	NCT02568267	NCT02568267	NCT02650401

ALK=Anaplastic lymphoma kinase; CNS=Central Nervous System; MHLW=Ministry of Health, Labor and Welfare; NTRK=neurotrophic receptor tyrosine kinase; PRIME= priority medicines; RPTD=Recommended phase II dose; ROS1=C-ros oncogene 1; WCLC=World Conference on Lung Cancer; ESMO=European Society for Medical Oncology; ASCO=American Society of Clinical Oncology

Gavreto (pralsetinib, RG6396)

Highly selective RET inhibitor

Indication	RET+ NSCLC, thyroid cancer and other advanced solid tumors	1L RET fusion-positive, metastatic NSCLC
Phase/study	**Phase I/II ARROW**	**Phase III AcceleRET Lung**
# of patients	N=647	N=250
Design	▪ **Part 1:** Gavreto 30-600mg dose escalation ▪ **Part 2:** Gavreto 400mg dose expansion	▪ **Arm A:** Gavreto 400mg ▪ **Arm B:** Platinum-based chemotherapy +/- pembrolizumab
Primary endpoint	▪ Safety and efficacy	▪ Progression-free survival
Status	▪ Data presented at ASCO (NSCLC) and ESMO (MTC) 2020 ▪ Filed in US and EU for RET fusion-positive NSCLC and US for RET-mutant MTC and RET fusion-positive thyroid cancer ▪ Approved in US Q3 2020 in RET fusion-positive NSCLC, in Q4 2020 in RET-mutant MTC and RET fusion-positive thyroid cancer ▪ Updated data presented at ASCO 2021 and 2022 ▪ Data published in Lancet Oncol 2021 Jul;22(7):959-969 and Lancet Diabetes & Endocrinology Aug 2021;9(8):491-501 ▪ Approved in EU for RET fusion-positive NSCLC Q4 2021	▪ Study initiated in Q1 2020
CT Identifier	NCT03037385	NCT04222972

In collaboration with Blueprint Medicines

NSCLC=non-small cell lung cancer; MTC=medullary thyroid cancer; RET=Rearranged during transfection; ASCO=American Society of Clinical Oncology; ESMO=European Society for Medical Oncology

Lunsumio (mosunetuzumab, CD20 x CD3, RG7828)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	3L+ FL, 3L+ DLBCL & other relapsed or refractory NHL	1L DLBCL	Relapsed or refractory DLBCL
Phase/study	**Phase I/II**	**Phase Ib/II**	**Phase Ib**
# of patients	N=746	N=160	N=262
Design	▪ Dose escalation study of Lunsumio as single agent and in combination with Tecentriq ▪ Expansion cohorts for r/r FL, r/r DLBCL and SC in r/r NHL	▪ Lunsumio plus CHOP ▪ Lunsumio plus CHP plus Polivy ▪ Lunsumio plus CHP-Polivy vs Rituximab plus CHP-Polivy	▪ Lunsumio plus Polivy, randomised cohorts ▪ **ARM A:** Lunsumio SC plus Polivy ▪ **ARM B:** Rituximab plus Polivy
Primary endpoint	▪ Safety, tolerability, dose/schedule, PK and response rates	▪ Safety/tolerability and response	▪ Safety/tolerability and response
Status	▪ Data in r/r NHL presented at ASH 2018 and 2019, and in r/r FL at ASH 2020 and ASH 2021 ▪ BTD granted by FDA Q2 2020 ▪ SC cohort FPI Q2 2021 ▪ Filed in EU and rolling submission submitted in US Q4 2021 ▪ Approved in EU Q2 2022 ▪ Filed in US (priority review) Q2 2022	▪ FPI Q1 2019 ▪ Data for Lunsumio plus CHOP presented at ASH 2020	▪ FPI Q3 2018 ▪ Initial data presented at ASCO and ASH 2021
CT Identifier	NCT02500407	NCT03677141	NCT03671018

FL=follicular lymphoma; DLBCL=diffuse large B cell lymphoma; r/r=relapsed/refractory; NHL=non-Hodgkin's lymphoma; R=Rituximab; SC=subcutaneous; CHOP=cyclophosphamide, doxorubicin, vincristine, and prednisone; CHP=cyclophosphamide, doxorubicin, and prednisone); PK=Pharmacokinetics; BTD=Breakthrough Therapy Designation; ASH=American Society of Hematology; ASCO=American Society of Clinical Oncology

Lunsumio (mosunetuzumab, CD20 x CD3, RG7828)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	1L DLBCL & 2L DLBCL following 1L induction	Relapsed or refractory 2L+ FL
Phase/study	**Phase I**	**Phase Ib**
# of patients	N=92 + 80 (cohort C)	N=27
Design	▪ **Cohort A:** Lunsumio monotherapy (after a response to prior systemic chemotherapy) ▪ **Cohort B:** Lunsumio monotherapy (1L treatment in elderly/frail) ▪ **Cohort C:** Lunsumio SC plus Polivy in 1L elderly/unfit	▪ Lunsumio plus lenalidomide safety run-in for phase III ▪ Lunsumio SC plus lenalidomide
Primary endpoint	▪ Safety/tolerability and response	▪ Safety/tolerability and response
Status	▪ FPI Q2 2019 – Cohort B ▪ FPI Q3 2019 – Cohort A ▪ Initial data presented at ASH 2020 (cohort B) ▪ Cohort C: FPI Q1 2021	▪ FPI Q3 2020 ▪ Initial data presented at ASH 2021
CT Identifier	NCT03677154	NCT04246086

FL=follicular lymphoma; DLBCL=diffuse large B cell lymphoma; r/r=relapsed/refractory; SC=subcutaneous; ASH=American Society of Hematology

Lunsumio (mosunetuzumab, CD20 x CD3, RG7828)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	2L+ FL	Relapsed or refractory FL	Relapsed or refractory CLL
Phase/study	**Phase III CELESTIMO**	**Phase Ib/II**	**Phase Ib/II**
# of patients	N=400	N=118	N=56
Design	▪ **ARM A**: Lunsumio plus lenalidomide ▪ **ARM B:** Rituxan plus lenalidomide	▪ **ARM A**: Lunsumio plus tiragolumab ▪ **ARM B:** Lunsumio plus tiragolumab plus Tecentriq ▪ Dose escalation phase ▪ Dose expansion phase	▪ Lunsumio monotherapy (3L+ CLL)
Primary endpoint	▪ Progression-free survival	▪ Phase Ib: Dose-limiting toxicity ▪ Phase II: Best complete response	▪ Safety, dose-limiting toxicity and RPTD
Status	▪ FPI Q4 2021	▪ FPI Phase Ib Q2 2022	▪ FPI Q1 2022
CT Identifier	NCT04712097	NCT05315713	

FL=follicular lymphoma; r/r=relapsed/refractory; RPTD=Recommended Phase II Dose; CLL=Chronic lymphocytic leukemia

Lunsumio (mosunetuzumab, CD20 x CD3, RG7828)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	2L+ SCT ineligible DLBCL
Phase/study	**Phase III** **SUNMO**
# of patients	N=222
Design	▪ **ARM A:** Lunsumio plus Polivy ▪ **ARM B:** R + GemOx
Primary endpoint	▪ Progression-free survival
Status	▪ FPI Q2 2022
CT Identifier	NCT05171647

DLBCL=diffuse large B cell lymphoma; SCT=stem cell transplant; R=Rituxan/MabThera; GemOx=Gemcitabin und Oxaliplatin

Ocrevus (ocrelizumab, RG1594)

Humanized monoclonal antibody selectively targeting CD20+ B cells

Indication	Relapsing multiple sclerosis (RMS)		Primary progressive multiple sclerosis (PPMS)
Phase/study	**Phase III OPERA I**	**Phase III OPERA II**	**Phase III ORATORIO**
# of patients	N=821	N=835	N=732
Design	96-week treatment period: ▪ **ARM A:** Ocrevus 2x300mg IV followed by 600mg IV every 24 weeks ▪ **ARM B:** Interferon β-1a (Rebif)	96-week treatment period: ▪ **ARM A:** Ocrevus 2x300mg IV followed by 600mg IV every 24 weeks ▪ **ARM B:** Interferon β-1a (Rebif)	120-week treatment period: ▪ **ARM A:** Ocrevus 2x300mg IV every 24 weeks ▪ **ARM B:** Placebo
Primary endpoint	▪ Annualized relapse rate at 96 weeks versus Rebif	▪ Annualized relapse rate at 96 weeks versus Rebif	▪ Sustained disability progression versus placebo by EDSS
Status	▪ Primary endpoint met Q2 2015, OLE ongoing ▪ Primary data presented at ECTRIMS 2015 ▪ Updated data presented at AAN and ECTRIMS 2017, AAN and EAN 2018 ▪ Data published in *NEJM* 2017; 376:221-234 ▪ Data published on COVID-19 in *Mult Scler Relat Disord* on Ocrevus treated people with MS, doi.org/10.1016/j.msard.2020.102725		▪ Primary endpoint met Q3 2015 ▪ Primary data presented at ECTRIMS 2015, updated data presented at AAN and ECTRIMS 2017, AAN and EAN 2018 ▪ Data published in *NEJM* 2017; 376:209-220
	▪ Approved in US Q1 2017 and EU Q1 2018		
CT Identifier	NCT01247324	NCT01412333	NCT01194570

IV=intravenous; EDSS=Expanded Disability Status Scale; OLE=Open label extension; ECTRIMS=European Committee for Treatment and Research in Multiple Sclerosis; AAN=Annual Meeting of the American Academy of Neurology; EAN=European Academy of Neurology; NEJM=New England Journal of Medicine

Ocrevus (ocrelizumab, RG1594)

Humanized monoclonal antibody selectively targeting CD20+ B cells

Indication	Relapsing and primary progressive multiple sclerosis (RMS & PPMS)	Primary progressive multiple sclerosis (PPMS)
Phase/study	**Phase IIIb** **ENSEMBLE PLUS**	**Phase IIIb** **ORATORIO-HAND**
# of patients	N=1,225	N ~ 1000
Design	• Substudy of ongoing phase IIIb, open-label, single-arm ENSEMBLE study • Shorter two-hour infusion time	120-week treatment period: ▪ **ARM A:** Ocrevus 600mg IV q24w ▪ **ARM B:** Placebo
Primary endpoint	▪ Safety, measured by the proportion of patients with IRRs following the first randomised 600 mg infusion (frequency/severity assessed during and 24-hours post infusion)	▪ Time to upper limb disability progression confirmed for at least 12 weeks
Status	• Filed in US and EU Q1 2020 • Approved in EU Q2 2020 and US Q4 2020 • Data published *Neurol, Neuroimmunol* and *Neuroinflamm* Sept 2020; 7(5), e807	▪ FPI Q3 2019
CT Identifier	NCT03085810	NCT04035005

IV=intravenous; IRR=Infusion Related Reaction

Ocrevus (ocrelizumab, RG1594)

Humanized monoclonal antibody selectively targeting CD20+ B cells

Indication	Primary progressive multiple sclerosis (PPMS)	Relapsing multiple sclerosis (RMS)	PPMS & RMS
Phase/study	**Phase IIIb GAVOTTE**	**Phase IIIb MUSETTE**	**Phase III Ocarina II[1]**
# of patients	N ~ 699	N ~ 786	N ~ 232
Design	120-week treatment period: ▪ **ARM A:** Ocrevus 600mg IV every 24 weeks ▪ **ARM B:** Ocrevus 1200mg if body weight <75kg or 1800mg if body weight > or equal to 75kg every 24 weeks	120-week treatment period: ▪ **ARM A:** Ocrevus 600mg IV every 24 weeks ▪ **ARM B:** Ocrevus 1200mg if body weight <75kg or 1800mg if body weight > or equal to 75kg every 24 weeks	▪ **ARM A:** Ocrevus IV ▪ **ARM B:** Ocrevus SC
Primary endpoint	▪ Superiority of Ocrevus higher dose versus approved dose on cCDP	▪ Superiority of Ocrevus higher dose versus approved dose on cCDP	▪ Serum Ocrevus area under the concentration-time curve (AUCW1-12) at week 12
Status	▪ FPI Q4 2020	▪ FPI Q4 2020 ▪ Recruitment completed Q4 2021	▪ FPI Q2 2022
CT Identifier	NCT04548999	NCT04544436	NCT05232825

[1]SC with Halozyme's rHuPH20/ Halozyme's human hyaluronidase
cCDP=composite confirmed disability progression; IV=intravenous; SC=Subcutaneous

Evrysdi (risdiplam, RG7916)

Oral SMN2 splicing modifier

Indication	Spinal muscular atrophy (SMA)		
Phase/study	**Phase II/III FIREFISH**	**Phase II/III SUNFISH**	**Phase II JEWELFISH**
# of patients	N=21 (Part 1), 41 (Part 2)	N=51 (Part 1), 180 (Part 2)	N=174
Design	Open-label study in infants with type 1 SMA ▪ **Part 1 (dose-finding):** At least 4 weeks ▪ **Part 2 (confirmatory):** 24 months	Randomized, double-blind, placebo-controlled study in adult and pediatric patients with type 2 or type 3 SMA: ▪ **Part 1 (dose-finding):** At least 12 weeks ▪ **Part 2 (confirmatory):** 24 months	▪ Open-label single arm study in adult and pediatric patients with previously treated SMA type 1, 2 and 3
Primary endpoint	▪ Safety, tolerability, PK/PD and efficacy	▪ Safety, tolerability, PK/PD and efficacy	▪ Safety, tolerability, PK/PD
Status	▪ 12-month data from Part 1 presented at AAN, CureSMA and EAN 2019; 16-month data presented at WMS 2019 ▪ Study met primary endpoint in part 2 Q1 2020 ▪ Part 2 1-year data presented at AAN 2020, part 1 2-year data at WMS 2020 ▪ Part 1 data published in *NEJM* 2021;384:915-923 ▪ Part 2 2-year data presented at AAN 2021 ▪ Part 2 1-year data published in *NEJM* 2021;385:427-435 ▪ 3-year data presented at EPNS 2022	▪ Recruitment completed for part 2 Q3 2018 ▪ 12-month data from Part 1 presented at AAN, CureSMA and EAN 2019; 16-month data presented at WMS 2019 ▪ Study met primary endpoint in part 2 Q4 2019 ▪ Part 2 1-year data presented at SMA Europe 2020, 2-year data at MDA 2021 and 3-year data at MDA 2022 ▪ Part 2 data 1-year published in Lancet Neurology, Dec 2021	▪ FPI Q1 2017 ▪ Data presented at WMS 2017, AAN 2018, WMS 2018, CureSMA 2019, WMS 2019, CureSMA 2020 and 2021 ▪ Recruitment completed Q1 2020
	▪ Orphan drug designation granted by FDA Q1 2017 and EU Q1 2019, PRIME designation in Q4 2018 ▪ Approved in US Q3 2020 and EU Q1 2021		
CT Identifier	NCT02913482	NCT02908685	NCT03032172

In collaboration with PTC Therapeutics and SMA Foundation

SMA=Spinal muscular atrophy; SMN=survival motor neuron; PK/PD=Pharmacokinetics/Pharmacodynamics; PRIME=priority medicines; AAN=American Academy of Neurology; WMS=World Muscle Society; EAN=European Academy of Neurology; NEJM=New England Journal of Medicine; MDA=Muscular Dystrophy Association; CureSMA=Annual SMA Conference; EPNS=European Paediatric Neurology Society

Evrysdi (risdiplam, RG7916)

Oral SMN2 splicing modifier

Indication	Spinal muscular atrophy (SMA)	
Phase/study	**Phase II** **RAINBOWFISH**	**Phase II/III** **MANATEE**
# of patients	N=25	N=180
Design	Open-label, single-arm, multicenter study in infants aged from birth to 6 weeks who have been genetically diagnosed with Spinal muscular atrophy but are not yet presenting with symptoms	**ARM A:** ▪ Part 1: GYM329 plus Evrysdi for 24 weeks, followed by GYM329 plus Evrysdi for 72 weeks ▪ Part 2: GYM329 plus Evrysdi for 72 weeks **ARM B:** ▪ Placebo plus Evrysdi comparator
Primary endpoint	▪ Proportion of participants with two copies of the SMN2 gene (excluding the known SMN2 gene modifier mutation c.859G>C) and baseline CMAP>=1.5 millivolt who are sitting without support	▪ Change from baseline in revised hammersmith scale (RHS) score after week 72 of treatment ▪ Safety, PK/PD and muscle biomarkers
Status	▪ FPI Q3 2019 ▪ Recruitment completed Q1 2022 ▪ Initial data presented at CureSMA , WMS 2021 and MDA 2022 ▪ Filed in US and EU Q4 2021 ▪ Approved in US Q2 2022	▪ FPI Part 1 Q2 2022 ▪ Orphan Drug Designation granted by FDA in Q4 2021 for GYM329
CT Identifier	NCT03779334	NCT05115110

In collaboration with PTC Therapeutics and SMA Foundation

SMN=survival motor neuron; CMAP=compound muscle action potential; PK/PD=Pharmacokinetics/Pharmacodynamics; WMS=World Muscle Society; CureSMA=Annual SMA Conference; MDA=Muscular Dystrophy Association

Enspryng (satralizumab, RG6168, SA237)

Anti-IL-6 receptor humanized monoclonal antibody

Indication	Neuromyelitis optica spectrum disorder (NMOSD)	
Phase/study	**Phase III** **SAkuraStar**	**Phase III** **SAkuraSky**
# of patients	N=95	N=83
Design	Enspryng monotherapy: • **ARM A:** Enspryng 120mg SC monthly • **ARM B:** Placebo SC monthly	Add-on therapy of Enspryng: • **ARM A:** Enspryng 120mg SC monthly • **ARM B:** Placebo SC monthly Both arms on top of baseline therapies: azathioprine, mycophenolate mofetil or oral corticosteroids
Primary endpoint	•Efficacy (time to first relapse), safety and PK/PD	▪ Efficacy (time to first relapse), safety and PK/PD
Status	▪ Primary endpoint met Q4 2018 ▪ Data presented at ECTRIMS 2019 ▪ Published in Lancet Neurology 2020; 19(5): 402-412	▪ FPI Q3 2017 ▪ Primary endpoint met Q3 2018 ▪ Data presented at ECTRIMS 2018 and AAN 2019 ▪ Published in *NEJM* 2019; 381:2114-2124
	▪ BTD granted by FDA Q4 2018 ▪ Filed in EU Q3 2019; US acceptance of filing Q4 2019 ▪ Approved in US Q3 2020 and EU Q2 2021	
CT Identifier	NCT02073279	NCT02028884

*Trials managed by Chugai (Roche opted-in)

BTD=Breakthrough therapy designation; PK/PD=Pharmacokinetics/Pharmacodynamics; SC=Subcutaneous; ECTRIMS=European Committee for Treatment and Research in Multiple Sclerosis; AAN=American Academy of Neurology; NEJM=New England Journal of Medicine

Enspryng (satralizumab, RG6168, SA237)

Anti-IL-6 receptor humanized monoclonal antibody

Indication	Generalised myasthenia gravis (MG)	Myelin oligodendrocyte glycoprotein antibody disease (MOGAD)
Phase/study	**Phase III** **Luminesce**	**Phase III** **METEOROID**
# of patients	N=240	N=152
Design	• **ARM A:** Enspryng plus standard of care • **ARM B:** Placebo plus standard of care	•**ARM A**: Enspryng at weeks 0, 2, 4 (loading doses) and maintenance doses q4w •**ARM B:** Placebo
Primary endpoint	▪ Mean change from baseline in total MG-ADL score at week 24 in AChR+ population	▪ Time from randomization to the first occurrence of a MOGAD relapse
Status	▪ Orphan Drug Designation granted in US Q1 2021 ▪ FPI Q4 2021	▪ FPI expected Q3 2022 ▪ Orphan Drug Designation granted by FDA in Q4 2021
CT Identifier	NCT04963270	NCT05271409

In collaboration with Chugai

MG-ADL= Myasthenia Gravis Activities of Daily Living; AChR=Acetylcholine receptor; MOGAD=Myelin Oligodendrocyte Glycoprotein Antibody Disease

Gazyva (obinutuzumab)

Immunology development program

Indication	Lupus nephritis		Membranous nephropathy
Phase/study	**Phase II NOBILITY**	**Phase III REGENCY**	**Phase III MAJESTY**
# of patients	N=126	N=252	N=140
Design	▪ **ARM A:** Gazyva 1000mg IV plus mycophenolate mofetil / mycophenolic acid ▪ **ARM B:** Placebo IV plus mycophenolate mofetil / mycophenolic acid	▪ **ARM A:** Gazyva 1000mg IV (6 doses through Week 52) plus mycophenolate mofetil ▪ **ARM B:** Gazyva1000 mg IV (5 doses through Week 52) plus mycophenolate mofetil ▪ **ARM C:** Placebo IV plus mycophenolate mofetil	▪ **ARM A:** Gazyva 1000mg IV dosed at baseline and weeks 0, 2, 24, and 26 on top of renin-angiotensin inhibitors ▪ **ARM B:** Tacrolimus treatment for 12 months
Primary endpoint	▪ Percentage of participants who achieve complete renal response (CRR)	▪ Percentage of participants who achieve complete renal response (CRR)	▪ Percentage of patients who achieve complete remission at week 104
Status	▪ Recruitment completed Q4 2017 ▪ Primary endpoint met Q2 2019 ▪ BTD granted by the FDA Q3 2019 ▪ Data presented at ASN and ACR 2019 ▪ Published in *Ann Rheum Dis* 2022 Jan;81(1):100-107	▪ FPI Q3 2020	▪ FPI Q2 2021
CT Identifier	NCT02550652	NCT04221477	NCT04629248

In collaboration with Biogen

BTD=Breakthrough therapy designation; IV=Intravenous; ASN=American Society of Nephrology; ACR=American College of Rheumatology

Gazyva (obinutuzumab)

Immunology development program

Indication	Systemic lupus erythematosus (SLE)
Phase/study	**Phase III** **ALLEGORY**
# of patients	N=200
Design	• **ARM A:** Gazyva 1000mg IV on Day 1 and Weeks 2, 24 and 26. • **ARM B:** Placebo IV
Primary endpoint	• Percentage of participants who achieve Systemic Lupus Erythematosus Responder Index (SRI) at week 52
Status	• FPI Q4 2021
CT Identifier	NCT04963296

In collaboration with Biogen
IV=Intravenous

Actemra/RoActemra (tocilizumab, RG-1569)

Interleukin 6 receptor inhibitor

Indication	Adult hospitalised with severe COVID-19 pneumonia	
Phase/study	**Phase III COVACTA[1]**	**Phase III REMDACTA[2]**
# of patients	N=450	N=650
Design	▪ **ARM A:** Actemra plus standard of care ▪ **ARM B:** Placebo plus standard of care	▪ **ARM A:** Remdesivir plus Actemra ▪ **ARM B:** Remdesivir plus placebo
Primary endpoint	▪ Clinical status assessed using 7-Category Ordinal Scale (Day 28)	▪ Time to hospital discharge or ready for discharge
Status	▪ FPI Q1 2020 ▪ Recruitment completed Q2 2020 ▪ Primary endpoint not met Q3 2020 ▪ Published in *NEJM* 2021; 384:1503-1516	▪ FPI Q2 2020 ▪ Recruitment completed Q1 2021 ▪ Primary endpoint not met Q1 2021 ▪ Published in *Intensive Care Med* 2021 doi: 10.1007/s00134-021-06507-x
	▪ Filed in EU Q3 2021 ▪ Approved in EU Q4 2021	
CT Identifier	NCT04320615	NCT04409262

[1]In collaboration with US Biomedical Advanced Research and Development Authority (BARDA); [2]In collaboration with Gilead Sciences, Inc.
NEJM=New England Journal of Medicine

Actemra/RoActemra (tocilizumab, RG-1569)

Interleukin 6 receptor inhibitor

Indication	Adult hospitalised with severe COVID-19 pneumonia	
Phase/study	**Phase II** **MARIPOSA**	**Phase III** **EMPACTA**
# of patients	N=100	N=379
Design	▪ **ARM A:** 8 mg/kg Actemra plus standard of care ▪ **ARM B:** 4mg/kg Actemra plus standard of care	Conducted in sites known to provide critical care to underserved and minority populations that often do not have access to clinical trials ▪ **ARM A:** Actemra plus standard of care ▪ **ARM B:** Placebo plus standard of care
Primary endpoint	▪ Pharmacodynamics and pharmacokinetics	▪ Cumulative proportion of participants requiring mechanical ventilation by day 28
Status	▪ FPI Q2 2020 ▪ Recruitment completed Q2 2020 ▪ Published in *Open Forum Infect Dis* 2021 Dec 4;9(1)	▪ FPI Q2 2020 ▪ Primary endpoint met Q3 2020 ▪ Published in *NEJM* 2021 Jan 7;384(1):20-30
	▪ Filed in EU Q3 2021 ▪ Approved in EU Q4 2021	
CT Identifier	NCT04363736	NCT04372186

NEJM=New England Journal of Medicine

Xolair

Humanized monoclonal antibody that selectively binds to IgE

Indication	Food allergy
Phase/study	**Phase III** **OUtMATCH[1]**
# of patients	N=225
Design	• Xolair by SC injection either q2w or q4w for 16 to 20 weeks
Primary endpoint	• Number of participants who successfully consume ≥600mg of peanut protein without dose-limiting symptoms
Status	• FPI Q3 2019
CT Identifier	NCT03881696

In collaboration with Novartis; [1] Sponsor of the study is the National Institute of Allergy and Infectious Diseases (NIAID)
IgE=Immunoglobulin E; SC=Subcutaneous

Lunsumio (mosunetuzumab, CD20 x CD3, RG7828)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	Systemic lupus erythematosus (SLE)
Phase/study	**Phase I**
# of patients	N=50
Design	▪ **ARM A**: Lunsumio SC on either Day 1 or on Days 1 and 8 ▪ **ARM B:** Fractionated (divided) dose of Lunsumio SC on Days 1 and 8
Primary endpoint	▪ Safety
Status	▪ FPI January 2022
CT Identifier	NCT05155345

SC=subcutaneous

Susvimo (PDS)

First eye implant to achieve sustained delivery of a biologic medicine

Indication	Wet age-related macular degeneration (wAMD)		
Phase/study	**Phase III Archway**	**Phase II+III extension Portal**	**Phase IIIb Velodrome**
# of patients	N=418	N=1,000	N=442
Design	▪ **ARM A:** Port delivery system with ranibizumab q24w ▪ **ARM B:** Intravitreal ranibizumab q4w	▪ Patients from LADDER or Archway will receive refills of 100mg/mL ranibizumab q24w (patients without the PDS will receive the PDS and subsequent refills)	▪ **ARM A:** Port delivery system with ranibizumab q36w ▪ **ARM B:** Port delivery system with ranibizumab q24w
Primary endpoint	▪ Change in BCVA from baseline at the average of week 36 and week 40	▪ Safety and long term efficacy	▪ Change in BCVA from baseline averaged over weeks 68 and 72
Status	▪ FPI Q3 2018 ▪ Recruitment completed Q2 2019 ▪ Study met primary endpoint Q2 2020 ▪ Primary endpoint data presented at ASRS 2020, 44/48 week data at Angiogenesis 2021 and 2-year data at Angiogenesis 2022 ▪ Filed in US (PRIME) and EU Q2 2021 ▪ Approved in US Q4 2021	▪ FPI Q3 2018	▪ FPI Q3 2021
CT Identifier	NCT03677934	NCT03683251	NCT04657289

BCVA=best corrected visual acuity; wAMD=wet age-related macular degeneration; ASRS=American Society of Retinal Specialists; PDS=Port Delivery System with ranibizumab; PRIME=Priority review

Susvimo (PDS)

First eye implant to achieve sustained delivery of a biologic medicine

Indication	Diabetic macular edema (DME)	Diabetic retinopathy (DR) without center-involved diabetic macular edema (DME)
Phase/study	**Phase III Pagoda**	**Phase III Pavilion**
# of patients	N=545	N=160
Design	▪ **ARM A:** Port delivery system with ranibizumab q24w ▪ **ARM B:** Intravitreal ranibizumab q4w	▪ **Arm A:** Intravitreal ranibizumab (X2) followed by PDS implant (refill q36w) ▪ **Arm B:** Q4w comprehensive clinical monitoring until participants receive PDS (refill q36w)
Primary endpoint	▪ Change in BCVA from baseline at the average of week 48 and week 52	▪ Percentage of participants with a ≥2-step improvement from baseline on the ETDRS-DRSS at Week 52
Status	▪ FPI Q3 2019 ▪ Recruitment completed Q2 2021	▪ FPI Q3 2020 ▪ Recruitment completed Q3 2021
CT Identifier	NCT04108156	NCT04503551

BCVA=best corrected visual acuity; ETDRS=Early Treatment Diabetic Retinopathy Study; DRSS=Diabetic Retinopathy Severity Scale; PDS=Port Delivery System with ranibizumab

Vabysmo (faricimab)

Bispecific antibody to simultaneously bind Ang-2 and VEGF-A

Indication	Center-involving diabetic macular edema (CI-DME)	
Phase/study	**Phase III YOSEMITE**	**Phase III RHINE**
# of patients	N=940	N=951
Design	▪ **ARM A:** Faricimab q8w ▪ **ARM B:** Faricimab PTI up to q16w ▪ **ARM C:** Aflibercept, q8w	▪ **ARM A:** Faricimab q8w ▪ **ARM B:** Faricimab PTI up to q16w ▪ **ARM C:** Aflibercept, q8w
Primary endpoint	▪ Change from baseline in BCVA at 1 year	▪ Change from baseline in BCVA at 1 year
Status	▪ FPI Q3 2018 ▪ Recruitment completed Q3 2019 ▪ Study met primary endpoint Q4 2020 ▪ Data presented at Angiogenesis 2021	▪ FPI Q4 2018 ▪ Recruitment completed Q3 2019 ▪ Study met primary endpoint Q4 2020 ▪ Data presented at Angiogenesis 2021
	▪ Filed in US and EU Q2 2021 ▪ Published in the Lancet 2022 Feb 19;399(10326):741-755. ▪ 2-year data presented at Angiogenesis 2022 ▪ Approved in US Q1 2022	
CT Identifier	NCT03622580	NCT03622593

Ang-2=Angiopoietin-2; VEGF=Vascular endothelial growth factor; PTI=Personalized Treatment Interval; BCVA=best corrected visual acuity

Vabysmo (faricimab)

Bispecific antibody to simultaneously bind Ang-2 and VEGF-A

Indication	Wet age related macular degeneration (wAMD)	
Phase/study	**Phase III TENAYA**	**Phase III LUCERNE**
# of patients	N=671	N=658
Design	▪ **ARM A:** Faricimab 6.0mg q16w flexible after 4 IDs ▪ **ARM B:** Aflibercept 2.0mg q8w after 3 IDs	▪ **ARM A:** Faricimab 6.0mg q16w flexible after 4 IDs ▪ **ARM B:** Aflibercept 2.0mg q8w after 3 IDs
Primary endpoint	▪ Change from baseline in BCVA week 40, 44 & 48	▪ Change from baseline in BCVA week 40, 44 & 48
Status	▪ FPI Q1 2019 ▪ Recruitment completed Q4 2019 ▪ Study met primary endpoint Q1 2021 ▪ Data presented at Angiogenesis 2021	▪ FPI Q1 2019 ▪ Recruitment completed Q4 2019 ▪ Study met primary endpoint Q1 2021 ▪ Data presented at Angiogenesis 2021
	▪ Filed in US and EU Q2 2021 ▪ Published in Lancet 2022 Feb 19;399(10326):729-740 ▪ Approved in US Q1 2022 ▪ 2-year data presented at ASRS 2022	
CT Identifier	NCT03823287	NCT03823300

BCVA=best corrected visual acuity; Ang-2=Angiopoietin-2; VEGF=Vascular endothelial growth factor; IDs=initiating doses; ASRS=American Society of Retina Specialists

Vabysmo (faricimab)

Bispecific antibody to simultaneously bind Ang-2 and VEGF-A

Indication	Macular edema (ME) secondary to branch retinal vein occlusion (RVO)	Macular edema (ME) secondary to central retinal vein occlusion (RVO)
Phase/study	**Phase III** **BALATON**	**Phase III** **COMINO**
# of patients	N=570	N=750
Design	▪ **ARM A:** Faricimab, q4w/PTI ▪ **ARM B:** Aflibercept, q4w	▪ **ARM A:** Faricimab, q4w/PTI ▪ **ARM B:** Aflibercept, q4w
Primary endpoint	▪ Change from baseline in BCVA at week 24	▪ Change from baseline in BCVA at week 24
Status	▪ FPI Q1 2021 ▪ Recruitment completed Q1 2022	▪ FPI Q1 2021 ▪ Recruitment completed Q1 2022
CT Identifier	NCT04740905	NCT04740931

PTI=Personalized Treatment Interval; BCVA=best corrected visual acuity; Ang-2=Angiopoietin-2; VEGF=Vascular endothelial growth factor

Xofluza (baloxavir marboxil, RG6152, S-033188)

Small molecule, novel CAP-dependent endonuclease inhibitor

Indication	Influenza		
Phase/study	**Phase III** **miniSTONE 1 (0-1 year old)**	**Phase III** **miniSTONE 2 (1- <12 years old)**	**Phase IIIb** **CENTERSTONE**
# of patients	N=30	N=176	N=3,160
Design	Xofluza on Day 1 (based on body weight and age) in healthy pediatric patients from birth to <1 year with influenza-like symptoms	Healthy pediatric patients 1 to <12 years of age with influenza-like symptoms ▪ **ARM A:** Xofluza ▪ **ARM B:** Tamiflu	Reduction of direct transmission of influenza from otherwise healthy patients to household contacts ▪ **ARM A:** Xofluza ▪ **ARM B:** Placebo
Primary endpoint	▪ Safety	▪ Safety	▪ Percentage of household contacts who are PCR-positive for influenza by day 5 post randomization of index patients
Status	• FPI Q1 2019	• Primary endpoint met Q2 2019 • Data presented at OPTIONS X 2019 • Filed in US Q1 2020 • Data published in *Pediatric Infectious Disease* 2020 Aug;39(8):700-705 • Not approved in the US, determining path forward with the FDA • Filed in EU Q4 2021	▪ FPI Q4 2019
CT Identifier	NCT03653364	NCT03629184	NCT03969212

In collaboration with Shionogi & Co., Ltd.
CAP=Catabolite Activating Protein

Roche Group development pipeline

Marketed products development programmes

Roche Pharma global development programmes

Roche Pharma research and early development (pRED)

Genentech research and early development (gRED)

Spark

Ipatasertib (RG7440, GDC-0068)

Highly selective small molecule inhibitor of Akt

Indication	1L castration-resistant prostate cancer (CRPC)
Phase/study	**Phase III** **IPATential150**
# of patients	N=1,100
Design	▪ **ARM A:** Ipatasertib plus abiraterone ▪ **ARM B:** Placebo plus abiraterone
Primary endpoint	▪ rPFS in patients with PTEN loss tumors and overall population
Status	▪ FPI Q2 2017 ▪ Recruitment completed Q1 2019 ▪ Study met co-primary endpoint in rPFS in patients with PTEN loss tumors Q2 2020 ▪ Data presented at ESMO 2020 and interim OS at ASCO 2022 ▪ Published in Lancet 2021; 398:131-142
CT Identifier	NCT03072238

Akt=Protein Kinase B; PTEN=Phosphatase and Tensin homolog; rPFS=Radiographic progression-free survival ; ESMO=European Society for Medical Oncology

Tiragolumab (anti-TIGIT, RG6058, MTIG7192A)

Monoclonal antibody targeting the immune checkpoint inhibitor TIGIT

Indication	1L NSCLC PD-L1 TPS>50%	Stage III unresectable 1L NSCLC
Phase/study	**Phase III** **SKYSCRAPER-01**	**Phase III** **SKYSCRAPER-03**
# of patients	N=500-560	N=800
Design	▪ **ARM A:** Tiragolumab plus Tecentriq ▪ **ARM B:** Placebo plus Tecentriq	▪ **ARM A:** Tiragolumab plus Tecentriq for up to 12 months ▪ **ARM B:** Durvalumab for up to 12 months
Primary endpoint	▪ Overall survival and progression-free survival	▪ Progression-free survival
Status	▪ FPI Q1 2020 ▪ Recruitment completed Q3 2021 ▪ Study did not meet its co-primary endpoint of PFS Q2 2022	▪ FPI Q3 2020
CT Identifier	NCT04294810	NCT04513925

NSCLC=Non-small cell lung cancer; ES-SCLC=Extensive stage small cell lung cancer; PD-L1=Programmed cell death-ligand 1; TPS=Tumor Proportion Score; PFS=Progression-free survival

Tiragolumab (anti-TIGIT, RG6058, MTIG7192A)

Monoclonal antibody targeting the immune checkpoint inhibitor TIGIT

Indication	Metastatic and/or recurrent PD-L1+ cervical cancer (CC)	Neoadjuvant and adjuvant NSCLC	1L non-squamous NSCLC
Phase/study	**Phase II SKYSCRAPER-04**	**Phase II SKYSCRAPER-05**	**Phase II/III SKYSCRAPER-06**
# of patients	N=172	N=82	N=500
Design	▪ **ARM A:** Tiragolumab plus Tecentriq ▪ **ARM B:** Tecentriq	▪ **ARM A:** (PD-L1 high) neoadjuvant tiragolumab plus Tecentriq followed by adjuvant tiragolumab plus Tecentriq or adjuvant chemotherapy ▪ **ARM B:** (PD-L1 all-comers) neoadjuvant tiragolumab plus Tecentriq plus chemo followed by adjuvant tiragolumab plus Tecentriq	▪ **ARM A:** Tiragolumab plus Tecentriq plus pemetrexed plus chemo followed by maintenance tiragolumab plus Tecentriq plus pemetrexed ▪ **ARM B:** Placebo plus pembrolizumab plus pemetrexed plus chemo followed by maintenance placebo plus pembrolizumab plus pemetrexed
Primary endpoint	▪ Objective response rate	▪ Pathologic complete response, major pathological response and safety	▪ Objective response rate, progression-free survival and overall survival
Status	▪ FPI Q2 2020	▪ FPI Q2 2021	▪ FPI Q4 2020
CT Identifier	NCT04300647	NCT04832854	NCT04619797

NSCLC=Non-small cell lung cancer; PD-L1=Programmed cell death-ligand 1

Tiragolumab (anti-TIGIT, RG6058, MTIG7192A)

Monoclonal antibody targeting the immune checkpoint inhibitor TIGIT

Indication	Locally advanced esophageal cancer (EC)	1L esophageal cancer (EC)	1L recurrent/metastatic PD-L1 positive squamous cell head and neck carcinoma (SCCHN)
Phase/study	**Phase III SKYSCRAPER-07**	**Phase III SKYSCRAPER-08**	**Phase II SKYSCRAPER-09**
# of patients	N=750	N=500	N=120
Design	▪ **ARM A:** Tiragolumab plus Tecentriq ▪ **ARM B:** Tecentriq plus placebo ▪ **ARM C:** Placebo plus placebo	▪ **ARM A:** Tiragolumab plus Tecentriq plus cisplatin and paclitaxel ▪ **ARM B:** Placebo plus placebo plus cisplatin and paclitaxel	▪ **ARM A:** Tiragolumab plus Tecentriq ▪ **ARM B:** Tecentriq plus placebo
Primary endpoint	▪ Progression-free survival (A vs C) ▪ Overall survival (A vs C, hierarchical, B vs C hierarchical)	▪ Overall survival and progression-free survival	▪ Objective response rate
Status	▪ FPI Q3 2020	▪ FPI Q4 2020 ▪ Recruitment completed Q4 2021	▪ FPI Q1 2021 ▪ Recruitment completed Q2 2022
CT Identifier	NCT04543617	NCT04540211	NCT04665843

NSCLC=Non-small cell lung cancer; PD-L1=Programmed cell death-ligand 1

Tiragolumab (anti-TIGIT, RG6058, MTIG7192A)

Monoclonal antibody targeting the immune checkpoint inhibitor TIGIT

Indication	Solid tumors	NSCLC	Relapsed or refractory multiple myeloma (MM) or r/r B-cell NHL
Phase/study	**Phase I**	**Phase II CITYSCAPE**	**Phase I**
# of patients	N=540	N=135	N=52
Design	▪ **Phase Ia:** Dose escalation and expansion of tiragolumab ▪ **Phase Ib:** Dose escalation and expansion of tiragolumab in combination with Tecentriq and/or other anti-cancer therapies	▪ **ARM A:** Tecentriq plus tiragolumab ▪ **ARM B:** Tecentriq monotherapy	▪ **Phase Ia:** Tiragolumab monotherapy ▪ **Phase Ib:** Tiragolumab plus daratumumab (r/r MM) or rituximab (r/r NHL)
Primary endpoint	▪ Safety, tolerability, PK variability and preliminary efficacy	▪ Overall response rate and progression-free survival	▪ Safety, tolerability, PK/PD and preliminary efficacy
Status	▪ FPI Q2 2016 ▪ Data presented at AACR 2020	▪ FPI Q3 2018 ▪ Recruitment completed Q2 2019 ▪ Data presented at ASCO 2020 and WCLC and ESMO IO 2021 ▪ BTD granted by FDA Q4 2020	▪ FPI Q2 2019
CT Identifier	NCT02794571	NCT03563716	NCT04045028

BTD=Breakthrough therapy designation; MM=Multiple myeloma; NSCLC=Non-small cell lung cancer; r/r=Relapsed refractory; NHL=Non-Hodgkin's lymphoma; PK=Pharmacokinetics; PD=Pharmacodynamics; ASCO=American Society of Clinical Oncology; AACR=American Association for Cancer Research; WCLC=World Conference on Lung Cancer; ESMO IO=European Society for Medical Oncology - Immuno-Oncology

Glofitamab (CD20-TCB, RG6026)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	Relapsed or refractory Non-Hodgkin's lymphoma (NHL)		
Phase/study	**Phase I**	**Phase Ib**	**Phase I**
# of patients	N=700	N=140	N=18-36
Design	**Cohort 1:** Single-agent dose escalation study ▪ Initial dose escalation ▪ Expansion cohort in r/r DLBCL ▪ Expansion cohort in r/r FL All patients will receive pretreatment with a single dose of Gazyva (1000mg) **Cohort 2:** Glofitamab plus Gazyva (i.e. continuous treatment with Gazyva)	Dose escalation and expansion ▪ **ARM A:** Glofitamab plus Tecentriq ▪ **ARM B:** Glofitamab plus Polivy	Glofitamab SC ▪ Part 1 dose escalation
Primary endpoint	▪ Efficacy, safety, tolerability and pharmacokinetics	▪ Safety	▪ Safety
Status	▪ FPI Q1 2017 ▪ Data presented at ASH 2018, ICML and ASH 2019; EHA and ASH 2020; ASCO, EHA, ICML and ASH 2021; ASCO and EHA 2022 ▪ Data published online March 2021 *J Clin Oncology* 39:18:1959-1970 ▪ Filed in EU April 2022	▪ Arm A: FPI Q2 2018 ▪ Data presented at ASH 2019 and ASH 2021 ▪ Arm B: FPI Q4 2020	▪ FPI Q3 2021
CT Identifier	NCT03075696	NCT03533283	ISRCTN17975931

DLBCL=diffuse large B cell lymphoma; FL=Follicular lymphoma; r/r=Relapsed or refractory; SC=subcutenous; ASCO=American Society of Clinical Oncology; ASH=American Society of Hematology; EHA=European Hematology Association; ICML=International Conference on Malignant Lymphoma

Glofitamab (CD20-TCB, RG6026)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	Non-Hodgkin's lymphoma (NHL)	2L+ SCT-ineligible DLBCL
Phase/study	**Phase Ib**	**Phase III** **STARGLO**
# of patients	Part I: 15-60 Part II: ~66-104	N=270
Design	▪ **Part I:** Dose-finding for the combination of glofitamab plus G/R-CHOP in r/r indolent NHL ▪ **Part II:** Dose expansion glofitamab plus G/R-CHOP or R-CHOP in 1L DLBCL ▪ **Part III:** Glofitamab plus R-CHP plus Polivy	▪ **ARM A:** Glofitamab plus gemcitabine and oxaliplatin, followed by up to 4 cycles of glofitamab monotherapy ▪ **ARM B:** Rituxan in combination with gemcitabine and oxaliplatin ▪ A single dose of Gazyva will be administered 7 days prior to the first dose of glofitamab
Primary endpoint	▪ Safety	▪ Overall survival
Status	▪ Part I: FPI Q1 2018 ▪ Part II: FPI Q1 2021 ▪ Data presented at ASH 2021	▪ FPI Q1 2021
CT Identifier	NCT03467373	NCT04408638

DLBCL=diffuse large B cell lymphoma; SCT=stem cell transplant; CHOP=cyclophosphamide, doxorubicin, vincristine and prednisone; R=Rituxan/MabThera; G=Gazyva; NHL=Non-Hodgkin's lymphoma; r/r=Relapsed or refractory
ASH=American Society of Hematology

Glofitamab (CD20-TCB, RG6026)

Bispecific anti-CD20/CD3 antibody engaging T and B cells simultaneously

Indication	1L ctDNA high risk DLBCL
Phase/study	**Phase II**
# of patients	N=40
Design	▪ Glofitamab plus R-CHOP (glofitamab is introduced as a consolidation to R-CHOP at cycle 3-8 in patients ctDNA+ at cycle 2)
Primary endpoint	▪ EOT PET-CR
Status	▪ FPI Q1 2022
CT Identifier	NCT04980222

ctDNA=circulating tumor DNA; DLBCL=diffuse large B cell lymphoma; R-CHOP=Rituxan, cyclophosphamide, doxorubicin, vincristine and prednisone; PET=positron emission tomography; EOT PET-CR=End of treatment PET-complete response rate

Inavolisib (RG6114, GDC-0077)

A potent, orally available, and selective PI3Kα inhibitor

Indication	PIK3CA-mutant HR+ metastatic breast cancer (mBC)	PIK3CA mutant solid tumors and metastatic ER+ HER2-neg breast cancer
Phase/study	**Phase III INAVO120**	**Phase I**
# of patients	N=400	N=256
Design	▪ **ARM A:** Inavolisib plus palbociclib plus fulvestrant ▪ **ARM B:** Placebo plus palbociclib plus fulvestrant	Monotherapy and in combination with standard of care (letrozole; letrozole plus palbociclib; fulvestrant) • **Stage 1:** Dose escalation • **Stage 2:** Dose expansion
Primary endpoint	▪ Progression-free survival	• Safety, tolerability and pharmacokinetics
Status	▪ FPI Q1 2020	• FPI Q4 2016 • Preclinical/molecule discovery data presented at AACR 2017 • Data presented at SABCS 2019, 2020 and 2021
CT Identifier	NCT04191499	NCT03006172

ER=Estrogen receptor; HR=Hormon receptor; HER2=Human Epidermal growth factor Receptor 2; PI3K=Phosphoinositide 3-Kinase; AACR=American Association for Cancer Research; SABCS=San Antonio Breast Cancer Symposium

Giredestrant (SERD (3),RG6171, GDC-9545)

A selective estrogen receptor degrader or downregulator

Indication	ER+ HER2-neg metastatic breast cancer (mBC)	ER+ HER2-neg Stage I-III operable breast cancer (BC)	Neoadjuvant ER+ breast cancer (BC)
Phase/study	**Phase I**	**Phase I**	**Phase II coopERA Breast Cancer**
# of patients	N=181	N=75	N=221
Design	▪ Dose escalation and expansion at RPTD ▪ Giredestrant monotherapy and in combination with palbociclib and/or LHRH agonist	▪ Open-label, pre-operative administration ▪ Dose escalation	• **ARM A:** Giredestrant followed by giredestrant plus palbociclib • **ARM B:** Anastrazole followed by anastrazole plus palbociclib
Primary endpoint	▪ Safety	▪ Safety, tolerability and PK/PD	▪ Safety, tolerability and PK/PD
Status	▪ FPI Q4 2017 ▪ Data presented at SABCS 2019, ASCO 2020, ASCO 2021 and SABCS 2021	▪ FPI Q3 2019 ▪ Data presented at ASCO 2021	▪ FPI Q3 2020 ▪ Data presented at ESMO and SABCS 2021; ASCO 2022
CT Identifier	NCT03332797	NCT03916744	NCT04436744

ER=Estrogen receptor; HER2=Human Epidermal growth factor Receptor; RPTD=Recommended phase II dose; LHRH=Luteinizing hormone–releasing hormone; PK/PD=Pharmacokinetics/Pharmacodynamics; SABCS=San Antonio Breast Cancer Symposium; ASCO=American Society of Clinical Oncology

Giredestrant (SERD (3),RG6171, GDC-9545)

A selective estrogen receptor degrader or downregulator

Indication	2L/3L ER+/HER2-negative metastatic breast cancer (mBC)	1L ER+ metastatic breast cancer (mBC)	Adjuvant ER+ breast cancer (BC)
Phase/study	**Phase II acelERA Breast Cancer**	**Phase III persevERA Breast Cancer**	**Phase III lidERA Breast Cancer**
# of patients	N=303	N=978	N=4,100
Design	▪ **ARM A:** Giredestrant monotherapy ▪ **ARM B:** Endocrine monotherapy (fulvestrant or aromatase inhibitor)	▪ **ARM A:** Giredestrant plus palbociclib ▪ **ARM B:** Letrozole plus palbociclib	▪ **ARM A:** Giredestrant monotherapy ▪ **ARM B:** Tamoxifen or aromatase inhibitor
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival	▪ Invasive disease-free survival
Status	▪ FPI Q4 2020 ▪ Recruitment completed Q4 2021 ▪ Study did not meet its primary endpoint Q2 2022	▪ FPI Q4 2020	▪ FPI Q3 2021
CT Identifier	NCT04576455	NCT04546009	NCT04961996

ER=Estrogen receptor; HER2=Human Epidermal growth factor Receptor

Giredestrant (SERD (3),RG6171, GDC-9545)

A selective estrogen receptor degrader or downregulator

Indication	1L ER+/HER2-positive breast cancer (BC)
Phase/study	**Phase III** **heredERA**
# of patients	N=812
Design	Induction Phesgo plus taxane followed by maintenance with either: ▪ **ARM A:** Giredestrant plus Phesgo ▪ **ARM B:** Phesgo
Primary endpoint	▪ Progression-free survival
Status	▪ FPI Q2 2022
CT Identifier	NCT05296798

ER=Estrogen receptor; HER2=Human Epidermal growth factor Receptor; Phesgo=FDC of Perjeta and Herceptin for SC administration;

zinpentraxin alfa (PRM-151, RG6354)

Recombinant human innate immunity protein pentraxin-2

Indication	Idiopathic pulmonary fibrosis (IPF)		Myelofibrosis
Phase/study	**Phase II**	**Phase III STARSCAPE**	**Phase II**
# of patients	N=117	N=658	N=125
Design	▪ Randomized, double-blind, placebo-controlled trial: 4-week screening period, 24-week randomized treatment period, 4-week follow-up visit (week 28) ▪ Zinpentraxin alfa at days 1, 3 and 5, then every 4 weeks vs placebo	▪ Randomized, double-blind, placebo-controlled trial: 4-week screening period, 52-week randomized treatment period ▪ Zinpentraxin alfa at days 1, 3 and 5, then every 4 weeks vs placebo	• Multiple dose study of zinpentraxin alfa
Primary endpoint	▪ Least-squares mean change in FVC percentage of predicted value from baseline to week 28	•Absolute change from baseline to week 52 in FVC	• Bone marrow response rate
Status	• Study met primary endpoint • Data published in JAMA 2018;319(22):2299-2307 and Lancet Respir Med 2019 Aug;7(8):657-664	▪ FPI Q1 2021	▪ Study completed Q1 2021
CT Identifier	NCT02550873	NCT04552899	NCT01981850

FVC=Forced vital capacity
JAMA=Journal of the American Medical Association

Crovalimab (RG6107; SKY59)

A humanized monoclonal antibody against complement C5

Indication	Paroxysmal nocturnal hemoglobinuria (PNH)	Paroxysmal nocturnal hemoglobinuria (PNH) patients switching from a C5 inhibitor
Phase/study	**Phase I/II** **COMPOSER**	**Phase III** **COMMODORE 1**
# of patients	N=59	N=250
Design	Healthy volunteers and treatment naïve and pretreated patients with PNH: ▪ **Part 1:** Single ascending dose study in healthy subjects ▪ **Part 2:** Intra-patient single ascending dose study in PNH patients ▪ **Part 3:** Multiple-dose study in PNH patients ▪ **Part 4:** Dose confirmation in PNH patients	▪ **ARM A:** Crovalimab ▪ **ARM B:** Eculizumab ▪ **ARM C:** Patients switching to crovalimab from ravulizumab, higher than labeled doses of eculizumab & C5 SNP patients (descriptive-arm)
Primary endpoint	▪ Safety, PK, PD	▪ Non-inferiority of crovalimab compared to eculizumab - mean % change in LDH level (measure of haemolysis) from baseline to week 25
Status	▪ Part 1: FPI Q4 2016 ▪ Part 2/3: FPI Q2 2017 ▪ Part 4: FPI Q2 2019 ▪ Nonclinical data published in Scientific Reports 2017 Apr; 7(1):1080 ▪ Data presented for Part 2 and 3 at ASH 2018 and 2019	▪ FPI Q3 2020
CT Identifier	NCT03157635	NCT04432584

In collaboration with Chugai

ASH=American Society of Hematology; PNH=Paroxysmal nocturnal hemoglobinuria; PK/PD=Pharmacokinetics/Pharmacodynamics; LDH=Lactate Dehydrogenase

Crovalimab (RG6107; SKY59)

A humanized monoclonal antibody against complement C5

Indication	Paroxysmal nocturnal hemoglobinuria (PNH) C5 inhibitor naive patients	Paroxysmal nocturnal hemoglobinuria (PNH) C5 inhibitor naive patients (China only)
Phase/study	**Phase III** **COMMODORE 2**	**Phase III** **COMMODORE 3**
# of patients	N=200	N=51
Design	▪ **ARM A:** Crovalimab ▪ **ARM B:** Eculizumab	▪ Crovalimab loading dose IV on Day 1, followed by weekly crovalimab SC doses for 4 weeks
Primary endpoint	▪ Non-inferiority of crovalimab compared to eculizumab: - % patients with transfusion avoidance from baseline through week 25 - % patients with haemolysis control, as measured by LDH <=1.5ULN from week 5-25	▪ Percentage of patients with transfusion avoidance from baseline through week 25 ▪ Mean percentage of participants with hemolysis control (week 5 through week 25)
Status	▪ FPI Q4 2020	▪ FPI Q1 2021 ▪ Recruitment completed Q3 2021 ▪ Study met its co-primary endpoints Q1 2022
CT Identifier	NCT04434092	NCT04654468

In collaboration with Chugai
LDH=Lactate Dehydrogenase; ULN=Upper Limit of Normal; IV=Intravenous; SC=Subcutaneous

Crovalimab (RG6107; SKY59)

A humanized monoclonal antibody against complement C5

Indication	Atypical hemolytic uremic syndrome (aHUS) study 1 - adults	Atypical hemolytic uremic syndrome (aHUS) study 2 - paediatrics
Phase/study	**Phase III COMMUTE-a**	**Phase III COMMUTE-p**
# of patients	N=90	N=35
Design	Single-arm study of aHUS patients ▪ **Cohort 1**: not previously treated with C5i ▪ **Cohort 2:** switching from C5i ▪ **Cohort 3:** known C5 polymorphism	Single-arm study of aHUS patients ▪ **Cohort 1:** not previously treated with C5i ▪ **Cohort 2:** switching from C5i ≤18y/o
Primary endpoint	▪ Cohort 1+3: proportion of patients with complete TMA response anytime between baseline and week 25 ▪ Cohort 2: proportion of patients with maintained TMA control from baseline through week 25	▪ Cohort 1: proportion of patients with complete TMA response anytime between baseline and week 25 ▪ Cohort 2: proportion of patients with maintained TMA control from baseline through week 25
Status	▪ FPI Q4 2021	▪ FPI Q4 2021
CT Identifier	NCT04861259	NCT04958265

In collaboration with Chugai

aHUS=Atypical Hemolytic Uremic Syndrome; C5i=C5 inhibitor; TMA=thrombotic microangiopathy

Crovalimab (RG6107; SKY59)

A humanized monoclonal antibody against complement C5

Indication	Sickle cell disease (SCD) acute treatment	Sickle cell disease (SCD) chronic VOC prevention
Phase/study	**Phase Ib CROSSWALK-a**	**Phase IIa CROSSWALK-c**
# of patients	N=30	N=90
Design	**ARM A:** Crovalimab **ARM B:** Placebo	**ARM A:** Crovalimab **ARM B:** Placebo
Primary endpoint	▪ Safety	▪ VOC rate, up to 48 weeks
Status	▪ FPI Q1 2022	▪ FPI Q1 2022
CT Identifier	NCT04912869	NCT05075824

SCD=Sickle Cell Disease; VOC=Vaso-occlusive crises

Crenezumab (RG7412)

Humanized monoclonal antibody targeting all forms of Ab

Indication	Alzheimer's prevention initiative (API) Colombia
Phase/study	**Phase II** Cognition study
# of patients	N=252
Design	▪ **ARM A:** PSEN1 E280A mutation carriers receive crenezumab SC or IV ▪ **ARM B:** PSEN1 E280A mutation carriers receive placebo ▪ **ARM C:** non-mutation carriers receive placebo
Primary endpoint	▪ Change on Alzheimer's Prevention Initiative (API) Composite Cognitive Test total score at 260 weeks treatment ▪ Annualized rate of change in an Episodic Memory Measure: Free and Cued Selective Reminding Task (FCSRT)
Status	▪ FPI Q4 2013 ▪ Recruitment completed Q1 2017 ▪ Study did not meet its co-primary endpoints Q2 2022
CT Identifier	NCT01998841

In collaboration with AC Immune
Ab=amyloid-beta; SC=Subcutaneous; IV=Intravenous

Gantenerumab (RG1450)

Fully human monoclonal antibody binding aggregated forms of Aß

Indication	Prodromal to mild Alzheimer's disease		
Phase/study	**Phase III GRADUATE 1**	**Phase III GRADUATE 2**	**Phase II GRADUATION**
# of patients	N=1,016	N=1,016	N=192
Design	104-week SC treatment period: ▪ **ARM A:** Gantenerumab ▪ **ARM B:** Placebo	104-week SC treatment period: ▪ **ARM A:** Gantenerumab ▪ **ARM B:** Placebo	104-week SC treatment period: gantenerumab SC treatment q1w dosing regimen
Primary endpoint	▪ Change in CDR-SOB at 27 months	▪ Change in CDR-SOB at 27 months	▪ Change from baseline in deposited amyloid (PET centiloid levels)
Status	▪ FPI Q2 2018 ▪ Recruitment completed Q2 2020	▪ FPI Q3 2018 ▪ Recruitment completed Q2 2020	▪ FPI Q4 2020 ▪ Recruitment completed Q3 2021
	▪ BTD granted by FDA Sep 2021		
CT Identifier	NCT03443973	NCT03444870	NCT04592341

In collaboration with MorphoSys AG

Aß=amyloid-beta; CDR-SOB=Clinical Dementia Rating Scale Sum of Boxes; SC=Subcutaneous; BTD=Breakthrough Therapy Designation; PET= positron emission tomography

Gantenerumab (RG1450)

Fully human monoclonal antibody binding aggregated forms of Aß

Indication	Prodromal Alzheimer's disease	Mild Alzheimer's disease	Cognitively unimpaired participants at risk for or at the earliest stages of Alzheimer's disease
Phase/study	**Phase II/III** **SCarlet RoAD[1]**	**Phase III** **Marguerite RoAD[1]**	**Phase III** **SKYLINE[2]**
# of patients	N=799	N=389	N=1200
Design	104-week SC treatment period: ▪ **ARM A:** Gantenerumab (225 mg) ▪ **ARM B:** Gantenerumab (105 mg) ▪ **ARM C:** Placebo	104-week SC treatment period: ▪ **ARM A:** Gantenerumab ▪ **ARM B:** Placebo	▪ **ARM A:** Gantenerumab q1w or q2w (patient preference) ▪ **ARM B:** Placebo
Primary endpoint	▪ Change in CDR-SOB at 2 years ▪ Sub-study: change in brain amyloid by PET at 2 years	▪ Change in ADAS-Cog and CDR-SOB at 2 years (co-primary)	▪ Cognitive composite (PACC5)
Status	▪ Phase I PET data: *Archives of Neurology*, 2012 Feb;69(2):198-207 ▪ Recruitment completed Q4 2013 ▪ Dosing stopped due to futility Q4 2014 ▪ FPI in open label extension study Q4 2015 ▪ Published in *Alzheimers Res Ther* 2017 Dec 8;9(1):95	▪ FPI Q1 2014 ▪ Recruitment stopped Q4 2015 ▪ FPI Q1 2016 for open label extension	▪ FPI Q2 2022
	▪ 36 OLE data published in J Prev Alzheimers Dis 2021;8(1):3-6		
CT Identifier	NCT01224106	NCT02051608	NCT05256134

[1]In collaboration with MorphoSys AG; [2]In collaboration with Banner Alzheimer's Institute

Aß=amyloid-beta; CDR-SOB=Clinical Dementia Rating Scale Sum of Boxes; PET= positron emission tomography; ADAS-cog=Alzheimer's Disease Assessment Scale cognitive subscale; SC=Subcutaneous; OLE=Open Label Extension; PACC5=Preclinical Alzheimer's Cognitive Composite

Tominersen (RG6042, HTT ASO)

Antisense oligonucleotide (ASO) targeting human HTT mRNA

Indication	Huntington's disease	
Phase/study	**Phase I/IIa**	**Phase II OLE**
# of patients	N=46	N=46
Design	▪ Multiple ascending doses of tominersen administered intrathecally to adult patients with early manifest Huntington's Disease	▪ Patients from phase I are enrolled into OLE
Primary endpoint	▪ Safety, tolerability and PK/PD	▪ Longer term safety, tolerability and PK/PD
Status	▪ FPI Q3 2015 ▪ Data presented at CHDI 2018 and AAN 2018 ▪ PRIME designation granted 2018 ▪ Published in *NEJM* 2019; 380:2307-2316	▪ FPI Q1 2018 ▪ PK/PD data presented at AAN 2019 ▪ Update presented at CHDI 2020 ▪ Study completed, patients moved to GEN-EXTEND OLE
CT Identifier	NCT02519036	NCT03342053

In collaboration with Ionis Pharmaceuticals

HTT=Huntingtin; PK/PD=Pharmacokinetics/Pharmacodynamics; PRIME=Priority medicines; OLE=Open Label Extension; AAN=American Academy of Neurology; NEJM=New England Journal of Medicine; CHDI=Huntington's Disease Association of Ireland

Tominersen (RG6042, HTT ASO)

Antisense oligonucleotide (ASO) targeting human HTT mRNA

Indication	Huntington's disease	
Phase/study	**Phase III Generation HD1**	**Phase III GEN-EXTEND**
# of patients	N=791	N=1,050
Design	▪ **ARM A:** Tominersen 120mg q2w ▪ **ARM B:** Tominersen 120mg q4m ▪ **ARM C:** Placebo q2w	OLE study in patients participating in prior Roche and Genentech sponsored studies • **ARM A:** Tominersen 120mg q2w • **ARM B:** Tominersen 120mg q4m
Primary endpoint	▪ cUHDRS globally ▪ TFC USA only	▪ Long term safety, tolerability
Status	▪ FPI Jan 2019 ▪ Q1 2019 protocol modified to allow for bi-monthly vs four-monthly dosing, FPI for new protocol July 2019 ▪ Recruitment completed Q2 2020 ▪ Dosing stopped in Q1 2021 based on IDMC recommendation regarding the potential benefit/risk profile for study participants. No new safety signals identified. ▪ Data presented at EHDN and CHDI 2022	• FPI Q2 2019 ▪ Dosing stopped in Q1 2021
CT Identifier	NCT03761849	NCT03842969

In collaboration with Ionis Pharmaceuticals

cUHDRS=composite Unified Huntington's Disease Rating Scale; TFC=total function capacity; HTT=Huntingtin; OLE=Open Label Extension; IDMC=Independent Data Monitoring Committee; CHDI=Huntington's Disease Association of Ireland; EHDN=European Huntington's Disease Network

Fenebrutinib (RG7845, GCD-0853)

Highly selective and reversible (noncovalent) bruton tyrosine kinase

Indication	Primary progressive multiple sclerosis (PPMS)	Relapsing multiple sclerosis (RMS)	
Phase/study	**Phase III FENtrepid**	**Phase III FENhance 1**	**Phase III FENhance 2**
# of patients	N=946	N=736	N=736
Design	▪ **ARM A:** Fenebrutinib twice daily oral ▪ **ARM B:** Ocrevus 2x300mg IV q24w	▪ **ARM A:** Fenebrutinib twice daily oral ▪ **ARM B:** Teriflunomide once daily oral	▪ **ARM A:** Fenebrutinib twice daily oral ▪ **ARM B:** Teriflunomide once daily oral
Primary endpoint	▪ Time to onset of cCDP12	▪ Time to onset of cCDP12 and annualized relapse rate	▪ Time to onset of cCDP12 and annualized relapse rate
Status	▪ FPI Q4 2020	▪ FPI Q1 2021	▪ FPI Q1 2021
CT Identifier	NCT04544449	NCT04586023	NCT04586010

IV=Intravenous; cCDP12=Composite 12-week confirmed disability progression

Roche Group development pipeline

Marketed products development programmes

Roche Pharma global development programmes

Roche Pharma research and early development (pRED)

Genentech research and early development (gRED)

Spark

pRED oncology development programs -1

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Oncology					
FAP-4-1BBL (RG7827)	Solid tumors	I	~150	FPI Q2 2018 Data presented at ESMO 2020 Recruitment completed Q2 2021	
	3L+ MSS mCRC	Ib	80	FPI Q3 2021 Combination study with cibisatamab	NCT04826003
CD19-4-1BBL (RG6076)	R/R B cell non-Hodgkin's lymphoma	I	362	Part I: FPI Q3 2019 Part II: FPI Q3 2020	NCT04077723
PD1-IL2v (RG6279)	Solid tumors	I	348	Part I: FPI Q2 2020; recruitment completed Q4 2021 Part II: FPI Q1 2022	NCT04303858
cibisatamab (CEA x CD3, RG7802)	CEA-positive solid tumors	Ia	149	FPI Q4 2014 Data presented at ASCO 2017	NCT02324257
		Ib	228	FPI Q1 2016 Data presented at ASCO 2017	NCT02650713
	3L+ MSS mCRC	Ib	46	FPI Q1 2019	NCT03866239
PD1-LAG3 (RG6139)	Solid tumors	I	320	FPI Q4 2019	NCT04140500
	Solid tumors	II	210	FPI Q2 2021 Randomized trial, compared with nivolumab	NCT04785820 TALIOS

pRED oncology development programs -2

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Oncology					
CD25 (RG6292)	Solid tumors	I	110	FPI Q4 2019	NCT04158583
	Advanced and metastatic solid tumors	I	160	Part I: FPI Q1 2021 Part II: FPI Q4 2021	NCT04642365
Anti-GPRC5D (RG6234)	Multiple myeloma	I	240	FPI Q4 2020	NCT04557150
HLA-A2-WT1 x CD3 (RG6007)	AML	I	160	FPI Q4 2020	NCT04580121
FAP-CD40 (RG6189)	Solid tumors	I	280	FPI Q2 2021	NCT04857138
HLA-A2-MAGE-A4 x CD3 (RG6129)	Solid tumors	I	180	FPI Q1 2022	NCT05129280
BRAFi (3) (RG6344)	Solid tumors	I	292	FPI Q1 2022	ISRCTN13713551
CD19xCD28 (RG6333)	R/R B cell non-Hodgkin's lymphoma	I	~200	FPI Q1 2022 Combination study with glofitamab	NCT05219513
EGFRvIIIxCD3 (RG6156)	Glioblastoma	I	~200	FPI Q2 2022	NCT05187624

pRED neuroscience development programs -1

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Neuroscience					
Brain Shuttle-gantenerumab (BS-gantenerumab, RG6102)	Alzheimer's disease	IIa	~120	FPI Q1 2021	NCT04639050
Brain Shuttle-CD20 (BS-CD20, RG6035)	Multiple sclerosis	I	30	FPI Q3 2021	ISRCTN16295177
ralmitaront (partial TAAR1 agonist, RG7906)	Schizophrenia	II	36	FPI Q4 2018; Recruitment completed Q3 2019	
		II	247	FPI Q4 2019	NCT03669640 (TWAIN I)
prasinezumab[1] (anti-αSynuclein, RG7935, PRX002)	Parkinson's disease	II	316	The study did not meet its primary endpoint, but showed a reduced clinical decline of core motor signs (MDS UPDRS partIII). Data presented at MDS & ADPD 2020-22. The Open Label Extension is ongoing.	NCT03100149 (PASADENA)
		IIb	575	FPI Q2 2021	NCT04777331 (PADOVA)
alogabat (GABA-Aa5 PAM, RG7816)	Autism spectrum disorder	II	105	FPI Q1 2021	NCT04299464 (Aurora)
NME (RG7637)	Neurodevelopmental disorders	I	80	FPI Q3 2020	NCT04475848
rugonersen (UBE3A LNA, RG6091)	Angelman syndrome	I	66	FPI Q3 2020	NCT04428281
NME (RG6182)	Neurodegenerative disorder	I	30	FPI Q4 2020	

Partner: [1]Prothena
BS=Brain Shuttle

pRED neuroscience development programs -2

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Neuroscience					
NME (RG6289)	Alzheimer's disease	I	138	FPI Q4 2021	
NME (RG6163)	Psychiatric disorders	I	84	FPI Q1 2022	

Partner: [1]Prothena; BS=Brain Shuttle

pRED immunology and ophthalmology development programs

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Immunology					
selnoflast (NLRP3i, RG6418)	Ulcerative colitis	Ib	18	FPI Q4 2021 Recruitment completed Q2 2022	
	Chronic obstructive pulmonary disease	Ib	102	FPI Q2 2022	
Ophthalmology					
NME (RG6179)[1]	DME	I	90	FPI Q3 2019	DOVETAIL
		II	160	FPI Q4 2021	NCT05151744 (BARDENAS)
		II	320	FPI Q4 2021	NCT05151731 (ALLUVIUM)
VEGF-Ang2 DutaFab (RG6120)	nAMD	I	~50	FPI Q4 2020	NCT04567303
NME (RG7774)	Retinal disease	II	135	FPI Q2 2020	NCT04265261 (CANBERRA)

Partner: [1]Sesen Bio

pRED infectious diseases development programs

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Infectious Diseases					
TLR7 agonist (3) (RG7854)	Chronic hepatitis B	I	150	FPI Q4 2016 Data presented at APASL 2019	NCT02956850
CpAM (RG7907)	Chronic hepatitis B	I/II	192	FPI Q4 2016 Data presented at EASL 2018, 2019 & 2020	NCT02952924
		I	22	FPI Q1 2021 Recruitment completed Q2 2021	NCT04729309
TLR7 agonist (3)/ CpAM/siRNA/ PDL1 LNA (RG7854/RG7907/RG6346/RG6084)	Chronic hepatitis B	II	275	FPI Q3 2020	NCT04225715 (PIRANGA)
PDL1 LNA (RG6084)	Chronic hepatitis B	I	35	FPI Q1 2019 Part Ia: completed Part Ib: initiated	
Abx MCP (RG6006)	A. baumannii infections	I	204	FPI Q4 2020	NCT04605718

Abx MCP=antibiotic macrocyclic peptide

Roche Group development pipeline

Marketed products development programmes

Roche Pharma global development programmes

Roche Pharma research and early development (pRED)

Genentech research and early development (gRED)

Spark

gRED oncology development programs

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Oncology					
KRAS G12C (RG6330)	Metastatic solid tumors with KRAS G12C mutation	I	270	FPI Q3 2020	NCT04449874
cevostamab (anti-FcRH5 x CD3; RG6160)	R/R multiple myeloma	I	300	FPI Q3 2017 Data presented at ASH 2020, ASH 2021	NCT03275103
runimotamab (HER2 x CD3, RG6194)	Metastatic HER2-expressing cancers	I	440	FPI Q2 2018	NCT03448042
NME (RG6286)	Locally advanced or metastatic colorectal cancer	I	67	FPI Q3 2020	NCT04468607
IL15/IL15Ra-Fc (RG6323)[1]	Solid tumors	I/II	250	FPI Q1 2020	NCT04250155
	R/R multiple myeloma	I	60	FPI Q2 2022	NCT05243342
autogene cevumeran (Individualized Neoantigen-Specific Therapy (iNeST); RG6180)[2]	Solid tumors	Ia/IIb	271	FPI Q4 2017 Data presented at AACR 2020 Recruitment completed Q1 2022	NCT03289962
	1L advanced melanoma	II	132	FPI Q1 2019	NCT03815058 (IMcode001)
SHP2i (RG6344)	Solid tumors	Ia	~50	FPI Q1 2020	NCT04252339
belvarafenib (RG6185)[3]	nRASmt CPI-experienced melanoma	Ib	83	FPI Q2 2021	NCT04835805
NME (RG6392)	Oncology	I	60	FPI Q4 2021	ISRCTN92655801

Partner: [1]Xencor, [2]BioNTech, [3]Hanmi

gRED immunology and ophthalmology development programs

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Immunology					
efmarodocokin alfa (IL-22Fc, RG7880)	aGVHD	Ib	18	FPI Q4 2020	NCT04539470
NME (RG6287, GDC-8264)	Inflammatory bowel disease	I	68	FPI Q1 2020 Recruitment completed Q3 2021	EUDRACT2019-002613-19
	Inflammatory diseases	I	16	FPI Q4 2021	
NME (RG6315, MTBT1466A)	Immunologic disorders	I	~24	FPI Q3 2020	
astegolimab (Anti-ST2, (RG6149, AMG 282, MSTT1041A)[1]	Chronic obstructive pulmonary disease	IIb	930	FPI Q4 2021	NCT05037929
NME (RG6341, GDC-6599)	Asthma	Ia/Ib	84	FPI Q4 2021	
Ophthalmology					
galegenimab (HtrA1, RG6147)	Geographic atrophy	II	360	FPI Q2 2019	NCT03972709 (GALLEGO)
NME (RG6312)	Geographic atrophy	Ia	63	FPI Q4 2020	NCT04615325
NME (RG6351)	Retinal disease	I	42-78	FPI Q2 2022	

Partner: [1]Amgen

gRED neuroscience and infectious diseases development programs

Molecule	Indication	Phase	# of patients	Status	CT Identifier
Neuroscience					
semorinemab (RG6100)[1]	Prodromal to mild Alzheimer's disease	II	457	FPI Q4 2017 Primary endpoint not met Q3 2020 Data presented at CTAD 2020	NCT03289143 (TAURIEL)
	Mild-to-moderate Alzheimer's disease	II	272	FPI Q1 2019 One of two co-primary endpoints met Q3 2021 Data presented at CTAD 2021 The Open Label Extension is ongoing	NCT03828747 (LAURIET)
Infectious Diseases					
LepB inhibitor (RG6319)	Complicated urinary tract infection	I	56	FPI Q1 2022	

Partner: [1]AC Immune

Roche Group development pipeline

Marketed products development programmes

Roche Pharma global development programmes

Roche Pharma research and early development (pRED)

Genentech research and early development (gRED)

Spark

Hemophilia A

Unique gene therapy platform

Molecule	SPK-8011 (RG6357)		SPK-8016 (RG6358)
Indication	**Hemophilia A**		**Hemophilia A with inhibitors to Factor VIII**
Phase/study	**Phase I**	**Phase I/II**	**Phase I/II**
# of patients	N=100	N=30	N=30
Design	▪ Long term follow up study of patients who have received SPK-8011 in any prior Spark-sponsored SPK-8011 study	▪ Gene transfer, dose-finding safety, tolerability, and efficacy study of SPK-8011	▪ Gene transfer, dose-finding safety, tolerability, and efficacy study of SPK-8016 in individuals with FVIII inhibitors
Primary endpoint	▪ Safety	▪ Safety and changes from baseline in FVIII activity levels at week 52	▪ Safety; peak and steady state FVIII activity levels at week 52
Status	▪ Ongoing	▪ FPI Q1 2017 ▪ Updated data presented at ISTH 2020 and 2021 ▪ Recruitment completed Q1 2021 ▪ Data published in *NEJM* 2021; 385:1961-1973	▪ FPI Q1 2019
CT Identifier	NCT03432520	NCT03003533	NCT03734588

ISTH=International Society on Thrombosis and Haemostasis; NEJM=New England Journal of Medicine

Pompe disease

Unique gene therapy platform

Molecule	**SPK-3006 (RG6359)**
Indication	**Pompe disease**
Phase/study	**Phase I/II RESOLUTE**
# of patients	N=20
Design	▪ Gene transfer study for late-onset Pompe disease
Primary endpoint	▪ Safety
Status	▪ FPI Q4 2020
CT Identifier	NCT04093349

Doing now what patients need next