
Pipeline summary

Marketed products additional indications

Global Development late-stage trials

pRED (Roche Pharma Research & Early Development)

gRED (Genentech Research & Early Development)

Roche Group YTD Sept 2014 sales

Diagnostics

Foreign exchange rate information

Changes to the development pipeline

Q3 2014 update

New to Phase I	New to Phase II	New to Phase III	New to Registration
<u>3 NMEs added by gRED</u> RG6046 SERD – ER-pos. HER2-neg. mBC (origin Seragon) RG7888 anti-OX40 MAb – solid tumors RG7893 Nav1.7 inhibitor – pain <u>1NME added by pRED</u> RG6061 HIF1 alpha locked nucleic acid (LNA) – solid tumors (origin Santaris) <u>1AI</u> RG7446 PD-L1 MAb + ipilimumab or IFN – solid tumors <u>1AI added following filing go decision</u> RG105 MabThera SC – CLL	<u>1 NME moved from Phase 1</u> RG7697 GIP/GLP-1 dual agonist – T2D	<u>2 NMEs moved from Phase 2</u> RG7853 alectinib – NSCLC RG7417 lampalizumab – geographic atrophy <u>1 AI added to reflect the pending FDA submission</u> RG435 Avastin – glioblastoma 1st line	<u>1NME following EU submission</u> RG7421 cobimetinib + Zelboraf – metastatic melanoma <u>1 AI following EU submission</u> RG1273 Perjeta – HER2-pos. BC neoadjuvant <u>1AI following US submission</u> RG3645 Lucentis – diabetic retinopathy
Removed from Phase I	Removed from Phase II	Removed from Phase III	Removed from Registration
<u>3 NMEs terminated by gRED</u> RG7600 mesothelin ADC – pancreatic cancer RG7636 ETBR ADC – metastatic melanoma RG7845 NME – heme tumors	<u>1 NME terminated by gRED</u> RG7667 CMV MAb – CMV <u>1AI</u> RG3616 Erivedge – AML		<u>1 AI following EU CHMP neg. opinion</u> RG435 Avastin – glioblastoma 1st line <u>1NME following approval in the EU</u> RG7159 Gazyvaro – CLL <u>1 AI following approval in the EU</u> RG1569 Actemra – early RA

Roche Group development pipeline

Phase I (28 NMEs + 10 AIs)

Oncology

RG105	MabThera SC	CLL
RG6016	LSD1 inh	AML
RG6046	SERD	ER+ (HER2-) mBC
RG6061	HIF1 alpha LNA	solid tumors
RG7116	HER3 MAb	solid tumors
RG7304	Raf & MEK dual inh	solid tumors
RG7388	MDM2 ant	solid & hem tumors
RG7446	PD-L1 MAb+Tarceva	NSCLC EGFR+
RG7446	PD-L1 MAb+Zelboraf	m. melanoma
RG7446	PD-L1 MAb+Avastin+chemo	solid tumors
RG7446	PD-L1 MAb+cobimetinib	solid tumors
RG7446	PD-L1 MAb+ipilimum./IFN	solid tumors
RG7446	PD-L1 MAb	solid tumors
RG7450	Steap 1 ADC	prostate ca.
RG7458	MUC16 ADC	ovarian & pancreatic ca.
RG7601	Bcl-2 inh + Gazyva	CLL
RG7601	Bcl-2 inh	heme indications
RG7666	PI3k inh	glioblastoma 2L
RG7741	ChK1 inh	solid tum & lymphoma
RG7775	MDM2 (4) IV prodrug	AML
RG7813	CEA IL2v	solid tumors
RG7841	ADC	solid tumors
RG7842	ERK inh	solid tumors
RG7882	ADC	ovarian ca
RG7888	OX40	solid tumors
CHU	PI3K inh	solid tumors

Other disease areas

RG7624	IL-17 MAb	autoimmune diseases
RG7795	TLR7 agonist	HBV
RG7641	aldosterone synth inh	kidney disease
CHU	URAT 1 inh	gout
RG7203	PDE10A inh	schizophrenia
RG7342	mGlu5 PAM	schizophrenia
RG7410	TAAR1 ago	schizophrenia
RG7893	Nav1.7 inh	pain
RG7800	SMN2 splicer	spinal muscular atrophy
RG7935	a-synuclein MAb	Parkinson's Disease
RG3645	Lucentis sust. deliv.	AMD/RVO/DME
RG7716	VEGF-ANG2 MAb	wAMD

	New Molecular Entity (NME)
	Additional Indication (AI)
	Oncology
	Immunology
	Infectious Diseases
	CardioMetabolism
	Neuroscience
	Ophthalmology
	Other
RG-No	Roche Genentech managed
CHU	Chugai managed

Roche Group development pipeline

Phase II (27 NMEs + 7 AIs)

RG7155	CSF-1R MAb	solid tumors & PVNS
RG7221	Ang2-VEGF MAb	colorectal cancer
RG7321	pictilisib (PI3K inh)	solid tumors
RG7440	ipatasertib (AKT inh)	solid tumors
RG7446	PD-L1 MAb	NSCLC 2 nd /3 rd line
RG7446	PD-L1 MAb + Avastin	RCC
RG7446	PD-L1 MAb	bladder cancer
RG7593	pinatuzumab vedotin (CD22 ADC)	hem tumors
RG7596	polatuzumab vedotin (CD79bADC)	hem tumors
RG7597	HER3/EGFR MAb	m. epithelial tumors
RG7599	lifastuzumab vedotin (NaPi2bADC)	Pt-resist. OC
RG7601	Bcl-2 inh	CLL rel/refract 17pdel
RG7601	Bcl-2 inh	DLBCL
RG7604	taselisib (PI3K inh beta sparing)	solid tumors
RG7686	glypican-3 MAb	liver cancer
RG1569	Actemra	systemic sclerosis
RG3637	lebrikizumab	idiopathic pulmonary fibrosis
RG7449	quilizumab	asthma
CHU	IL-31R MAb	atopic dermatitis
RG7128	mericitabine	HCV
RG7227	danoprevir	HCV
RG7745	Flu A MAb	influenza
RG7790	setrobuvir	HCV
RG7929	LptD antibiotic	antibacterial
RG7697	GIP/GLP-1 dual ago	type 2 diabetes
RG1512	inclacumab	ACS/CVD
RG1577	MAO-B inh	Alzheimer's
RG1578	decoglutran (mGlu2 NAM)	depression
RG1662	GABRA5 NAM	Down Syndrome
RG1678	bitopertin	obsessive compulsive dis.
RG7090	basimglurant (mGlu5 NAM)	TRD
RG7314	V1 receptor antag	autism
RG7412	crenezumab	Alzheimer's
RG6013	FIXa /FX bispecific MAb	hemophilia A

Phase III (9 NMEs + 19 AIs)

RG435 ¹	Avastin	glioblastoma 1 st line
RG435	Avastin	NSCLC adj
RG435 ¹	Avastin	ovarian cancer 1 st line
RG435 ¹	Avastin	rel. ovarian ca. Pt-sensitive
RG1273	Perjeta	HER2+ mBC 2 nd line
RG1273	Perjeta	HER2+ BC adj
RG1273	Perjeta	HER2+ gastric cancer
RG3502	Kadcyla	HER2+ gastric cancer
RG3502	Kadcyla +/- Perjeta	HER2+ mBC 1 st line
RG3502	Kadcyla	HER2+ BC adj
RG3502	Kadcyla + Perjeta	HER2+ BC adj
RG3502	Kadcyla + Perjeta	HER2+ BC neoadj
RG7159	Gazyva (obinutuzumab)	DLBCL
RG7159	Gazyva (obinutuzumab)	iNHL relapsed
RG7159	Gazyva (obinutuzumab)	iNHL front-line
RG7204	Zelboraf	melanoma adj
RG7446	PD-L1 MAb	NSCLC 2 nd line
RG7601	Bcl-2 inh	CLL rel/refract
RG7853	alectinib (ALK inhibitor)	NSCLC
RG1569	Actemra	giant cell arteritis
RG3637	lebrikizumab	severe asthma
RG7413	etrolizumab	ulcerative colitis
CHU	Suvenyl	enthesopathy
CHU	IL-6R MAb	neuromyelitis optica
RG1450	gantenerumab	Alzheimer's
RG1594	ocrelizumab	RMS
RG1594	ocrelizumab	PPMS
RG7417	lampalizumab (factor D)	geo. atrophy

Registration (1 NMEs + 4 AIs)

RG435 ²	Avastin	recurrent cervical cancer
RG435 ³	Avastin	rel. ovarian ca. Pt-resistant
RG1273 ²	Perjeta	HER2+ BC neoadj
RG7421	cobimetinib + Zelboraf	m. melanoma
RG3645 ⁴	Lucentis	diabetic retinopathy

- 1 US only: FDA submission pending
- 2 Approved in US, submitted in EU
- 3 Approved in EU, submitted in US
- 4 Submitted in US

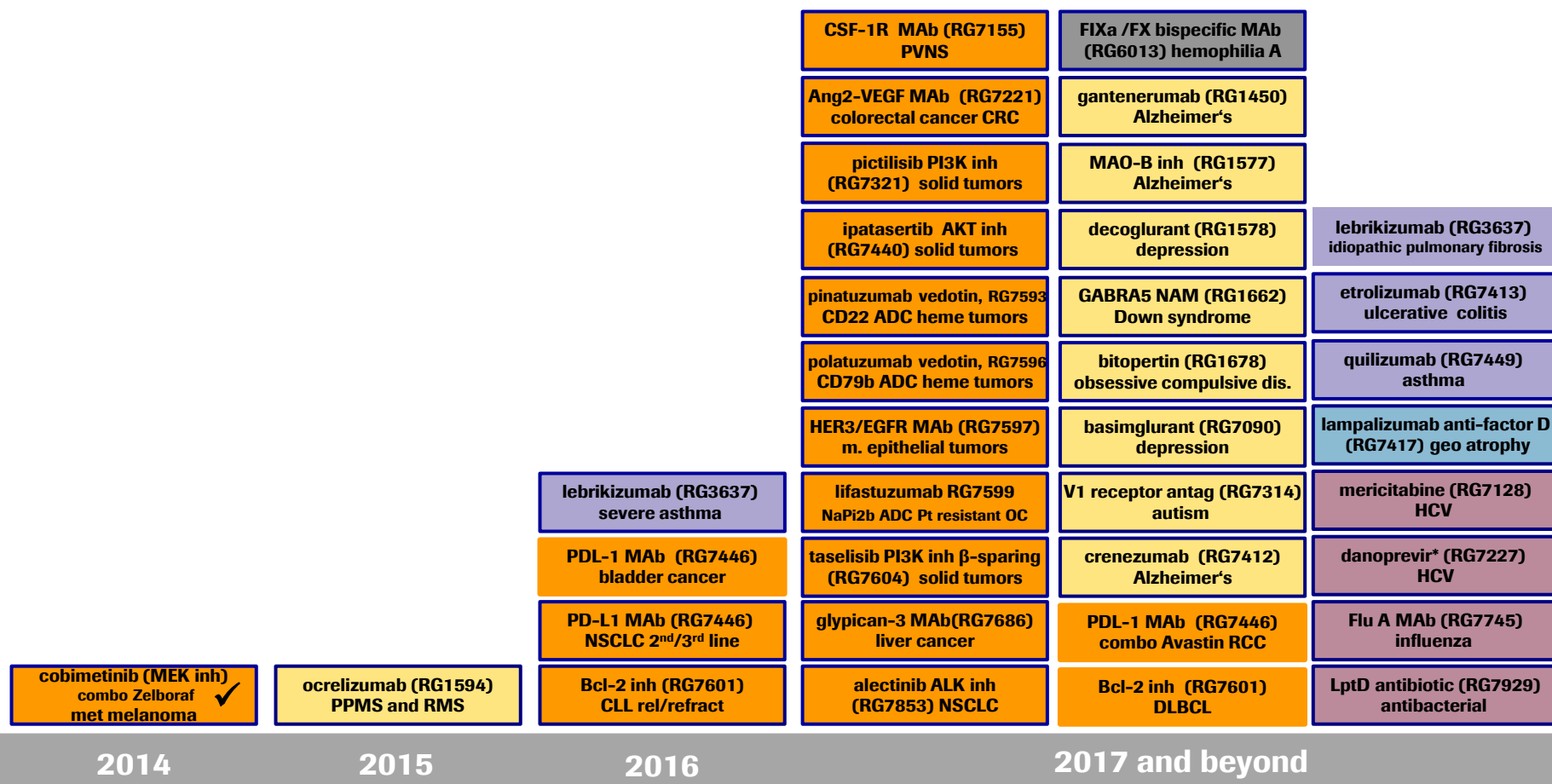
New Molecular Entity (NME)
 Additional Indication (AI)

Oncology
 Immunology
 Infectious Diseases
 CardioMetabolism
 Neuroscience
 Ophthalmology
 Other

RG-No Roche Genentech managed
CHU Chugai managed
RG105 MabThera is branded as Rituxan in US and Japan
RG1569 Actemra is branded as RoActemra in EU
RG7159 Gazyva is branded as Gazyvaro in EU

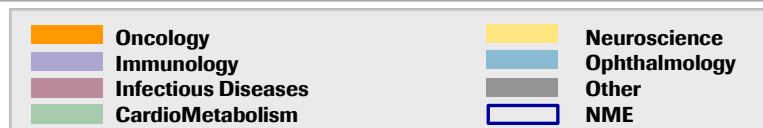
NME submissions and their additional indications

Projects currently in phase 2 and 3



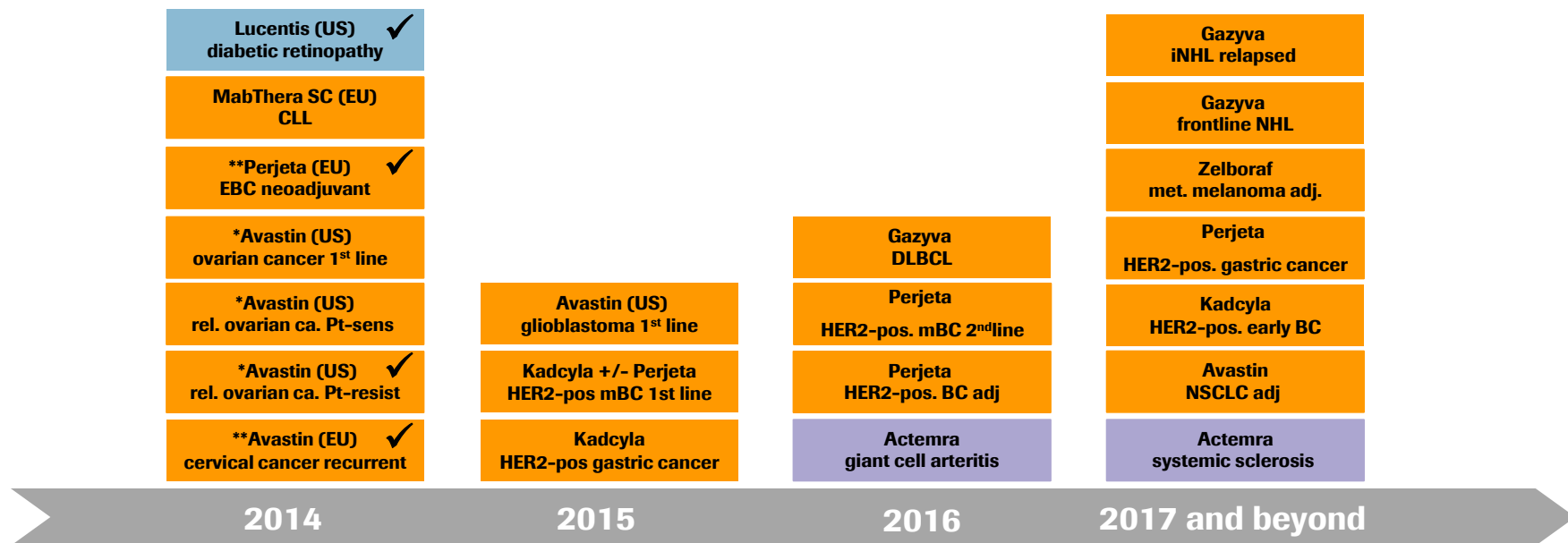
Unless stated otherwise, submissions are planned to occur in US and EU
 * lead market China

Status as of October 16, 2014



Submissions of additional indications for existing products

Projects currently in phase 2 and 3

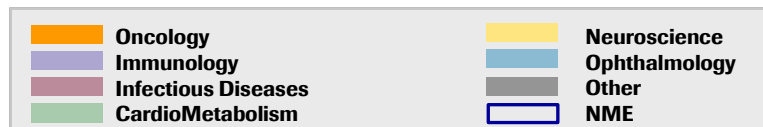


✓ Indicates submission to health authorities has occurred.

* approved in EU; ** approved in US

Unless stated otherwise, submissions are planned to occur in US and EU.

Status as of October 16, 2014



Major granted and pending approvals 2014

Approved

Pending approvals

US

Avastin
cervical cancer
August 2014

Esbriet*
idiopathic pulmonary fibrosis
October 2014

Xolair
chronic idiopathic urticaria
March 2014

Avastin
rel. ovarian ca. Pt-resist
Filed May 2014

Lucentis
diabetic retinopathy
Filed August 2014

EU

MabThera
NHL sc formulation
March 2014

Actemra
RA sc formulation
April 2014

Gazyvaro
CLL
July 2014

Actemra
early RA
September 2014

Avastin
rel. ovarian ca. Pt-resist
August 2014

Esbriet*
idiopathic pulmonary fibrosis
March 2011

Avastin
cervical cancer
Filed April 2014

Perjeta
BC neoadjuvant
Filed September 2014

cobimetinib + Zelboraf
m. melanoma
Filed September 2014

Japan-Chugai

alectinib ALECENSA
ALK-pos rec/adv NSCLC
July 2014

Zelboraf
m. melanoma
Filed April 2014

Status as of October 16, 2014; * Newly acquired asset (InterMune)

Oncology	Neuroscience
Immunology	Ophthalmology
Infectious Diseases	Other
CardioMetabolism	NME

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gRED (Genentech Research & Early Development)

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Avastin

Ovarian cancer clinical development programme

Indication	Front-line metastatic ovarian cancer	
Phase/study	Phase III GOG-0218	Phase III ICON7
# of patients	N=1,873	N=1,528
Design	▪ ARM A: Paclitaxel and carboplatin for 6 cycles plus 5 cycles of concurrent placebo followed by placebo alone for up to 22 cycles (15 months) ▪ ARM B: Paclitaxel and carboplatin for 6 cycles plus 5 cycles of concurrent Avastin followed by placebo alone for up to 22 cycles (15 months) ▪ ARM C: Paclitaxel and carboplatin for 6 cycles plus 5 cycles of concurrent Avastin followed by Avastin alone for up to 22 cycles (15 months)	▪ ARM A: Paclitaxel and carboplatin for 6 cycles ▪ ARM B: Paclitaxel and carboplatin plus concurrent Avastin for 6 cycles followed by Avastin alone for up to 18 cycles (12 months)
Avastin dose	▪ 15 mg/kg q3 weeks	▪ 7.5 mg/kg q3 weeks
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival
Status	▪ Study met its primary endpoint in Q1 2010 ▪ Data presented at ASCO 2010 and 2011 ▪ Results: NEJM 2011 Dec 29;365(26):2484-96	▪ Study met its primary endpoint Q3 2010 ▪ Data presented at ESMO 2010 and ASCO 2011 ▪ Results: NEJM 2011 Dec 29;365(26):2473-83 ▪ OS data presented at ECC 2013
	▪ EMA approval Q4 2011 ▪ Re-evaluate FDA submission in 2014	

Avastin

Ovarian cancer clinical development programme

Indication	Relapsed Platinum-sensitive ovarian cancer	Relapsed Platinum-resistant ovarian cancer
Phase/study	Phase III OCEANS	Phase III AURELIA
# of patients	N=484	N=361
Design	▪ ARM A: Carboplatin, gemcitabine, and concurrent placebo for 6-10 cycles, followed by placebo alone until disease progression ▪ ARM B: Carboplatin, gemcitabine, and concurrent Avastin for 6-10 cycles, followed by Avastin alone until disease progression.	▪ ARM A: Paclitaxel, topotecan or liposomal doxorubicin ▪ ARM B: Paclitaxel, topotecan or liposomal doxorubicin plus Avastin
Avastin dose	▪ 15 mg/kg q3 weeks	▪ 10 mg/kg q2 weeks or 15 mg/kg q3 weeks
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival
Status	▪ Study met its primary endpoint Q1 2011 ▪ Data presented at ASCO 2011 ▪ EMA approval received Q4 2012 ▪ Re-evaluate FDA submission in 2014	▪ Study met its primary endpoint Q2 2012 ▪ Data presented at ASCO 2012 ▪ Results published in JCO 2014 May 1;32(13):1309-16 ▪ Approved in EU Q3 2014 ▪ Filed with the FDA Q2 2014 ▪ FDA priority review granted Q3 2014

Avastin

Cervical and brain cancer clinical development programmes

Indication	Stage IVB, recurrent or persistent cervical cancer	Newly diagnosed glioblastoma
Phase/study	Phase III GOG-240	Phase III AVAglio
# of patients	N=452	N=920
Design	▪ ARM A: Paclitaxel, cisplatin ▪ ARM B: Paclitaxel, cisplatin plus Avastin ▪ ARM C: Paclitaxel, topotecan ▪ ARM D: Paclitaxel, topotecan plus Avastin	▪ ARM A: Concurrent radiation and temozolomide plus placebo; followed by maintenance TMZ plus placebo for 6 cycles; then placebo until disease progression ▪ ARM B: Concurrent radiation and TMZ plus Avastin; followed by maintenance TMZ plus Avastin for 6 cycles; then Avastin (15mg/kg q3 weeks) monotherapy until disease progression
Avastin dose	▪ 15 mg/kg q3 weeks	▪ 10 mg/kg q2 weeks or 15 mg/kg q3 weeks
Primary endpoint	▪ Overall survival	▪ Progression-free survival ▪ Overall survival
Status	▪ Study met its primary endpoint Q1 2013 ▪ Results published in NEJM Feb. 2014; 370(8):734-43 ▪ Filed globally Q2 2014 ▪ FDA approval granted Q3 2014	▪ Co-primary endpoint of PFS met Q3 2012 ▪ Overall survival data presented at ASCO 2013 ▪ Filed in EU Q1 2013 ▪ Negative CHMP opinion Q3 2014

Avastin

Lung and breast cancer development programmes

Indication	Adjuvant lung cancer	First-line HER2-negative metastatic breast cancer
Phase/study	Phase III ECOG 1505	Phase III MERiDiAN
# of patients	N=1,500	N=480
Design	▪ ARM A: Cisplatin plus vinorelbine, docetaxel, gemcitabine or pemetrexed ▪ ARM B: Cisplatin plus vinorelbine, docetaxel, gemcitabine or pemetrexed plus Avastin up to 12 months	▪ ARM A: Paclitaxel + Avastin ▪ ARM B: Paclitaxel + Placebo
Avastin dose	▪ 15 mg/kg q3 weeks	▪ 10 mg/kg q2 weeks
Primary endpoint	▪ Overall survival	▪ PFS in ITT ▪ PFS in patients with high plasma VEGF-A
Status	▪ Recruitment completed Q4 2013 ▪ Expect data in 2016	▪ Recruitment completed ▪ Expect data in 2015

Erivedge

A novel small molecule inhibitor of the hedgehog signaling pathway

Indication	Locally advanced or metastatic basal cell carcinoma	Idiopathic pulmonary fibrosis
Phase/study	Phase II STEVIE	Phase II
# of patients	N=1,200	N=129
Design	▪ Single ARM: 150 mg Erivedge orally once daily	▪ ARM A: Erivedge 150mg daily ▪ ARM B: placebo
Primary endpoint	▪ Safety: Incidence of adverse events	▪ Change in FVC
Status	▪ FPI Q2 2011	▪ FPI pending in anticipation of trial design amendment to incorporate new standard of care pirfenidone.

Gazyva/Gazyvaro

Type II, glycoengineered anti-CD20 monoclonal antibody

Indication	Front-line chronic lymphocytic leukaemia Patients with comorbidities	Previously untreated or relapsed/refractory chronic lymphocytic CLL
Phase/study	Phase III CLL11	Phase III GREEN
# of patients	N=781	N=800
Design	▪ ARM A: Gazyva 1000mg iv plus chlorambucil ▪ ARM B: MabThera/Rituxan plus chlorambucil ▪ ARM C: Chlorambucil alone	▪ Single-arm cohort study: Gazyva alone or in combination with different chemotherapy regimens (FC, Bendamustine or Clb), investigation of different strategies to reduce IRRs
Primary endpoint	▪ Progression-free survival	▪ Safety in combination with different chemotherapy regimens
Status	▪ Filed globally Q2 2013 ▪ FDA approval granted Q4 2013 ▪ Positive CHMP opinion Q2 2014 ▪ Full data published NEJM Mar 2014; 370(12):1101-10	▪ FPI Q4 2013

Gazyva/Gazyvaro

Type II, glycoengineered anti-CD20 monoclonal antibody

Indication	Diffuse large B-cell lymphoma (DLBCL)	Indolent non-Hodgkin's lymphoma MabThera/Rituxan refractory	Front-line indolent non-Hodgkin's lymphoma
Phase/study	Phase III GOYA	Phase III GADOLIN Induction and maintenance study	Phase III GALLIUM Induction and maintenance study
# of patients	N=1,400	N=410	N=1,400
Design	▪ ARM A: Gazyva 1000mg iv plus CHOP ▪ ARM B: MabThera/Rituxan plus CHOP	▪ ARM A: Gazyva 1000mg iv plus bendamustine followed by Gazyva maintenance ▪ ARM B: bendamustine	▪ ARM A: Gazyva 1000mg iv plus chemotherapy followed by Gazyva maintenance ▪ ARM B: MabThera/Rituxan plus chemotherapy followed by MabThera/Rituxan maintenance ▪ Chemotherapy: ▪ For follicular lymphoma: CHOP, CVP or bendamustine ▪ For non-follicular lymphoma: physician's choice
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival	▪ Progression-free survival
Status	▪ Recruitment completed Q2 2014 ▪ Expect data in 2015	▪ FPI Q2 2010 ▪ Expect data in 2017	▪ Recruitment completed ▪ Expect data in 2017

Kadcyla

Evaluating new treatment options in HER2-positive early breast cancer

Indication	HER2-positive neoadjuvant breast cancer	HER2-positive early breast cancer high-risk patients	Operable HER2-positive early breast cancer
Phase/study	Phase III KRISTINE	Phase III KATHERINE	Phase III KAITLIN
# of patients	N=432	N=1,484	N=2,500
Design	Before surgery patients will receive 6 cycles of: ▪ ARM A: Herceptin plus Perjeta plus docetaxel plus carboplatin ▪ ARM B: Kadcyla plus Perjeta After surgery patients will receive: ▪ ARM A: Herceptin plus Perjeta ▪ ARM B: Kadcyla plus Perjeta	▪ ARM A: Kadcyla 3.6mg/kg q3w ▪ ARM B: Herceptin	Following surgery and anthracycline-based therapy: ▪ ARM A: Herceptin 6mg/kg q3w plus Perjeta 420 mg/kg q3w plus taxane ▪ ARM B: Kadcyla 3.6mg/kg q3w plus Perjeta 420mg/kg q3w
Primary endpoint	▪ Pathologic Complete Response (pCR)	▪ Invasive disease-free survival (IDFS)	▪ Invasive disease-free survival (IDFS)
Status	▪ FPI Q2 2014	▪ FPI Q1 2013	▪ FPI Q1 2014

Kadcyla

Evaluating new treatment options in HER2-positive breast and gastric cancer

Indication	Previously untreated HER2 pos. metastatic breast cancer	Previously Treated Locally Advanced Or Metastatic Her2- Positive Gastric Cancer
Phase/study	Phase III MARIANNE	Phase II/III GATSBY
# of patients	N=1,092	N=412
Design	▪ ARM A: Herceptin plus taxane ▪ ARM B: Kadcyla 3.6mg/kg q3w plus Perjeta ▪ ARM C: Kadcyla 3.6 mg/kg q3w plus placebo	▪ ARM A: Kadcyla 3.6mg/kg q3w ▪ ARM B: Kadcyla 2.4mg/kg weekly ▪ ARM C: Docetaxel or paclitaxel
Primary endpoint	▪ Progression-free survival assessed by IRF	▪ Phase II: Dose-finding ▪ Phase III: Overall survival
Status	▪ Recruitment completed Q2 2012 ▪ Expect data in 2014	▪ FPI Q3 2012

MabThera/Rituxan

Oncology development programme

Indication	Previously untreated chronic lymphocytic leukemia
Phase/study	Phase Ib SAWYER Subcutaneous study <i>Study being conducted ex-US</i>
# of patients	N=225
Design	- Two-stage design: - Stage 1 (dose-finding, N=55) - Stage 2 (N=170): CLL dose confirmation: ARM A: MabThera iv plus chemotherapy (fludarabine and cyclophosphamide) ARM B: MabThera 1600mg sc plus chemotherapy (fludarabine and cyclophosphamide)
Primary endpoint	- Part 1: PK (dose selection) - Part 2: PK of MabThera iv versus MabThera sc (arm A vs arm B)
Status	- FPI (stage 2) Q3 2012 - Stage 1 data presented at ASH 2012

Perjeta

First in a new class of HER dimerization inhibitors

Indication	Neoadjuvant HER2-positive breast cancer		Adjuvant HER2-positive breast cancer
Phase/ study	Phase II NEOSPHERE	Phase II TRYPHAENA	Phase III APHINITY
# of patients	N=417	N=225	N=4,803
Design	▪ ARM A: Herceptin plus docetaxel ▪ ARM B: Perjeta (840mg loading, 420mg q3w) plus Herceptin and docetaxel ▪ ARM C: Perjeta plus Herceptin ▪ ARM D: Perjeta plus docetaxel	▪ ARM A: FEC followed by Taxane with Herceptin and pertuzumab (H+P given concurrently) ▪ ARM B: FEC followed by Taxane with Herceptin + pertuzumab (H+P given sequentially) ▪ ARM C: TCH + pertuzumab (H+P given concurrently)	▪ ARM A: Perjeta (840mg loading, 420 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	▪ Pathologic complete response (pCR)	▪ Safety	▪ Invasive disease-free survival (IDFS)
Status	▪ Positive data presented at SABCS 2010 ▪ Biomarker data presented SABCS 2011	▪ Positive safety and efficacy data presented at SABCS 2011	▪ Recruitment completed Q3 2013 ▪ Expect data in 2016
	▪ Filed in US Q2 2013 ▪ FDA approval granted Q3 2013 ▪ Filed in EU Q3 2014		

Perjeta

First in a new class of HER dimerization inhibitors

Indication	Second-line HER2-positive metastatic breast cancer	Advanced HER2-positive gastric cancer	Neoadjuvant/adjuvant HER2-positive breast cancer
Phase/study	Phase III PHEREXA	Phase III JACOB	Phase II BERENICE
# of patients	N=450	N=780	N=400
Design	▪ ARM A: Herceptin plus Xeloda ▪ ARM B: Perjeta plus Herceptin and Xeloda	▪ ARM A: Perjeta (840mg loading, 420mg q3w) plus Herceptin and chemotherapy ▪ ARM B: Placebo plus Herceptin and chemotherapy	Neoadjuvant treatment: ▪ ARM A: ddAC q2w x4 cycles followed by weekly paclitaxel for 12 weeks, with P+H x4 cycles ▪ ARM B: FEC+P+H x4 cycles followed by docetaxel+P+H x4 cycles Adjuvant treatment: ▪ P+H q3w to complete 1 year of HER2 therapy ▪ Hormonal and radiation therapy as indicated
Primary endpoint	▪ Progression-free survival	▪ Overall survival	▪ Safety
Status	▪ Recruitment completed Q3 2013 ▪ Expect data in 2015	▪ FPI Q2 2013	▪ FPI Q3 2014

Zelboraf

A selective novel small molecule that inhibits mutant BRAF

Indication	Adjuvant therapy in patients with resected cutaneous BRAF mutation positive melanoma
Phase/study	Phase III BRIM8
# of patients	N=725
Design	▪ 52-week treatment ▪ ARM A: Zelboraf 960mg bid ▪ ARM B: Placebo
Primary endpoint	▪ Disease-free survival
Status	▪ FPI Q3 2012

Actemra/RoActemra

Interleukin 6 receptor inhibitor

Indication	Early moderate-to-severe rheumatoid arthritis	Systemic sclerosis
Phase/study	Phase III FUNCTION	Phase II faSScinat Proof-of-concept study
# of patients	N=1,162	N=86
Design	104 week treatment ARM A: Actemra IV 8 mg/kg q4w plus placebo MTX ARM B: Actemra IV 8 mg/kg q4w plus MTX ARM C: Actemra IV 4 mg/kg q4w plus MTX ARM D: MTX alone	- Blinded 48-week treatment with weekly dosing: ARM A: Actemra SC 162mg ARM B: Placebo SC - Open-label weekly dosing at weeks 49 to 96: - Actemra SC 162mg
Primary endpoint	DAS28 remission at 24 weeks, 1 year and 2 years	- Change in modified Rodnan skin score (mRSS) at week 24 - Safety
Status	- Primary endpoint met Q3 2012 - Data presented at EULAR 2013 - Approved in EU Q3 2014	- Primary endpoint not met, trend with improved efficacy with longer treatment observed - Adboard to advise on phase 3 held in June 2014 - Study is ongoing in a blinded manner to week 48

Actemra/RoActemra

Interleukin 6 receptor inhibitor

Indication	Giant Cell Arteritis
Phase/study	Phase III GiACTA
# of patients	N=250
Design	- Part 1: 52-week blinded period ARM A: Actemra SC 162mg qw + 26 weeks prednisone taper ARM B: Actemra SC 162mg q2w + 26 weeks prednisone taper ARM C: Placebo+ 26 weeks prednisone taper ARM D: Placebo+ 52 weeks prednisone taper - Part II: 104-week open label extension – patients in remission followed off of the study drug; Patients with active disease receive open label Actemra SC 162mg qw
Primary endpoint	Proportion of patients in sustained remission at week 52
Status	FPI Q3 2013

Pipeline summary

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gRED (Genentech Research & Early Development)

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Alectinib (ALK inhibitor, RG7853, AF802)

New CNS-active inhibitor of anaplastic lymphoma kinase

Indication	ALK-positive crizotinib-naïve advanced NSCLC	ALK-positive advanced NSCLC after progression on crizotinib treatment	Treatment naïve ALK-positive advanced NSCLC
Phase/study	Phase I/II	Phase I/II ACCALIA	Phase III ALEX
# of patients	N=70	N=270	N=286
Design	▪ Part 1: Dose escalation monotherapy ▪ Part 2: Monotherapy, dose selected based on the results of Part 1	▪ Part 1: Dose escalation monotherapy ▪ Part 2: Monotherapy, dose selected based on the results of Part 1	▪ ARM A: alectinib 600mg BID ▪ ARM B: crizotinib 250mg BID
Primary endpoint	▪ Safety and efficacy	▪ Safety and efficacy	▪ Progression-free survival
Status	▪ Study in crizotinib-naïve patients in Japan completed; crizotinib-failure patients in US ongoing ▪ Data presented at ECC 2013 ▪ Japan study results: Lancet Oncology 2013 Jun;14(7):590-8 ▪ Approved in Japan with brand name Alecensa July 2014	▪ Phase II FPI Q2 2013 ▪ Phase I full cohort including CNS data published in Lancet Oncology 2014, Sept.15(10):1119-28	▪ FPI Q3 2014
▪ Breakthrough therapy designation granted by the FDA June 2013			

Anti-PDL1 (MPDL3280A, RG7446)

Novel approach in cancer immunotherapy

Indication	Metastatic NSCLC 2 nd line	Locally advanced or metastatic NSCLC PD-L1 positive	Locally advanced or metastatic NSCLC PD-L1 positive	Locally advanced or metastatic NSCLC (2 nd /3 rd line)	Non-small cell lung cancer
Phase/study	Phase III OAK	Phase II FIR	Phase II BIRCH	Phase II POPLAR	Phase I
# of patients	N=850	N=130	N=635	N=287	N=32
Design	▪ RG7446 1200mg q3w ▪ docetaxel	▪ Single arm study ▪ 1200mg of RG7446 q3w for maximum of 16 cycles	▪ Single arm study ▪ 1200mg of RG7446 q3w for maximum of 16 cycles	▪ ARM A: RG7446 1200mg IV q3w, up to 16 cycles ▪ ARM A: Docetaxel IV q3w	▪ RG7446 plus Tarceva¹
Primary endpoint	▪ Overall survival	▪ Overall response rate	▪ Objective response rate	▪ Overall survival	▪ Safety
Status	▪ FPI Q1 2014	▪ Recruitment completed Q2 2014	▪ FPI Q1 2014	▪ Recruitment completed Q2 2014	▪ FPI Q1 2014

¹Tarceva is a registered trademark of OSI Pharmaceuticals, LLC, a subsidiary of Astellas US, LLC;

Anti-PDL1 (MPDL3280A, RG7446)

Novel approach in cancer immunotherapy

Indication	Untreated advanced renal cell carcinoma	Locally advanced or metastatic urothelial bladder cancer	Solid tumors	Locally advanced or metastatic solid tumors
Phase/study	Phase II	Phase II	Phase I	Phase I
# of patients	N=150	N=330	N=154	N=200
Design	▪ ARM A: RG7446 plus Avastin ▪ ARM B: RG7446; following PD: RG7446 plus Avastin ▪ ARM C: sunitinib; following PD: RG7446 plus Avastin	▪ Cohort 1: Treatment-naïve and cisplatin-ineligible patients ▪ Cohort 2: Patients with disease progression following or during platinum-containing treatment	▪ ARM A: RG7446 + Avastin ▪ ARM B: RG7446 + Avastin + FOLFOX ▪ ARM C: RG7446 + Avastin + carboplatin+paclitaxel ▪ ARM D: RG7446 + Avastin + carboplatin+ pemetrexed ▪ ARM E: RG7446 + Avastin + carboplatin+ nab-paclitaxel	▪ ARM A: RG7446 plus ipilimumab ▪ ARM B: RG7446 plus interferon alpha-2b
Primary endpoint	▪ Progression free survival	▪ Objective response rate	▪ Safety/PK	▪ Safety
Status	▪ FPI Q1 2014	▪ FPI Q2 2014	▪ FPI Q2 2012	▪ FPI Q3 2014

Anti-PDL1 (MPDL3280A, RG7446)

Novel approach in cancer immunotherapy

Indication	Previously untreated metastatic melanoma BRAF mutation positive	Locally advanced or metastatic tumors	Relapsed/Refractory follicular lymphoma and DLBCL	Solid tumors
Phase/study	Phase I	Phase I	Phase I	Phase I
# of patients	N=44	N=90	N=52	N=344
Design	Three-arm study with different doses of RG7446-Zelboraf¹ combination	ARM A: Dose-finding – RG7446 plus cobimetinib² ARM B: Dose-expansion – RG7446 plus cobimetinib	Stage 1: safety evaluation - RG7446 plus Gazyva Stage 2: expansion - RG7446 plus Gazyva	Dose escalation study
Primary endpoint	Safety/PK	Safety	Safety	Safety/PK
Status	FPI Q4 2012	FPI Q4 2013	Expect FPI Q4 2014	- FPI Q2 2011 - Initial efficacy data presented at ASCO 2013 - Updated data presented at ECC 2013 - Data from bladder cohort presented at ASCO 2014

¹Zelboraf in collaboration with Plexxikon, a member of Daiichi Sankyo Group;

²Cobimetinib in collaboration with Exelixis

Bcl-2 inhibitor (RG7601, ABT-199/GDC-0199)

Novel small molecule Bcl-2 selective inhibitor

Indication	Relapsed or Refractory CLL	Relapsed CLL and SLL	Relapsed/Refractory or previously untreated CLL	Relapsed/Refractory or previously untreated CLL
Phase/study	Phase III MURANO	Phase Ib	Phase I	Phase I
# of patients	N=370	N=50	N=70	N=74
Design	▪ ARM A: RG7601 plus Rituxan ▪ ARM B: Rituxan plus bendamustine	▪ Dose-escalation study in combination with MabThera/Rituxan	▪ RG7601 in combination with MabThera/Rituxan and bendamustine	▪ RG7601 in combination with Gazyva
Primary endpoint	▪ Safety/MTD	▪ Safety/MTD	▪ Safety/MTD	▪ Safety/MTD
Status	▪ FPI Q1 2014	▪ FPI Q3 2012 ▪ Data presented at ASCO 2014	▪ FPI Q2 2013	▪ FPI Q1 2014

Joint project with AbbVie in collaboration with WEHI (The Walter and Eliza Hall Institute)

CLL=Chronic Lymphocytic Leukemia; NHL=Non-Hodgkin's Lymphoma; SLL=Small Lymphocytic Lymphoma

ASCO=American Society of Clinical Oncology

Bcl-2 inhibitor (RG7601, ABT-199/GDC-0199)

Novel small molecule Bcl-2 selective inhibitor

Indication	Relapsed/Refractory CLL with 17p deletion	CLL patients with relapse or refractory to B-cell receptor signaling pathway inhibitor therapy	Relapsed or Refractory follicular non-Hodgkin's lymphoma
Phase/study	Phase II	Phase II	Phase II
# of patients	N=100	N=40	N=156
Design	Single-agent RG7601	- RG7601 after ibrutinib therapy - RG7601 after idelalisib therapy	ARM A: RG7601 plus Rituxan ARM B: RG7601 plus Rituxan plus bendamustine ARM C: Rituxan plus bendamustine
Primary endpoint	Safety/MTD	Overall response rate	Overall response rate
Status	Recruitment completed Q2 2014	FPI Q3 2014	Expect FPI Q4 2014

Bcl-2 inhibitor (RG7601, ABT-199/GDC-0199)

Novel small molecule Bcl-2 selective inhibitor

Indication	Front-line DLBCL	Relapsed or Refractory NHL	Relapsed/Refractory CLL and NHL
Phase/study	Phase I/II	Phase I	Phase I
# of patients	N=230	N=40	N=121
Design	Dose finding: ▪ ARM A: RG7601+R-CHOP ▪ ARM B: RG7601+G-CHOP Expansion: ▪ RG7601+R(or G)-CHOP	▪ Dose escalation of RG7601 in combination with Rituxan and bendamustine	▪ Dose-escalation study
Primary endpoint	▪ Safety and efficacy	▪ Safety/MTD	▪ Safety/PK/Response rate
Status	▪ FPI Q2 2014	▪ FPI Q2 2012 ▪ Study resumed Q3 2013	▪ FPI Q2 2011 ▪ CLL and NHL data presented at ASCO 2013 ▪ Updated CLL and SLL data presented at ASH 2013 ▪ Updated CLL, SLL and NHL (DLBCL and FL) data presented at ASCO 2014

Joint project with AbbVie in collaboration with WEHI (The Walter and Eliza Hall Institute)

CLL=Chronic Lymphocytic Leukemia; NHL=Non-Hodgkin's Lymphoma; SLL=Small Lymphocytic Lymphoma

ASH=American Society of Hematology; ASCO=American Society of Clinical Oncology

Bcl-2 inhibitor (RG7601, ABT-199/GDC-0199)

Novel small molecule Bcl-2 selective inhibitor

Indication	Relapsed/Refractory multiple myeloma	Relapsed/Refractory multiple myeloma
Phase/study	Phase I	Phase I
# of patients	N=30	N=30
Design	- Patients receiving Bortezomib and Dexamethasone as standard therapy: - Dose escalation cohort: RG7601+bortezomib+dexamethasone - Safety expansion cohort: RG7601+bortezomib+dexamethasone	- Dose escalation cohort - Safety expansion cohort
Primary endpoint	Safety/MTD	Safety/MTD
Status	FPI Q4 2012	FPI Q4 2012

Bcl-2 inhibitor (RG7601, ABT-199/GDC-0199)

Novel small molecule Bcl-2 selective inhibitor

Indication	Acute myelogenous leukemia (AML)	
Phase/study	Phase I	Phase Ib
# of patients	N=32	N=89
Design	▪ Dose escalation of RG7601	▪ RG7601 (dose escalation) +decitabine ▪ RG7601 (dose escalation) +azacitidine
Primary endpoint	▪ Overall response rate	▪ Safety
Status	▪ Recruitment completed	▪ FPI Q4 2014

Cobimetinib (RG7421, GDC-0973)

Selective small molecule inhibitor of mitogen-activated protein kinase kinase

Indication	Previously untreated metastatic melanoma BRAF mutation positive	Locally advanced or metastatic tumors	Locally advanced or metastatic tumors with mutant KRAS
Phase/study	Phase III coBRIM	Phase I	Phase I
# of patients	N=495	N=90	N=50
Design	▪ ARM A: Zelboraf¹ plus cobimetinib ▪ ARM B: Zelboraf¹ plus placebo	▪ ARM A: Dose-finding - cobimetinib plus RG7446 (anti-PDL1) ▪ ARM B: Dose-expansion - cobimetinib plus RG7446 (anti-PDL1)	▪ Dose finding of cobimetinib plus RG7597 (anti-HER3/EGFR DAF)
Primary endpoint	▪ Progression-free survival	▪ Safety	▪ Safety
Status	▪ Enrollment completed Q1 2014 ▪ Primary endpoint met July 2014 ▪ Data presented at ESMO 2014 ▪ Results published in NEJM 2014 Sept. 29 ▪ Filed in EU Q3 2014 ▪ US filing expected Q4 2014	▪ FPI Q4 2013	▪ FPI Q4 2013

In collaboration with Exelixis

¹Zelboraf In collaboration with Plexxikon, a member of Daiichi Sankyo Group;

ESMO=European Society for Medical Oncology; NEJM=New England Journal of Medicine

Pictilisib (RG7321, GDC-0941)

Pan-PI3 kinase inhibitor with potential activity in multiple cancers

Indication	2L ER-positive metastatic breast cancer	Previously untreated advanced or recurrent NSCLC	Locally recurrent or metastatic HER2-negative HR-positive breast cancer
Phase	Phase II FERGI	Phase II FIGARO	Phase II PEGGY
# of patients	N=340	N=302	N=180
Design	▪ ARM A: pictilisib plus hormonal therapy ▪ ARM B: apitolisib plus hormonal therapy (ARM B discontinued) ▪ ARM C: Hormonal therapy + placebo	▪ ARM A: Pictilisib + carboplatin + paclitaxel ▪ ARM B: Placebo + carboplatin + paclitaxel ▪ ARM C: Pictilisib+ carboplatin + paclitaxel + bevacizumab ▪ ARM D: Pictilisib+ carboplatin + paclitaxel + bevacizumab	▪ ARM A: Pictilisib+ paclitaxel ▪ ARM B: Placebo + paclitaxel
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival	▪ Progression-free survival
Status	▪ Recruitment completed Q1 2014	▪ FPI Q1 2012	▪ Recruitment completed Q2 2014

Polatuzumab vedotin (RG7596)

Antibody drug conjugate targeting CD79b for the treatment of B-cell malignancies

Indication	Non-Hodgkin's lymphoma	Non-Hodgkin's lymphoma	Relapsed or Refractory follicular lymphoma and DLBCL
Phase	Phase II ROMULUS	Phase Ib	Phase Ib/II
# of patients	N=120	N=90	N=224
Design	▪ ARM A: RG7593 plus Rituxan ▪ ARM B: RG7596 plus Rituxan	▪ Dose escalation study in combination with Rituxan and chemotherapy	▪ Plb: dose escalation ▪ P2: RG7596+ BR vs. BR ▪ P2 expansion: RG7596+Gazyva non-randomised
Primary endpoint	▪ Safety and anti-tumor activity	▪ Safety	▪ Safety
Status	▪ Recruitment completed Q1 2014 ▪ Initial data presented at ASCO 2014	▪ FPI Q4 2013	▪ Expect FPI Q4 2014

Taselisib(RG7604, GDC-0032)

Beta isoform sparing PI3 kinase inhibitor targeting commonly mutated oncogene

Indication	Solid tumors and HER2-negative HR-positive breast cancer	HER2-negative locally recurrent or metastatic breast cancer	PI3KCAmut-pos. 2L squamous NSCLC Lung Master Protocol
Phase	Phase I/II	Phase I	Phase II Lung-MAP
# of patients	N=320	N=65	N=120
Design	▪ Phase I ▪ RG7604 ▪ RG7604 plus letrozole or fulvestrant ▪ Phase II ▪ RG7604 (multiple doses) plus letrozole or fulvestrant	▪ RG7604 plus docetaxel ▪ RG7604 plus paclitaxel	▪ RG7604 vs. chemo
Primary endpoint	▪ Safety/PK/efficacy	▪ Safety	▪ Progression-free survival
Status	▪ Recruitment completed Q2 2014 ▪ Data presented at SABCS 2013 ▪ Biomarker data presented at AACR 2014	▪ FPI Q2 2013	▪ FPI Q2 2014

Bitopertin (GlyT-1, RG1678)

A small molecule first-in-class glycine reuptake inhibitor (GRI)

Indication	Obsessive-compulsive disorder
Phase/study	Phase II SKYLYTE
# of patients	N=99
Design	▪ 16-week treatment period ▪ Background therapy of selective serotonin reuptake inhibitors (SSRI) ▪ARM A: bitopertin daily (30 mg) ▪ARM B: bitopertin daily (10 mg) ▪ARM C: Placebo
Primary endpoint	▪ Change in total score on Yale-Brown Obsessive Compulsive Scale
Status	▪ FPI Q4 2012

Gantenerumab (RG1450)

Fully human monoclonal antibody against amyloid-beta

Indication	Prodromal Alzheimer's Disease	Mild Alzheimer's Disease
Phase/study	Phase II/III SCarlet RoAD	Phase III Marguerite Road
# of patients	N=799	N=1,000
Design	▪ 104-week subcutaneous treatment period ▪ ARM A: Gantenerumab (225 mg) ▪ ARM B: Gantenerumab (105 mg) ▪ ARM C: Placebo	▪ 104-week subcutaneous treatment period ▪ ARM A: Gantenerumab ▪ 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 ADCS-ADL at 2 years (co-primary)
Status	▪ Phase I PET data: Archives of Neurology 2012 Feb;69(2):198-207 ▪ Enrollment completed Q4 2013 ▪ Data expected in 2016	▪ FPI Q1 2014

Etrolizumab (RG7413)

A humanized monoclonal antibody against beta 7 integrin

Indication	Ulcerative colitis patients who are TNF naïve		
Phase/study	Phase III HIBISCUS I Induction study	Phase III HIBISCUS II Induction study	Phase III GARDENIA Sustained remission study
# of patients	N=350	N=350	N=720
Design	▪ ARM A: etrolizumab 105mg SC q4w + adalimumab placebo ▪ ARM B: etrolizumab placebo + adalimumab ▪ ARM C: etrolizumab placebo + adalimumab placebo	▪ ARM A: etrolizumab 105mg SC q4w + adalimumab placebo ▪ ARM B: etrolizumab placebo + adalimumab ▪ ARM C: etrolizumab placebo + adalimumab placebo	Time on treatment 54 weeks ▪ ARM A: etrolizumab 105mg SC q4w + placebo IV ▪ ARM B: placebo SC q4w + adalimumab SC
Primary endpoint	▪ Induction of remission compared with placebo as determined by the Mayo Clinic Score (MCS) at week 10	▪ Induction of remission compared with placebo as determined by the Mayo Clinic Score (MCS) at week 10	▪ Proportion of patients in sustained clinical remission as determined by Mayo Clinic Score (MCS) at weeks 10, 30 and 54
Status	▪ Expect FPI Q4 2014	▪ Expect FPI Q4 2014	▪ Expect FPI Q4 2014

Etrolizumab (RG7413)

A humanized monoclonal antibody against beta 7 integrin

Indication	UC patient who are TNF naïve and refractory or intolerant to immunosuppressant and/or corticosteroid treatment	UC patient who are refractory or intolerant of TNF inhibitors
Phase/study	Phase III LAUREL Maintenance study	Phase III HICKORY Induction and maintenance study
# of patients	N=350	N=800
Design	Induction phase: ▪ ARM A: open label etrolizumab 105mg SC q4w Maintenance study: ▪ ARM B: etrolizumab 105mg SC q4w ▪ ARM C: placebo	Cohort 1 (open-label): ▪ ARM A: etrolizumab induction + placebo maintenance ▪ ARM B: etrolizumab induction + maintenance Cohort 2 (blinded): ▪ ARM A: etrolizumab induction + maintenance ▪ ARM B: placebo induction + maintenance
Primary endpoint	▪ Maintenance of remission (at week 62) among randomized patients in remission at Week 10 as determined by the Mayo Clinic Score (MCS)	▪ Clinical Remission (Mayo Clinic Score, MCS) at Week 14 ▪ Remission maintenance (by MCS, at Week 66) among patients with remission at Week 14
Status	▪ FPI Q3 2014	▪ FPI Q2 2014

Etrolizumab (RG7413)

A humanized monoclonal antibody against beta 7 integrin

Indication	Moderate to severe ulcerative colitis	Moderate to severe ulcerative colitis
Phase/study	Phase II SPRUCE Open label extension study	Phase III COTTONWOOD Open label extension study
# of patients	N=116	N=2,600
Design	Patients who were enrolled in EUCALYPTUS study and meet enrollment criteria will receive etrolizumab 105 sc q4w	Patients who were previously enrolled in etrolizumab phase III studies and meet enrollment criteria will receive etrolizumab 105 sc q4w
Primary endpoint	Safety	- Long-term efficacy as determined by partial Mayo Clinic Score (pMCS) - Incidence of adverse events
Status	Recruitment completed	Expect FPI Q4 2014

HCV: Mericitabine, danoprevir, setrobuvir

IFN-free combination of different direct-acting antivirals in treatment-naïve patients

Indication	Hepatitis C patients Treatment-naïve or null-responders to interferon-based treatment
Phase/study	Phase II ANNAPURNA
# of patients	N=110
Design	▪ ARM A: GT1a including setrobuvir, danoprevir, ritonavir, ribavirin and mericitabine ▪ ARM B: GT1a including setrobuvir, danoprevir, ritonavir, ribavirin and mericitabine ▪ ARM C: GT1a including setrobuvir, danoprevir, ritonavir and ribavirin ▪ ARM D: GT1b including setrobuvir, danoprevir, ritonavir, ribavirin and mericitabine ▪ ARM E: GT1b including setrobuvir, danoprevir, ritonavir and ribavirin
Primary endpoint	▪ Sustained virological response at week 12 after the end of the study treatment
Status	▪ FPI Q2 2012 ▪ Recruitment Part 1 completed in Q4 2012 ▪ Data presented at APASL 2014 ▪ Publication expected in 2015

HCV: Danoprevir (RG7227)

IFN-based triple regimen for treatment-naïve patients of Asian origin conducted in China

Indication	Treatment-naïve patients of Asian origin with chronic hepatitis C genotype 1 with or without cirrhosis
Phase/study	Phase II DAPSANG
# of patients	N=61
Design	▪ Without cirrhosis: ▪ ARM A: Danoprevir 125 mg bid + Ritonavir 100mg bid+ Pegasys + Copegus for 12 weeks ▪ With compensated cirrhosis: ▪ ARM B: Danoprevir 125 mg bid + Ritonavir 100mg bid+ Pegasys + Copegus for 24 weeks
Primary endpoint	▪ Safety:
Status	▪ Recruitment completed Q4 2013 ▪ Study ongoing

Lampalizumab (RG7417)

Antibody fragment to selectively block activation of alternative complement pathway

Indication	Geographic atrophy (GA) secondary to age-related macular degeneration	
Phase/study	Phase III CHROMA	Phase III SPECTRI
# of patients	N=936	N=936
Design	▪ ARM A: Lampalizumab 10mg q4w ▪ ARM B: Lampalizumab 10mg q6w ▪ ARM C: Placebo	▪ ARM A: Lampalizumab 10mg q4w ▪ ARM B: Lampalizumab 10mg q6w ▪ ARM C: Placebo
Primary endpoint	▪ Primary: change in GA area ▪ Secondary: change in BCVA and in additional measures of visual function	▪ Primary: change in GA area ▪ Secondary: change in BCVA and in additional measures of visual function
Status	▪ FPI Q3 2014 ▪ Design presented at EURETINA 2014	▪ FPI Q3 2014 ▪ Design presented at EURETINA 2014

Lebrikizumab (RG3637)

A humanized monoclonal antibody designed to bind specifically to IL-13

	Severe uncontrolled adult asthma	
Indication	Adult patients whose asthma is uncontrolled with inhaled corticosteroids and a second controller medication	
Phase/study	Phase III LAVOLTA I	Phase III LAVOLTA II
# of patients	N=1,050	N=1,050
Design	▪ Subcutaneous lebrikizumab q4w on top of SOC for 52 weeks safety follow-up ▪ ARM A: Lebrikizumab high dose ▪ ARM B: Lebrikizumab low dose ▪ ARM C: Placebo ▪ Patients will be tested for periostin level	▪ Subcutaneous lebrikizumab q4w on top of SOC for 52 weeks safety follow-up ▪ ARM A: Lebrikizumab high dose ▪ ARM B: Lebrikizumab low dose ▪ ARM C: Placebo ▪ Patients will be tested for periostin level
Primary endpoint	▪ Rate of asthma exacerbations during the 52-week placebo-controlled period	▪ Rate of asthma exacerbations during the 52-week placebo-controlled period
Status	▪ FPI Q3 2013	▪ FPI Q3 2013

Lebrikizumab (RG3637)

A humanized monoclonal antibody designed to bind specifically to IL-13

Indication	Adolescent patients whose asthma is uncontrolled with inhaled corticosteroids and a second controller medication	Idiopathic pulmonary fibrosis
Phase/study	Phase III ACOUSTICS	Phase II RIFF
# of patients	N=375	N=250
Design	▪ Subcutaneous lebrikizumab q4w on top of SOC for 52 weeks with 52 week double-blind active treatment extension ▪ ARM A: Lebrikizumab high dose, week 1-104 or week 52-104 ▪ ARM B: Lebrikizumab low dose, week 1-104 or week 52-104 ▪ ARM C: Placebo, week 1-52	▪ ARM A: Lebrikizumab SC q4w ▪ ARM B: Placebo
Primary endpoint	▪ Rate of asthma exacerbations during the 52-week placebo-controlled period	▪ Progression-free survival
Status	▪ FPI Q3 2013	▪ FPI Q4 2013

Lebrikizumab (RG3637)

A humanized monoclonal antibody designed to bind specifically to IL-13

Indication	Adult asthma	Adult asthma mild-to-moderate patients
Phase/study	Phase II VOCALS	Phase III STRETTO
# of patients	N=225	N=300
Design	▪ ARM A: Lebrikizumab high dose SC q4w ▪ ARM B: Lebrikizumab low dose SC q4w ▪ ARM C: Placebo	▪ ARM A: Lebrikizumab SC q4w ▪ ARM B: Placebo ▪ ARM C: Montelukast
Primary endpoint	▪ Relative change in OCS dose at week 44	▪ Absolute change in FEV1 at week 12
Status	▪ FPI Q1 2014	▪ FPI Q2 2014

Ocrelizumab (RG1594)

2nd generation anti-CD20 monoclonal antibody

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=800	N=800	N=630
Design	96-week treatment period: ARM A: Ocrelizumab 2x 300 mg iv followed by 600 mg iv every 24 weeks ARM B: Interferon β-1a	96-week treatment period: ARM A: Ocrelizumab 2x 300 mg iv followed by 600 mg iv every 24 weeks ARM B: Interferon β-1a	120-week treatment period: ARM A: Ocrelizumab 2x 300 mg 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 Expanded Disability Status Scale (EDSS)
Status	- Enrolment completed Q1 2013 - Expect data in 2015	- Enrolment completed Q1 2013 - Expect data in 2015	- Enrolment completed Q1 2013 - Expect data in 2015

Pipeline summary

Marketed products additional indications

Global Development late-stage trials

pRED (Roche Pharma Research & Early Development)

gRED (Genentech Research & Early Development)

Roche Group YTD Sept 2014 sales

Diagnostics

Foreign exchange rate information

Oncology development programmes

Small molecules

Molecule	MDM2 (4) antagonist (RG7388)		MDM2 (4) ant. IV prodrug (RG7775)	LSD1 inhibitor (RG6016)	Raf/MEK inhibitor (RG7304, CKI27)
Indication	Solid tumors	Acute myeloid leukemia	Advanced cancers including AML	Acute Leukemia	Solid tumors
Phase	Phase I	Phase I	Phase I	Phase I	Phase I
# of patients	N=100	N=100	N=90	N=30	N=52
Design	Multiple ascending dose-escalation study	Multiple ascending dose-escalation study	Dose-escalation study ARM A: patients with advanced solid tumors ARM B: patients with r/r AML	Multiple ascending dose-escalation study	Dose-escalation to MTD
Primary endpoint	MTD	MTD	MTD	MTD	MTD and tumor assessment
Status	- Completed Q2 2013 - Data presented at ASCO 2014	FPI Q1 2013	FPI Q2 2014	FPI Q1 2014	- Initiated Q4 2008 - Enrolment stopped in Q4 2010
Collaborator				Oryzon Genomics, S.A.	Chugai

Oncology development programmes

Monoclonal antibodies

Molecule	Anti-glypican-3 MAb (RG7686, GC33)	
Indication	Metastatic liver cancer (hepatocellular carcinoma)	2L metastatic liver cancer (hepatocellular carcinoma)
Phase	Phase Ib	Phase II
# of patients	N= 40-50	N=171
Design	- Study US monotherapy - Study Japan monotherapy - Dose escalation study in combo with SOC	- Adaptive design study Double blind randomized 2:1 RG7686 : placebo - Patients are stratified according to the level of GPC-3 expression in tumor
Primary endpoint	Safety and tolerability	Progression-free survival
Status	- Recruitment completed Q4 2013 - Dose escalation completed for US and Japan monotherapy and combination therapy studies	- Recruitment completed Q1 2013 - Results under internal review
Collaborator	Chugai	

Oncology development programmes

Monoclonal antibodies (continued)

Molecule	GE-huMAb HER3 (RG7116)		
Indication	Solid tumors	HER2-low and HER3-positive metastatic breast cancer	1L mNSCLC of squamous histology
Phase	Phase I	Phase I	Phase Ib/II
# of patients	N=105	N=40	N=53
Design	- Multiple ascending dose study with extension cohorts and imaging sub-study - Combination arms with HER1-targeted therapies (erlotinib, cetuximab)	Multiple ascending dose of RG7116 in combination with Perjeta and paclitaxel	RG7116 in combination with carboplatin and paclitaxel
Primary endpoint	Safety, PK	Safety	Safety, ORR
Status	- FPI Q4 2011 - Initial data presented at ASCO 2013	FPI Q3 2013	Expect FPI Q4 2014

Oncology development programmes

Monoclonal antibodies (continued)

Molecule	Ang2-VEGF MAb (RG7221)		CSF-1R huMAb (RG7155)	CEA-IL2v (RG7813)
Indication	Solid tumors	Metastatic colorectal cancer	Solid tumors and PVNS	Solid tumors
Phase	Phase I	Phase II McCAVE	Phase I/II	Phase I
# of patients	N≈80	N=140	N≈140	N~110
Design	Multiple ascending dose study with extension cohorts in solid tumors to assess the PD effects and platinum resistant ovarian cancer	ARM A: Induction: Avastin+mFOLFOX-6; followed by maintenance: Avastin+5-FU/LV ARM B: Induction: RG7221+mFOLFOX-6; followed by maintenance: RG7221+5-FU/LV	Multiple ascending dose study +/- paclitaxel with extension cohorts	Single and multiple dose escalation study with extension cohorts
Primary endpoint	Safety, PK	PFS	Safety, PK, PD & preliminary clinical activity	Safety, PK, PD
Status	- FPI Q4 2012 - Dose escalation data presented at ASCO 2014	FPI Q2 2014	- FPI Q4 2011 - Biomarker data presented at AACR 2013 and AACR 2014 - Data presented at ASCO 2014	FPI Q4 2013

Neuroscience development programmes

	Metabolic glutamate receptor pathway			
Molecule	Decoglurant (mGlu2 NAM, RG1578)	Basimglurant (mGlu5 NAM, RG7090)		
Indication	Adjunctive Treatment of Major Depressive Disorder	Adjunctive Treatment of Major Depressive Disorder	Fragile X Syndrome	
Phase/study	Phase II ArtDeCo	Phase II Marigold	Phase II Fragxis	Phase II FoXtail
# of patients	N=480	N=300	N=180	N=45 Pediatric patients
Design	▪ ARM A: decoglurant 5 mg ▪ ARM B: decoglurant 15 mg ▪ ARM C: decoglurant 30 mg ▪ ARM D: matching placebo	▪ ARM A: basimglurant 0.5 mg ▪ ARM B: basimglurant 1.5 mg ▪ ARM C: matching placebo	▪ ARM A: basimglurant 0.5 mg ▪ ARM B: basimglurant 1.5 mg ▪ ARM C: matching placebo	▪ ARM A: basimglurant dose A ▪ ARM B: basimglurant dose B ▪ ARM C: matching placebo
Primary Endpoint	▪ Efficacy - Montgomery Asberg Depression Rating Scale	▪ Efficacy - Montgomery Asberg Depression Rating Scale	▪ Efficacy, safety and tolerability	▪ Safety ▪ Exploratory efficacy and tolerability
Status	▪ Recruitment completed ▪ Data in-house under review	▪ Study completed ▪ Data in-house under review ▪ To be presented at ECNP 2014	▪ Primary endpoint not met as reported to patient advocacy groups in Sept. 2014	▪ Primary endpoint (safety) met (not powered for efficacy) as reported to patient advocacy groups in Sept. 2014

Neuroscience development programmes

Molecule	PDE10A inhibitor (RG7203)	TAAR1 agonist (RG7410)	GABRA5 NAM (RG1662)	mGlu5 PAM (RG7342)
Indication	Schizophrenia	Schizophrenia	Down Syndrome	Schizophrenia
Phase	Phase I	Phase I	Phase IIB CLEMATIS	Phase I
# of patients	N=26	N= up to 40	N=180	N=93
Design	▪ Multiple dose, double-blind study in schizophrenia patients ▪ ARM A: RG7203 plus risperidone ▪ ARM B: placebo plus risperidone	▪ Double-blind, randomized, placebo controlled, sequential multiple ascending dose study in HVs	▪ For 26 weeks patients will receive: ▪ ARM A: RG1662 120mg twice daily ▪ ARM B: RG1662 120mg twice daily ▪ ARM C: Placebo	▪ Single ascending dose of RG7342
Primary endpoint	▪ Safety, tolerability, PK	▪ Safety and tolerability in HVs	▪ Cognition and adaptive behavior	▪ Safety, tolerability, PK and food effect
Status	▪ Study completed	▪ FSI Q3 2014	▪ FPI Q2 2014	▪ FPI Q3 2014

Neuroscience development programmes

Molecule	Monoamine oxidase type B (MAO-B) inhibitor (RG1577, EVT-302)	V1 receptor antagonist (RG7314)	SMN2 splicing modifier (RG7800)	
Indication	Alzheimer's Disease	Autism	Spinal muscular atrophy	
Phase	Phase IIb MAYfloweR RoAD	Phase II VANILLA	Phase I	Phase Ib MOONFISH
# of patients	N=495	N=150	N=48	N=48
Design	52-week oral treatment ARM A: RG1577 (dose 1) ARM B: RG1577 (dose 2) ARM C: placebo	Multi-center, randomized, double-blind, placebo-controlled proof-of-concept study in individuals with Autism Spectrum Disorder (ASD)	Healthy volunteer study ARM A: RG7800 Single dose ARM B: Placebo	Randomized, double-blind, 12-week, placebo-controlled multiple dose study in adult and pediatric patients
Primary endpoint	Changes in ADAS-Cog at 52 weeks	Safety and efficacy	Safety, PK	Safety and tolerability
Status	Recruitment completed Q1 2014	FSI Q3 2013	Completed	Expect FPI Q4 2014
Collaborator	Evotec		PTC Therapeutics/ SMA Foundation	

Neuroscience development programmes

Molecule	Anti- α Syn (RG7935, PRX002)	
Indication	Parkinson's disease	
Phase	Phase I	Phase I
# of patients	N=40	N=up to 60
Design	▪ Double-blind, placebo-controlled, single ascending dose study of RG7935/PRX002 in healthy subjects	▪ Double-blind, placebo-controlled, multiple ascending dose study of RG7935/PRX002 in patients with Parkinson's disease
Primary endpoint	▪ Safety, tolerability, PK	▪ Safety and tolerability
Status	▪ FSI Q2 2014	▪ FPI July 2014
Collaborator	Prothena	

Infectious diseases programmes

Molecule	TLR7 agonist (RG7795)	LptD antibiotic (RG7929)
Indication	Chronic hepatitis B	Pseudomonas infections (including MDR strains)
Phase	Phase I	Phase II
# of patients	N=50	N=~50
Design	▪ Healthy volunteer study ▪ ARM A: Single ascending dose of RG7795 ▪ ARM B: Placebo	▪ Patient and HV study
Primary endpoint	▪ Safety	▪ Safety, PK/PD and efficacy
Status	▪ FPI Q4 2013	▪ FPI Q4 2013 ▪ QIDP and fast track designation granted Q2 2014
Collaborator		Polyphor

Metabolic development programmes

Molecule	Inclacumab (P-selectin huMAb, RG1512)	
Indication	Prevention of saphenous vein graft disease Patients undergoing coronary artery bypass graft (CABG) surgery	Acute Coronary Syndrome (ACS) Patients undergoing percutaneous coronary intervention (PCI)
Phase/study	Phase II SELECT-CABG	Phase II SELECT-ACS
# of patients	N=384	N=516
Design	▪ 32-week treatment period ▪ ARM A: Inclacumab (20 mg/kg) ▪ ARM B: Placebo	▪ Single infusion ▪ ARM A: Inclacumab (5 mg/kg) ▪ ARM B: Inclacumab (20 mg/kg) ▪ ARM C: Placebo
Primary Endpoint	▪ Saphenous vein graft re-occlusion	▪ Procedural damage (troponin)
Status	▪ Recruitment completed Q2 2012 ▪ Data to be published in 2014	▪ Recruitment completed ▪ Data presented at ACC 2013
	▪ Candidate for partnering-out	
Collaborator	Genmab	

Metabolic, ophthalmology and hemophilia development programmes

Molecule	GLP-1/GIP dual agonist (MAR709, RG7697)	Aldosterone synthase inhibitor (RG7641)	Anti-VEGF/Ang2 (RG7716)	Factor IX/X bispecific (RG6013, ACE910)
Indication	Type 2 diabetes	Metabolic diseases	Wet age-related macular degeneration	Hemophilia A Study in Japanese patients
Phase/study	Phase II	Phase I	Phase I	Phase I/II
# of patients	N=105	N=96	N=30	N≈18
Design	▪ ARM A: RG7697 SC ▪ ARM B: Liraglutide ▪ ARM C: Placebo	▪ ARM A: RG7641 single dose ▪ ARM B: Placebo	▪ Healthy volunteer study ▪ Single ascending dose of RG7716	▪ Extension study in patients from Phase I ▪ Phase I enrolled 64 HVs and 18 patients
Primary Endpoint	▪ HbA1c	▪ Safety	▪ Safety	▪ Exploratory efficacy and safety
Status	▪ FPI Q3 2014	▪ Recruitment completed Q2 2014	▪ FPI Q4 2013	▪ Phase I recruitment completed Q2 2014 in Japan ▪ Extension study FPI Q3 2013
Collaborator	Marcadia Biotech, Inc. acquisition			Chugai

Pipeline summary

Marketed products additional indications

Global Development late-stage trials

pRED (Roche Pharma Research & Early Development)

gRED (Genentech Research & Early Development)

Roche Group YTD Sept 2014 sales

Diagnostics

Foreign exchange rate information

Oncology development programmes

Monoclonal antibodies

	Growth factor signaling			
Molecule	Anti-HER3 EGFR DAF MAb (RG7597)			
Indication	Metastatic/recurrent SCCHN	KRAS wild-type metastatic colorectal cancer	1L recurrent/metastatic squamous cell carcinoma of head and neck	Locally advanced or metastatic tumors with mutant KRAS
Phase/study	Phase II MEHGAN	Phase II DARECK	Phase Ib	Phase I
# of patients	N=170	N=130	N=21	N=50
Design	▪ ARM A: RG7597 ▪ ARM B: Cetuximab	▪ ARM A: RG7597+FOLFIRI ▪ ARM B: Cetuximab+FOLFIRI	▪ Evaluating safety/tolerability with two chemo backbones ▪ Arm A: Cisplatin/5-FU ▪ Arm B: Carboplatin/Paclitaxel	▪ Dose finding of RG7597 plus cobimetinib¹
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival	▪ Safety, DLT, PK	▪ Safety
Status	▪ Recruitment completed Q2 2013 ▪ Primary endpoint to met ▪ Data presented at ESMO 2014	▪ Recruitment completed Q4 2013	▪ Recruitment completed Q2 2014	▪ FPI Q4 2013

¹cobimetinib in collaboration with Exelixis

SCCHN=Squamous Cell Carcinoma of the Head and Neck

FOLFOX=Folinic acid, Fluorouracil, Oxaliplatin; FOLFIRI=Folinic acid, Fluorouracil, Irinotecan

Oncology development programmes

Monoclonal antibodies

Molecule	Anti-OX40 (RG7888, MOXR0916)
Indication	Solid tumors
Phase I	Phase I
# of patients	N=400
Design	▪ RG7888 dose escalation and expansion
Primary endpoint	▪ Safety
Status	▪ FPI Q3 2014

Oncology development programmes

Antibody drug conjugates

	Antibody drug conjugates (ADCs)		
Molecule	Anti-STEAP1 ADC (RG7450)	Anti-MUC16 ADC (RG7458)	NME ADC (RG7882)
Indication	Prostate cancer	Ovarian and pancreatic cancer	Pt. resistant ovarian cancer or unresectable pancreatic cancer
Phase	Phase I	Phase I	Phase I
# of patients	N=67	N=57	N=75
Design	▪ Dose escalation study	▪ Dose escalation study	▪ Dose escalation study
Primary endpoint	▪ Safety	▪ Safety/PK	▪ Safety/PK
Status	▪ FPI Q1 2011 ▪ Data presented at ASCO 2013-2014 and AACR 2014	▪ Recruitment completed Q2 2014 ▪ Safety and PK data presented at AACR 2013	▪ FPI Q2 2014
Collaborator	Seattle Genetics and Agensys	Seattle Genetics	

Oncology development programmes

Antibody drug conjugates (continued)

	Antibody drug conjugates (ADCs)		
Molecule	Lifastuzumab vedotin (anti-NaPi2b ADC, RG7599)		
Indication	NSCLC and ovarian cancer	Platinum-sensitive ovarian cancer and NSCLC	Platinum-resistant ovarian cancer
Phase	Phase I	Phase Ib	Phase II HERAEA
# of patients	N=96	N=52	N=92
Design	▪ Dose escalation study	▪ Dose escalation of RG7599 in combination with carboplatin, with or without Avastin	▪ ARM A: RG7599 ▪ ARM B: Pegylated liposomal doxorubicin
Primary endpoint	▪ Safety	▪ Safety, PK	▪ Progression-free survival
Status	▪ FPI Q2 2011 ▪ Data presented at ASCO 2014	▪ FPI Q4 2013	▪ FPI Q1 2014
Collaborator	Seattle Genetics		

Oncology development programmes

Antibody drug conjugates (continued)

	Antibody drug conjugates (ADCs)	
Molecule	Pinatuzumab vedotin (RG7593) vs. polatuzumab vedotin (RG7596)	NME ADC (RG7841)
Indication	Non-Hodgkin's Lymphoma	Refractory solid tumors
Phase	Phase II ROMULUS	Phase I
# of patients	N=120	N=115
Design	▪ Pinatuzumab vedotin plus Rituxan ▪ Polatuzumab vedotin plus Rituxan	▪ Dose escalation study
Primary endpoint	▪ Safety and anti-tumor activity	▪ Safety
Status	▪ Recruitment completed Q1 2014 ▪ Interim data presented at ASCO and EHA 2014	▪ FPI Q2 2014
Collaborator	Seattle Genetics	

Oncology development programmes

Small molecules

Molecule	Ipatasertib (AKT inhibitor, GDC-0068, RG7440)		
Indication	2L Castration-resistant prostate cancer	1L metastatic gastric or gastroesophageal junction adenocarcinoma	1L triple-negative breast cancer
Phase	Phase II A.MARTIN	Phase II JAGUAR	Phase II LOTUS
# of patients	N=262	N=120	N=120
Design	▪ ARM A: Ipatasertib (400mg) + abiraterone ▪ ARM B: Ipatasertib (200mg) + abiraterone ▪ ARM C: Placebo + abiraterone	▪ ARM A: Ipatasertib + mFOLFOX6 ▪ ARM B: Placebo + mFOLFOX6	▪ ARM A: Ipatasertib + paclitaxel ▪ ARM B: Placebo + paclitaxel
Primary endpoint	▪ Progression-free survival	▪ Progression-free survival	▪ Progression-free survival
Status	▪ FPI Q3 2013	▪ FPI Q3 2013	▪ FPI Q3 2014
Collaborator	Array BioPharma		

Oncology development programmes

Small molecules (continued)

Molecule	Ipatasertib (AKT inhibitor, GDC-0068, RG7440)	
Indication	Solid tumors	Solid tumors
Phase	Phase Ib	Phase Ib
# of patients	N=120	N=62
Design	▪ Dose escalation with: ▪ ARM A: Docetaxel ▪ ARM B: Fluoropyrimidine plus oxaliplatin ▪ ARM C: Paclitaxel ▪ ARM D: Enzalutamide	▪ Dose escalations study of ipatasertib in combination with cobimetinib* (MEK inhibitor)
Primary endpoint	▪ Safety	▪ Safety/PK
Status	▪ FPI Q3 2011 ▪ Data presented at ASCO and ESMO 2012	▪ FPI Q2 2012 ▪ Data presented at AACR 2014
Collaborator	Array BioPharma	

*cobimetinib in collaboration with Exelixis

ASCO=American Society of Clinical Oncology; ESMO=European Society for Medical Oncology; AACR=American Association for Cancer Research

Oncology development programmes

Small molecules (continued)

Molecule	ChK1 inhibitor (RG7741, GDC-0575)	ERK inhibitor (RG7842, GDC-0994)	PI3 Kinase inhibitor (RG7666, GDC-0084)	Selective estrogen receptor degrader (SERD) (RG6046, GDC-0810/ARN-810)
Indication	Solid tumors or lymphoma	Solid tumors	Progressive or recurrent high-grade glioma	Metastatic ER+ HER2- breast cancer
Phase	Phase I	Phase I	Phase I	Phase I
# of patients	N=170	N=78	N=68	N=72
Design	▪ Stage 1: Dose escalation ▪ Stage 2: Cohort expansion	▪ Stage 1: Dose escalation ▪ Stage 2: Cohort expansion	▪ Dose escalation study	▪ Dose escalation study
Primary endpoint	▪ Safety/PK	▪ Safety, MTD, PK	▪ Safety/PK	▪ Safety
Status	▪ FPI Q2 2012	▪ FPI Q2 2013	▪ FPI Q2 2012	▪ FPI Q1 2013
Collaborator	Array BioPharma			Seragon acquisition

Immunology development programmes

Molecule	Quilizumab (Anti-M1 prime, RG7449)		anti-IL17 (RG7624)
Indication	Allergic asthma - inadequately controlled	Chronic spontaneous urticaria	Autoimmune diseases
Phase/study	Phase IIb COSTA	Phase II QUAIL	Phase Ib
# of patients	N=560	N=30	N=21
Design	▪ SC administration on top of SOC ▪ ARM A: Quilizumab 300mg ▪ ARM B: Quilizumab 150mg ▪ ARM C: Quilizumab 450mg ▪ ARM D: Placebo	▪ ARM A: Quilizumab sc ▪ ARM B: Placebo sc	▪ Randomized, double-blind, placebo-controlled, multiple ascending dose escalation study
Primary endpoint	▪ Rate of protocol-defined exacerbations from baseline to week 36	▪ Efficacy and safety	▪ Safety and tolerability
Status	▪ Recruitment completed Q3 2013	▪ Recruitment completed Q2 2014	▪ Enrolment completed Q2 2012 ▪ Next study in preparation
Collaborator			NovImmune

Neuroscience development programmes

Molecule	Crenezumab (Anti-A β , RG7412)		
Indication	Alzheimer's Disease		Alzheimer's Prevention initiative (API) Colombia
Phase/study	Phase II ABBY Cognition study	Phase II BLAZE Biomarker study	Phase II Cognition study
# of patients	N=450	N=91	N=300
Design	▪ ARM A: Crenezumab sc ▪ ARM B: Crenezumab iv ▪ ARM C: Placebo	▪ ARM A: Crenezumab sc ▪ ARM B: Crenezumab iv ▪ ARM C: Placebo	▪ ARM A: 100 carriers receive crenezumab sc ▪ ARM B: 100 carriers receive placebo ▪ ARM C: 100 non-carriers receive placebo
Primary endpoint	▪ Change in cognition (ADAS-cog) and Clinical Dementia Rating, Sum of Boxes (CDR-SOB) score from baseline to week 73	▪ Change in brain amyloid load from baseline to week 69	▪ Change on Alzheimer's Prevention Initiative (API) Composite Cognitive Test total score
Status	▪ Enrolment completed Q3 2012 ▪ Data presented at AAIC 2014	▪ Enrolment completed Q3 2012 ▪ Cognition data presented at AAIC 2014 ▪ Biomarker data to be presented at CTAD 2014	▪ FPI Q4 2013
Collaborator	AC Immune		AC Immune and Banner Alzheimer's Institute

Neuroscience and infectious diseases development programmes

Molecule	Nav1.7 (RG7893, GDC-0276)	Anti-Flu A (RG7745)
Indication	Pain	Influenza
Phase/study	Phase I	Phase IIa
# of patients	N=74	N=100
Design	Phase 1, randomized, placebo-controlled, double blinded study to determine safety, tolerability, and pharmacokinetics in healthy volunteers	- Healthy volunteers in an influenza challenge model - ARM A: RG7745 - ARM B: Placebo - ARM C: Tamiflu
Primary endpoint	Safety, tolerability, and pharmacokinetics of single and multiple doses	Reduction in viral activity
Status	FPI Q3 2014	- Data positive with 98% reduction of viral load at 3600mg dose - Presented at ISIRV 2014
Collaborator	Xenon Pharmaceuticals Inc.	

Pipeline summary

Marketed products additional indications

Global Development late-stage trials

pRED (Roche Pharma Research & Early Development)

gRED (Genentech Research & Early Development)

Roche Group YTD Sept 2014 sales

Diagnostics

Foreign exchange rate information

Geographical sales split by divisions and Group*

CHFm	YTD Sep 2013	YTD Sep 2014	% change CER
Pharmaceuticals Division	27,190	26,965	+4
United States	11,429	11,528	+5
Europe	6,952	7,070	+3
Japan	2,492	2,406	+7
International	6,317	5,961	+3
Diagnostics Division	7,677	7,792	+6
United States	1,706	1,736	+6
Europe	3,024	3,058	+2
Japan	358	333	+3
International	2,589	2,665	+12
Group	34,867	34,757	+5
United States	13,135	13,264	+5
Europe	9,976	10,128	+2
Japan	2,850	2,739	+7
International	8,906	8,626	+6

CER = Constant Exchange Rates (avg full year 2013)

* Geographical sales split shown here does not represent operational organization

Pharma Division sales YTD Sep 2014 (vs. 2013)

Top 20 products

	Global		US		Europe		Japan		International	
	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER
MabThera/Rituxan	5,124	3	2,483	0	1,517	6	165	2	959	6
Avastin	4,749	6	2,002	5	1,472	3	519	11	756	10
Herceptin	4,679	7	1,451	10	1,696	3	200	4	1,332	9
Lucentis	1,260	5	1,260	5	-	-	-	-	-	-
Tarceva	971	0	486	7	229	-11	73	17	183	-10
Actemra/RoActemra	897	24	292	31	320	22	156	23	129	17
Pegasy	811	-17	167	-34	194	-29	49	37	401	-4
Xolair	701	24	701	24	-	-	-	-	-	-
Perjeta	633	255	380	190	157	319	58	*	38	*
Xeloda	623	-43	171	-63	76	-68	67	-8	309	-8
CellCept	623	-4	153	1	164	-7	42	-8	264	-5
Activase/TNKase	559	14	522	14	-	-	-	-	37	14
Valcyte/Cymevene	545	14	295	18	135	12	-	-	115	8
Tamiflu	452	14	252	10	73	*	63	-20	64	-15
Pulmozyme	427	8	281	11	92	-1	0	-	54	10
Kadcyla	371	148	213	46	111	*	21	-	26	*
NeoRec./Epogin	342	-10	-	-	145	-13	43	-37	154	7
Mircera	304	6	-	-	76	1	144	5	84	14
Zelboraf	230	-8	53	-42	145	2	-	-	32	69
Madopar	206	-8	-	-	79	-5	12	-2	115	-10

CER = Constant Exchange Rates (avg full year 2013)

* over +500%

Pharma Division sales YTD Sep 2014 (vs. 2013)

Recently launched products

	Global		US		Europe		Japan		International	
	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER
Erivedge	90	93	57	29	29	*	-	-	4	*
Gazyva	32	-	30	-	1	-	-	-	1	-

Pharma Division CER sales growth¹ in %

Global top 20 products

	Q3/13	Q4/13	Q1/14	Q2/14	Q3/14
MabThera/Rituxan	12	7	3	5	1
Avastin	14	13	9	4	6
Herceptin	7	7	3	9	9
Lucentis	21	22	8	4	2
Tarceva	5	4	-5	3	0
Actemra/RoActemra	33	23	23	21	28
Pegasy	-16	-20	-19	-10	-22
Xolair	14	17	15	22	33
Perjeta	262	394	274	277	227
Xeloda	6	-3	-19	-50	-61
CellCept	-2	-10	-1	-11	0
Activase/TNKase	18	19	-1	26	19
Valcyte/Cymevene	0	26	12	12	19
Tamiflu	115	-27	9	-36	121
Pulmozyme	0	18	3	8	13
Kadcyla	-	-	474	105	103
NeoRec./Epogin	-16	-14	-9	-8	-12
Mircera	29	23	21	2	-1
Zelboraf	38	26	-2	-9	-13
Madopar	3	9	-20	3	-6

CER = Constant Exchange Rates (avg full year 2013)

¹ Q3-Q4/13 vs. Q3-Q4/12; Q1-Q3/14 vs. Q1-Q3/13

* over 500%

Pharma Division CER sales growth¹ in %

Top 20 products by region

	US				Europe				Japan				International			
	Q4	Q1	Q2	Q3	Q4	Q1	Q2	Q3	Q4	Q1	Q2	Q3	Q4	Q1	Q2	Q3
MabThera/Rituxan	2	-2	8	-4	0	6	8	4	8	20	-17	5	26	12	-2	9
Avastin	4	6	6	3	9	8	2	1	12	27	-5	13	49	7	9	15
Herceptin	3	4	17	10	-2	2	4	4	11	23	-12	5	21	0	12	14
Lucentis	22	8	4	2	-	-	-	-	-	-	-	-	-	-	-	-
Tarceva	-8	-6	16	11	-3	-12	-12	-9	25	42	6	7	27	-6	-9	-13
Actemra/RoActemra	20	22	30	39	21	20	20	25	24	49	5	21	31	3	28	21
Pegasy	-55	-40	-14	-51	-14	-19	-32	-38	-28	16	45	48	1	-7	3	-8
Xolair	17	15	22	33	-	-	-	-	-	-	-	-	-	-	-	-
Perjeta	201	161	205	202	*	*	287	228	-	-	-	375	*	*	*	341
Xeloda	-8	-15	-80	-93	-9	-57	-71	-76	0	8	-23	-6	9	-3	-2	-19
CellCept	5	-7	-6	16	-5	-10	-5	-6	8	5	-14	-12	-24	6	-16	-3
Activase/TNKase	22	0	27	18	-	-	-	-	-	-	-	-	-13	-4	19	27
Valcyte/Cymevene	19	26	8	21	28	5	10	22	-	-	-	-	38	-7	24	12
Tamiflu	-24	-9	22	155	-	*	-71	49	-47	-17	-74	238	-72	-8	-44	37
Pulmozyme	16	2	10	20	1	2	-1	-5	12	150	-	38	41	10	9	10
Kadcyla	-	315	16	3	-	-	*	*	-	-	-	-	-	-	*	*
NeoRec./Epogin	-	-	-	-	-19	-14	-10	-15	-22	-26	-45	-38	-2	7	12	3
Mircera	-	-	-	-	29	8	-1	-3	21	36	-12	-2	20	8	32	1
Zelboraf	-1	-40	-46	-38	17	12	8	-12	-	-	-	-	425	98	56	65
Madopar	-	-	-	-	-1	-9	-2	-5	3	4	-6	-2	18	-29	7	-7

CER = Constant Exchange Rates (avg full year 2013)

* over 500%

¹ Q4/13 vs. Q4/12; Q1-Q3/14 vs. Q1-Q3/13

CER sales growth (%)

Quarterly development

2013 vs. 2012

2014 vs. 2013

Q1 Q2 Q3 Q4 Q1 Q2 Q3

Pharmaceuticals Division

United States

Europe

Japan

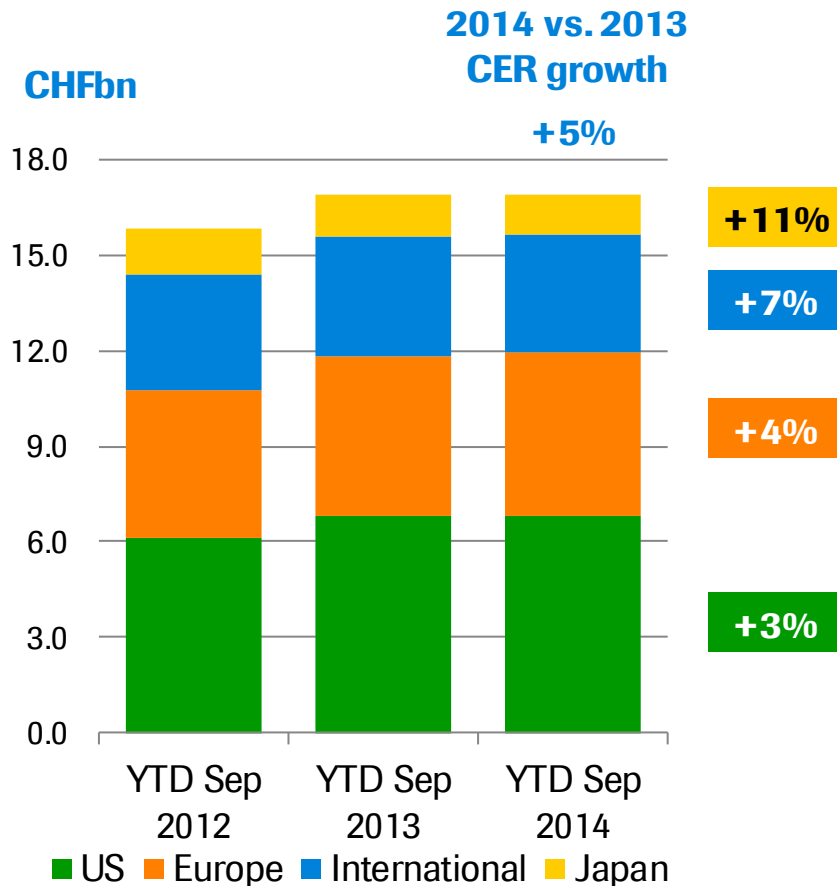
International

Diagnostics Division

Roche Group

7	4	9	7	4	4	4
13	7	16	5	3	8	4
1	2	3	2	5	1	1
2	2	4	2	19	-4	8
8	2	5	18	1	3	6
1	4	7	5	7	5	7
6	4	8	7	5	4	5

YTD Sept 2014: Oncology franchise



YTD Sept 2014 sales of CHF 16.910bn

US

- HER2 Franchise (including strong uptake of PERJETA & Kadcyla) and Avastin driving growth and performance

Europe

- Growth mainly driven by Perjeta and Kadcyla as well as MabThera (+6%)

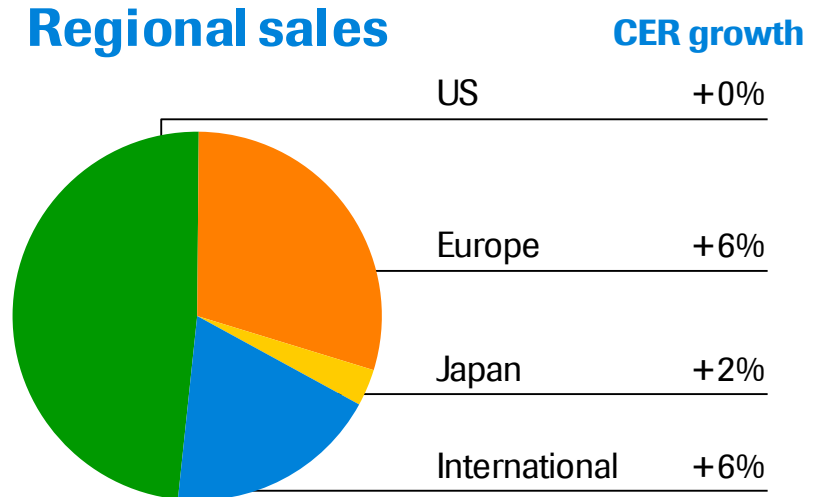
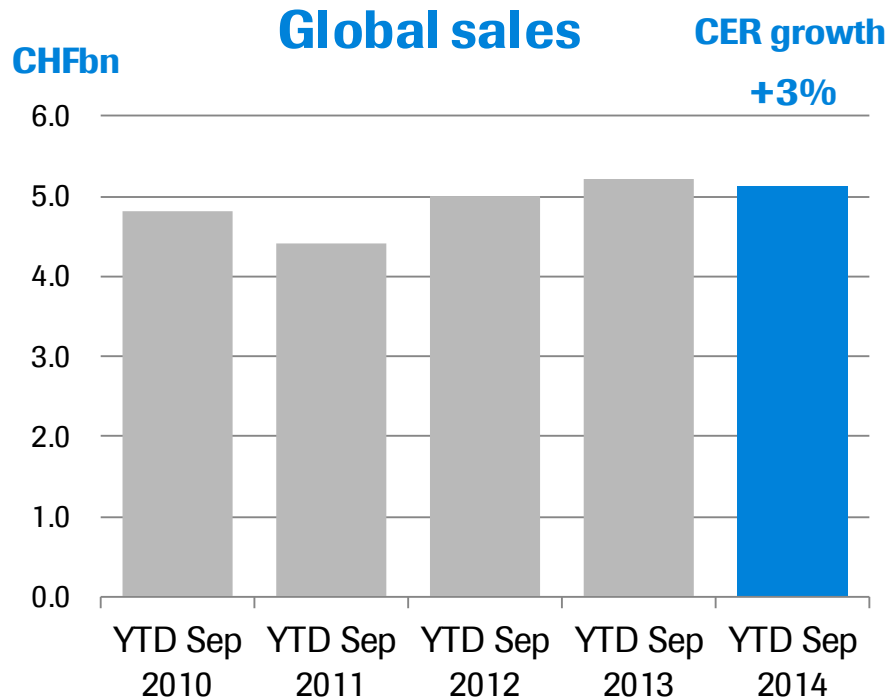
International

- Continued strong growth for Avastin and Herceptin

Japan

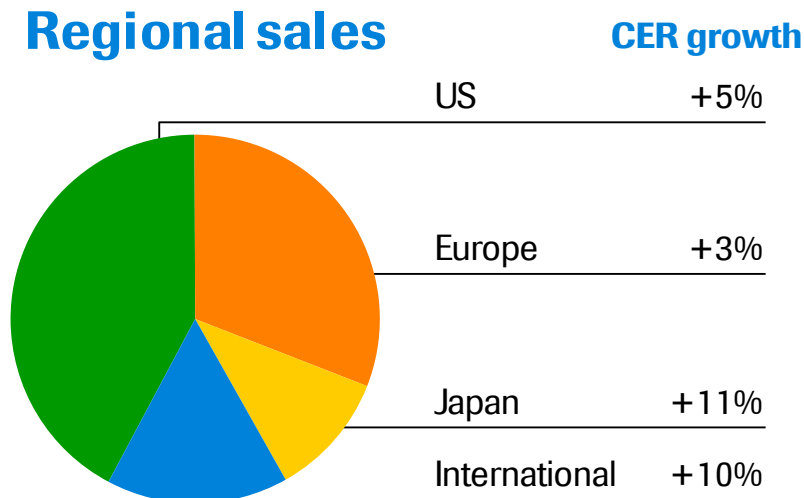
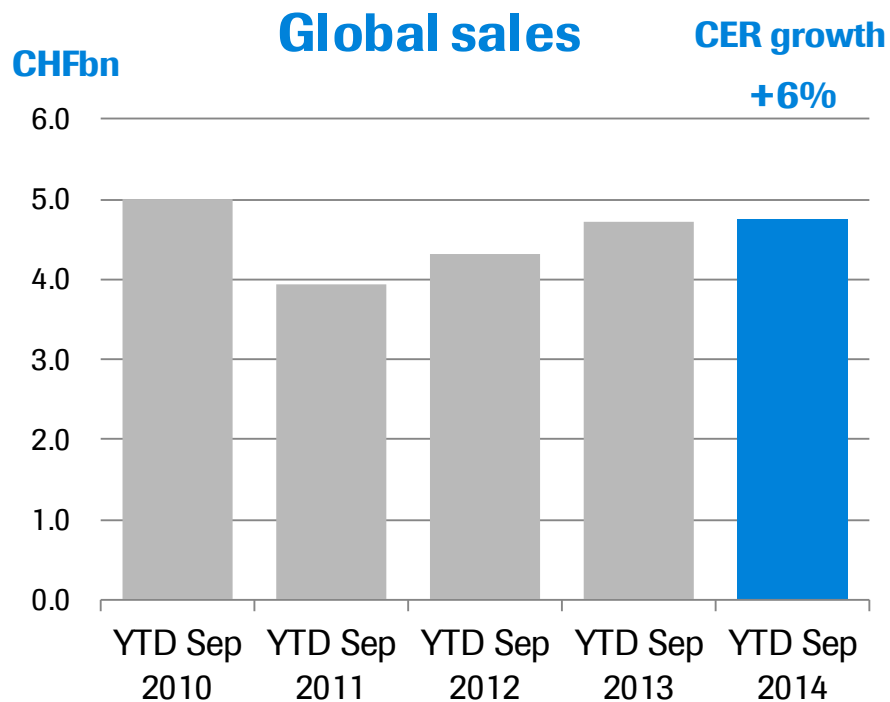
- Growth driven largely by Avastin

MabThera/Rituxan



YTD Sept 2014 sales of CHF 5.124bn

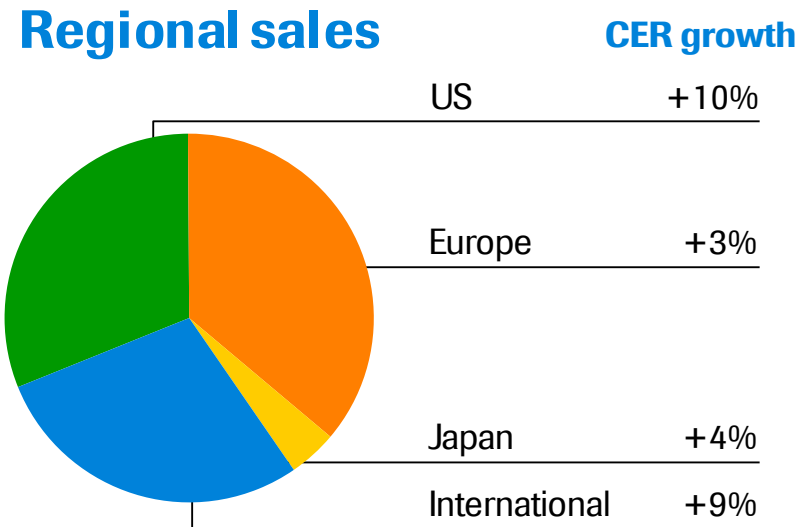
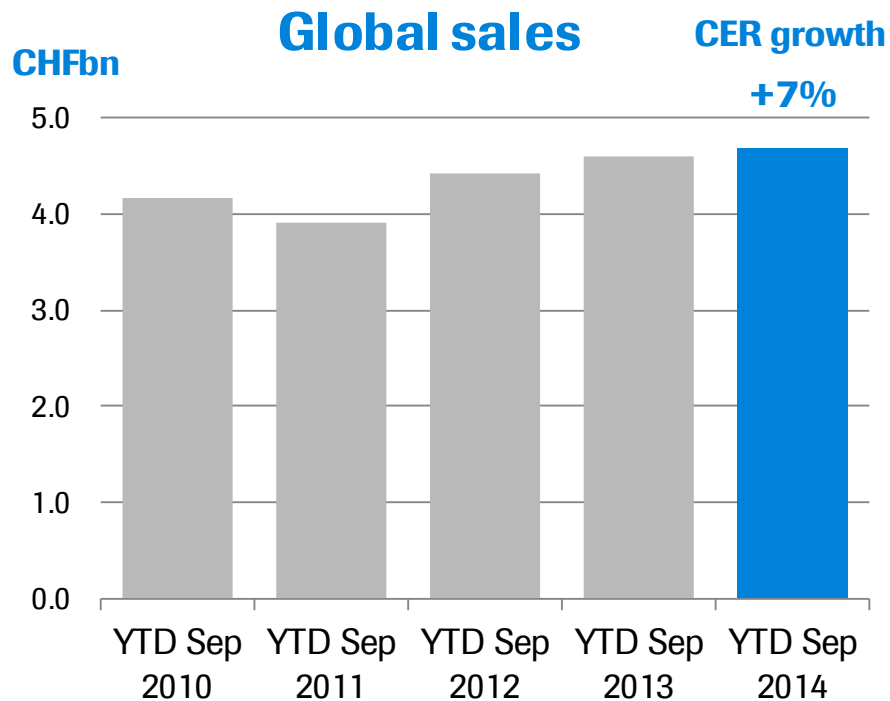
- Overall solid growth with high penetration in developed markets
- Europe: Strong growth driven by increased market share in follicular lymphoma, as well as CLL (1L)
- US: Sales stable but comparison distorted by 340B baseline effect
- International: Driven by strong growth in Latin America



YTD Sept 2014 sales of CHF 4.749bn

- Europe: Strong growth driven by further uptake in ovarian and breast cancer
- US: Continued increase in mCRC use associated with TML awareness
- Japan: Driven by increased demand in CRC, breast and ovarian cancer, as well as GBM
- International: Driven by launches for ovarian cancer and uptake in colorectal cancer

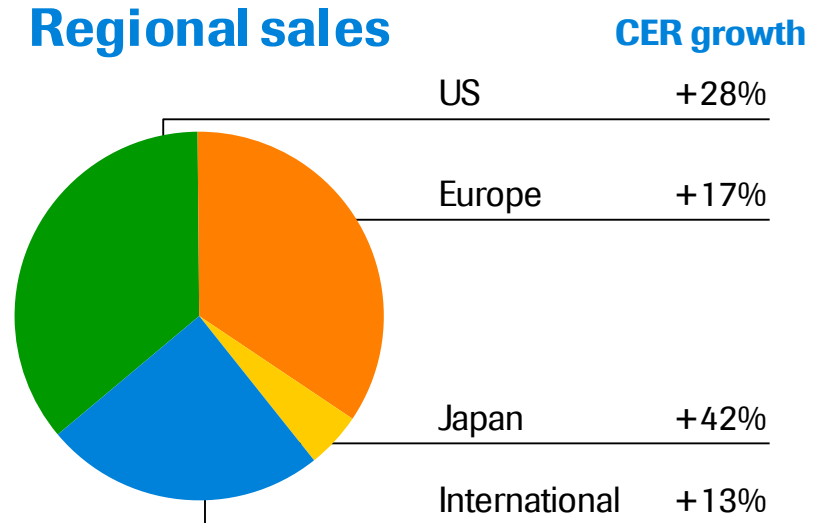
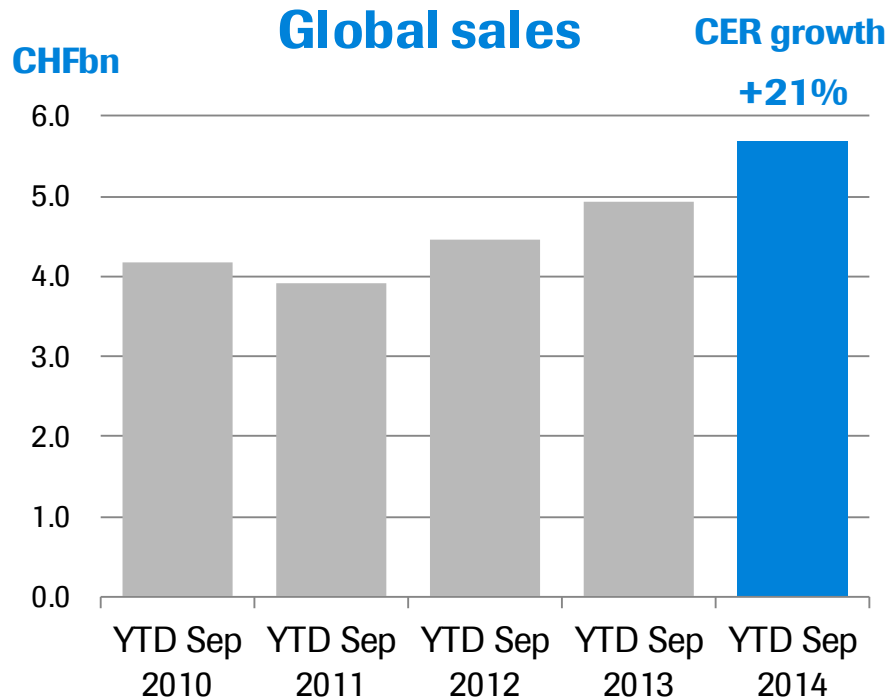
Herceptin



YTD Sept 2014 sales of CHF 4.679bn

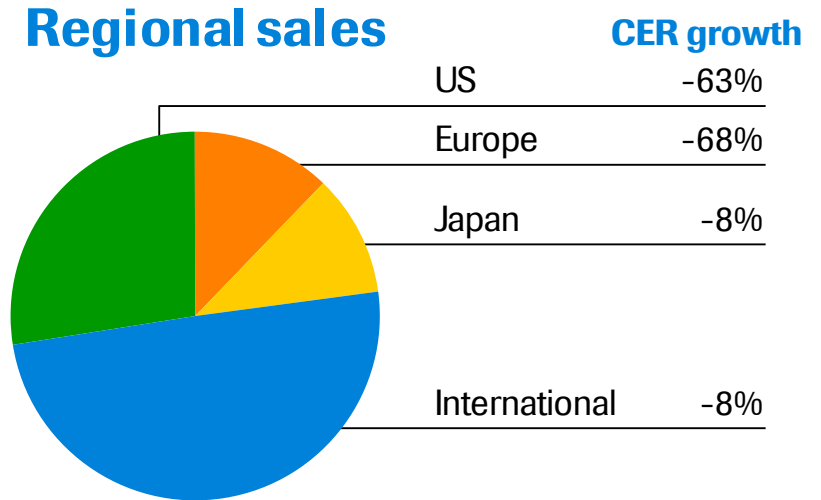
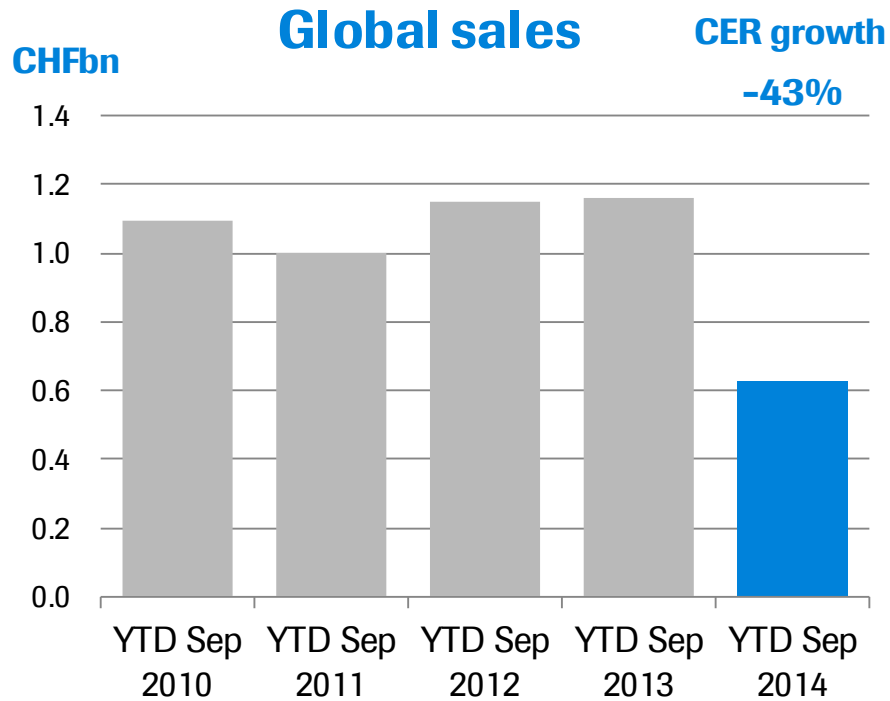
- Positive growth in all regions driven by higher volumes / prolonged treatment times
- US: Continued growth in mBC (1L)
- Europe: Strong demand in Germany, Spain and the UK
- Japan: Increased usage in combination with Perjeta in breast, as well as in gastric cancer
- International: Growth driven by China and Brazil

HER2 franchise (Herceptin, Perjeta & Kadcyła)



YTD Sept 2014 sales of CHF 5.683bn

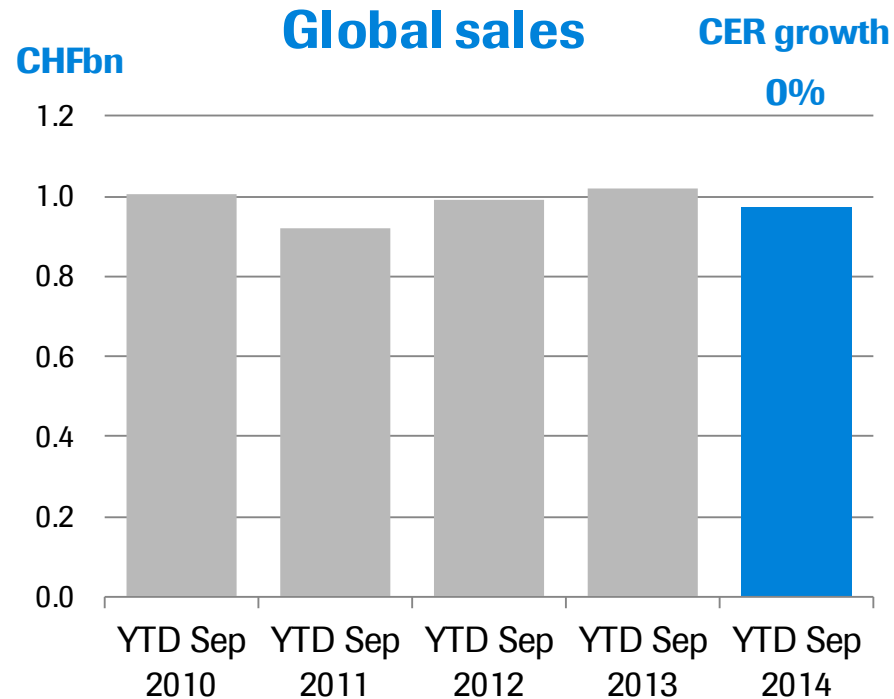
- Growth was driven by continued strong uptake of Perjeta in 1/2L mBC and in the neoadjuvant setting (US), as well as by Kadcyła in 2L mBC
- Continued strong growth in Herceptin benefiting from higher volumes / prolonged treatment times



YTD Sept 2014 sales of CHF 0.623bn

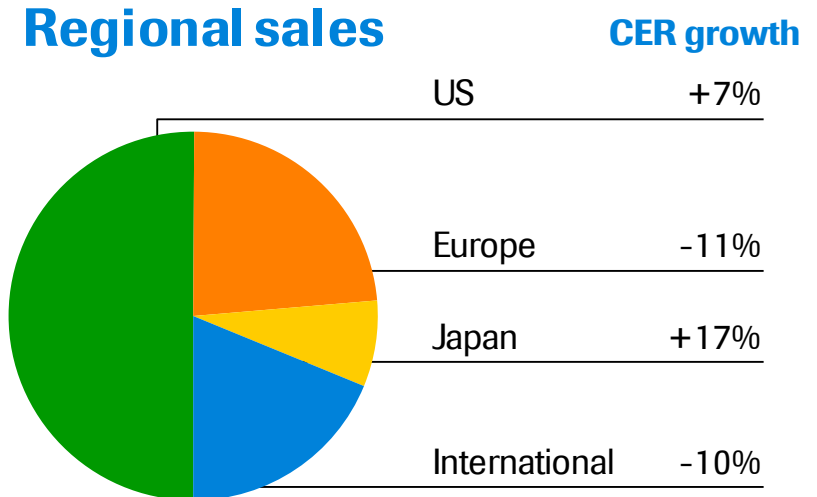
- US: LoE impacts with loss of exclusivity in Feb 2014
- Europe: LoE impacts with loss of exclusivity in Dec 2013

Tarceva

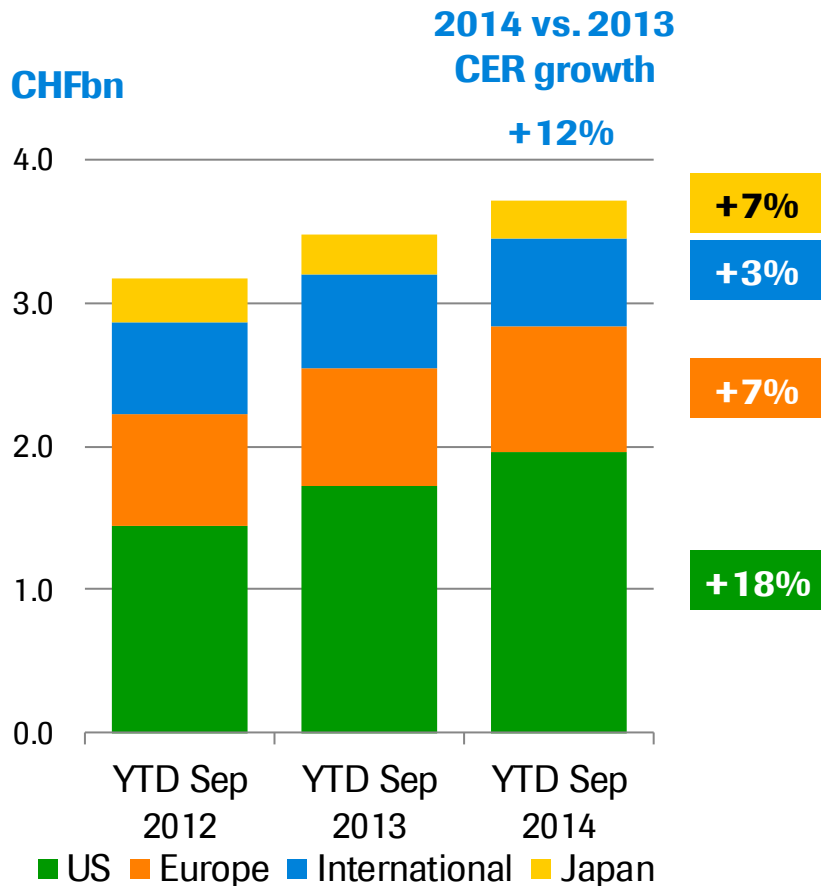


YTD Sept 2014 sales of CHF 0.971bn

- US: Increased use in 1L EGFR Mut+
- Europe: Increased competition in 1L EGFR Mut+ market
- International: local competition in China



YTD Sept 2014: Immunology



YTD Sept 2014 sales of CHF 3.715bn

- Strong growth of Actemra/RoActemra in all regions and MabThera/Rituxan in Europe & International, CellCept declining

Actemra/RoActemra

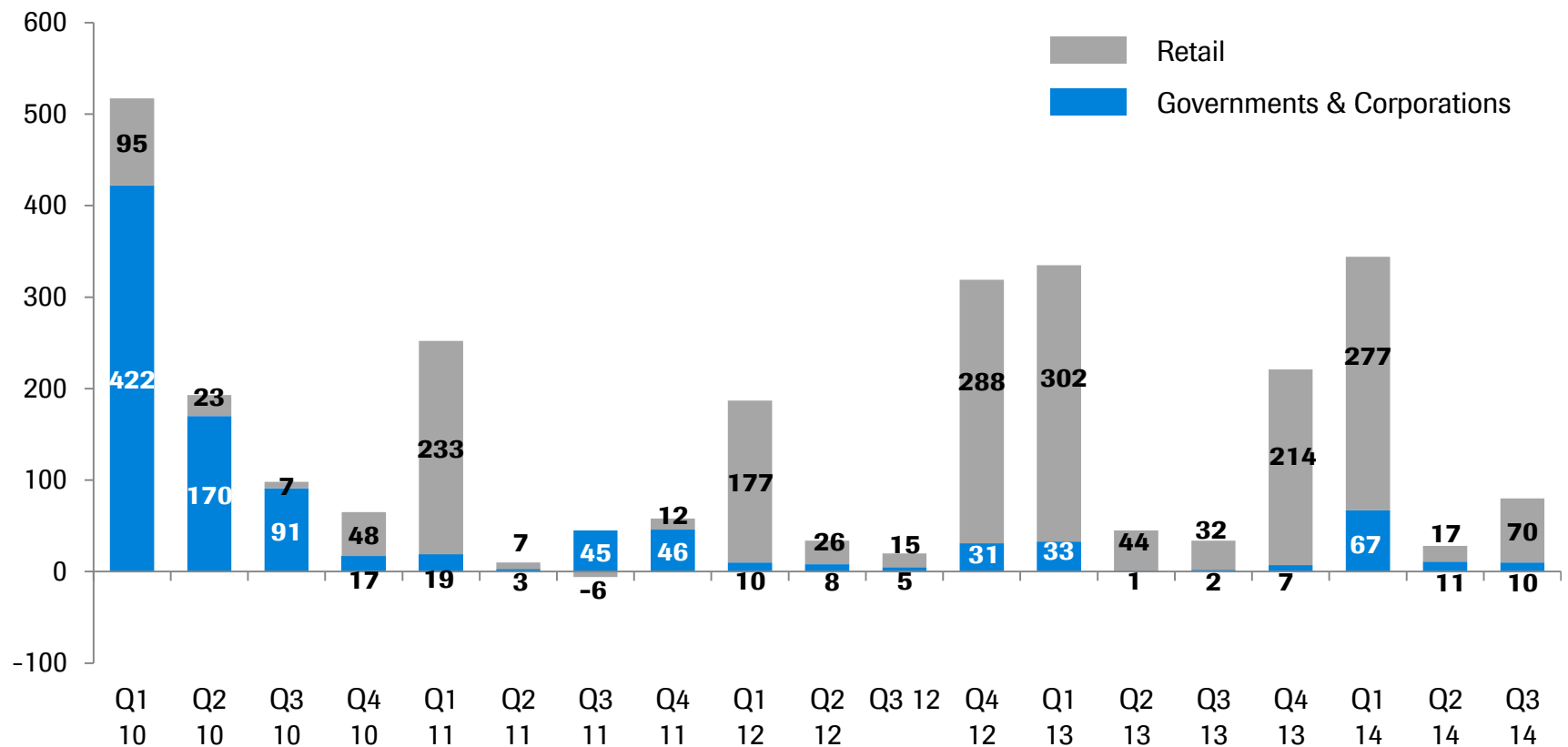
- EU: Continued growth driven by a strong mono patient shares in all lines, very encouraging SC
- US: Growth is driven by strong IV demand and SC patient share uptake (~80% of new patients)

CellCept

- Decline of sales due to patent expiry in EU end 2010 (YTD Sept CHF 623m, -4%)

Tamiflu quarterly sales 2010 - 2014

Retail and Governments/Corporations



Pipeline summary

Marketed products additional indications

Global Development late-stage trials

pRED (Roche Pharma Research & Early Development)

gRED (Genentech Research & Early Development)

Roche Group YTD Sept 2014 sales

Diagnostics

Foreign exchange rate information

Diagnostics Division CER growth

By Region and Business Area (vs. 2013)

	Global		North America		EMEA ¹		RoW	
	% CER		% CER		% CER		% CER	
	CHFm	growth	CHFm	growth	CHFm	growth	CHFm	growth
Professional Diagnostics	4,397	8	897	7	1,920	4	1,580	15
Diabetes Care	1,721	1	320	-4	1,065	1	336	7
Molecular Diagnostics	1,165	5	417	9	458	2	290	4
Tissue Diagnostics	509	10	295	6	143	16	71	18
Diagnostics Division	7,792	6	1,929	6	3,586	3	2,277	12

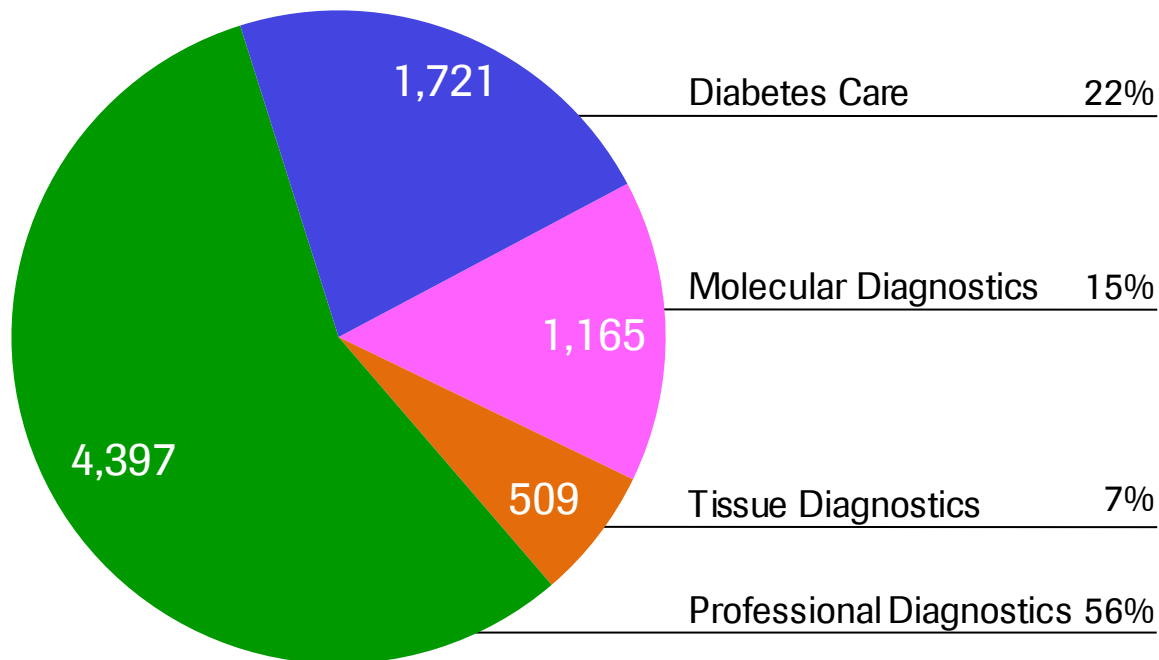
Diagnostics Division quarterly sales and CER growth¹

	Q2 13		Q3 13		Q4 13		Q1 14		Q2 14		Q3 14	
	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER	CHFm	% CER
Professional Diagnostics	1,480	8	1,426	9	1,521	10	1,392	9	1,512	8	1,493	8
Diabetes Care	666	-4	576	3	678	-4	538	5	602	-4	581	4
Molecular Diagnostics	403	6	383	4	416	3	370	4	392	3	403	8
Tissue Diagnostics	165	4	159	8	184	10	156	4	178	14	175	13
Dia Division	2,714	4	2,544	7	2,799	5	2,456	7	2,684	5	2,652	7

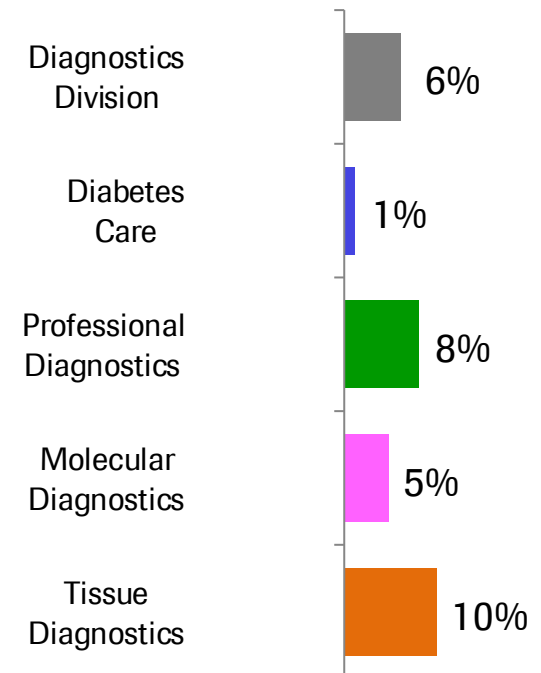
YTD Sept 2014: Diagnostics Division sales

Growth driven by Professional Diagnostics

CHF 7,792m



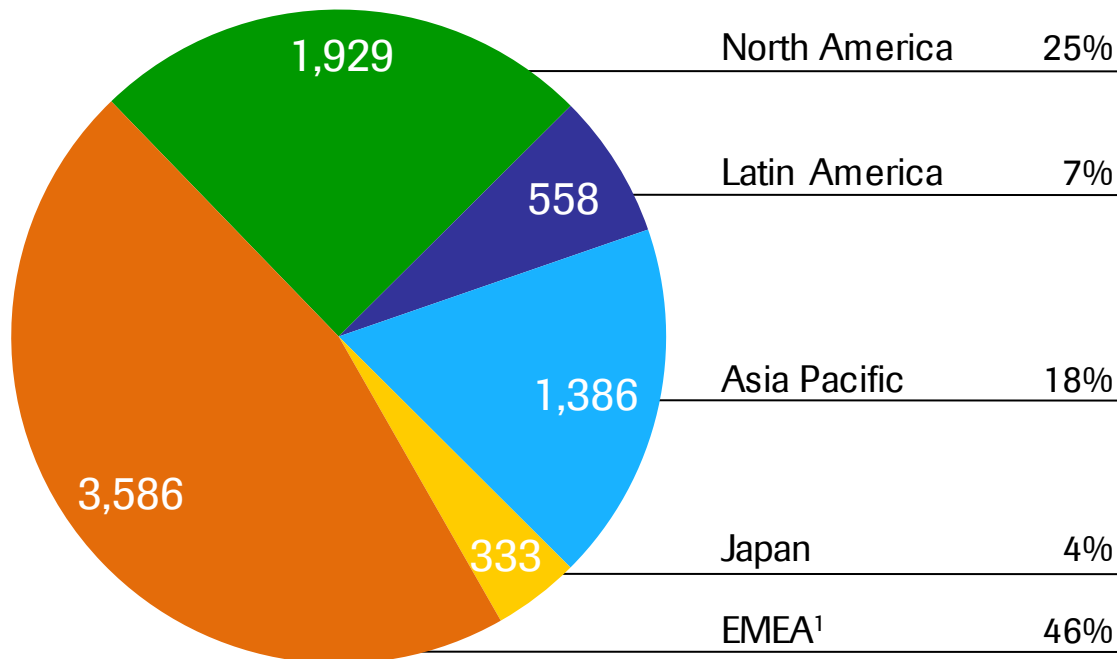
CER sales growth



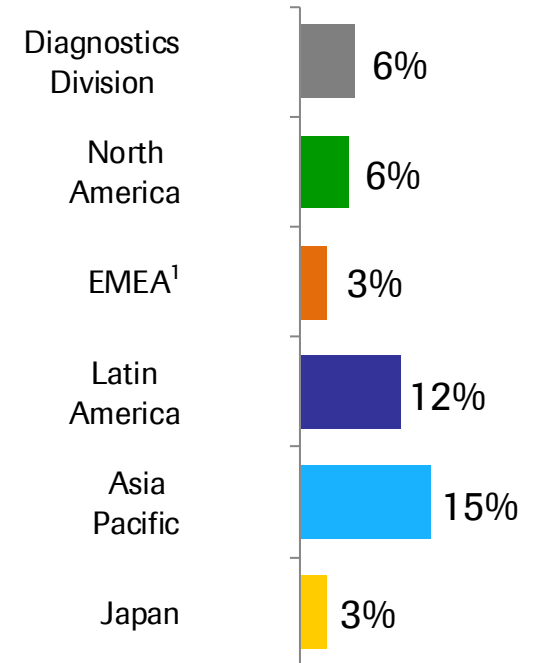
YTD Sept 2014: Diagnostics Division sales

Growth driven by Asia Pacific

CHF 7,792 m

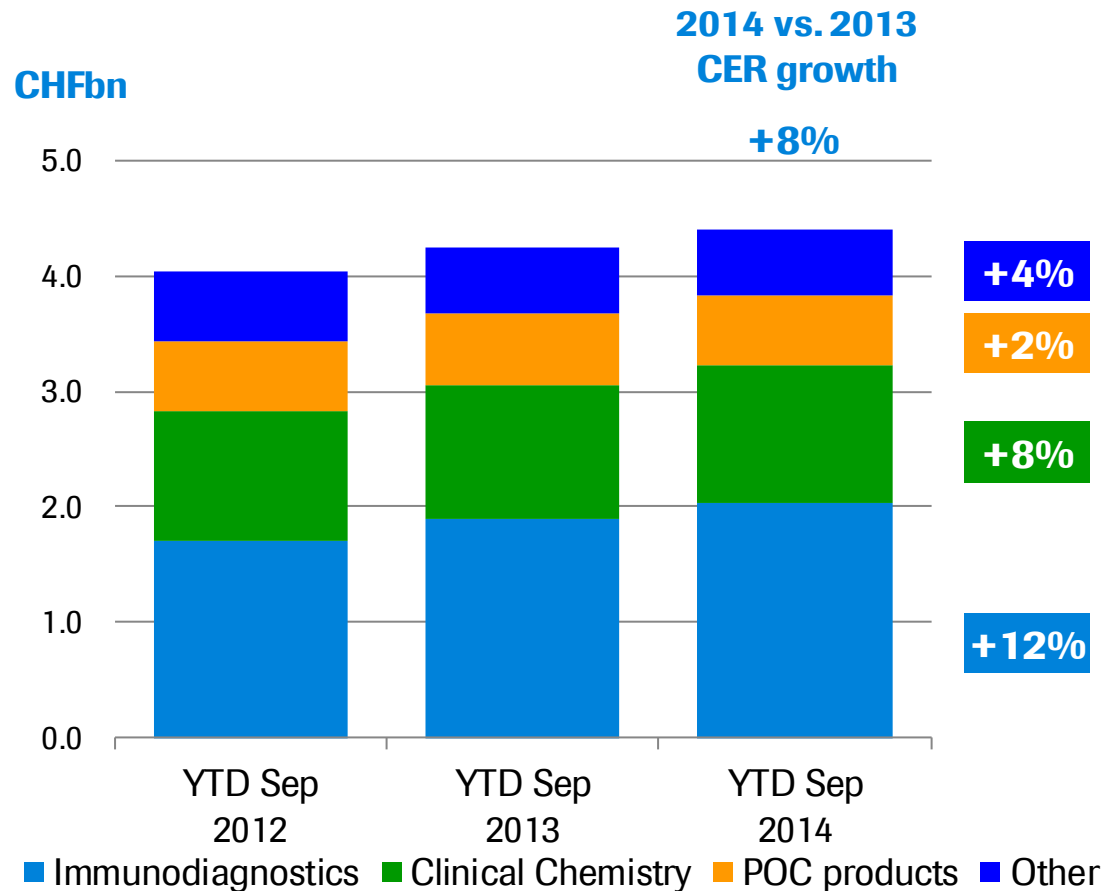


CER sales growth



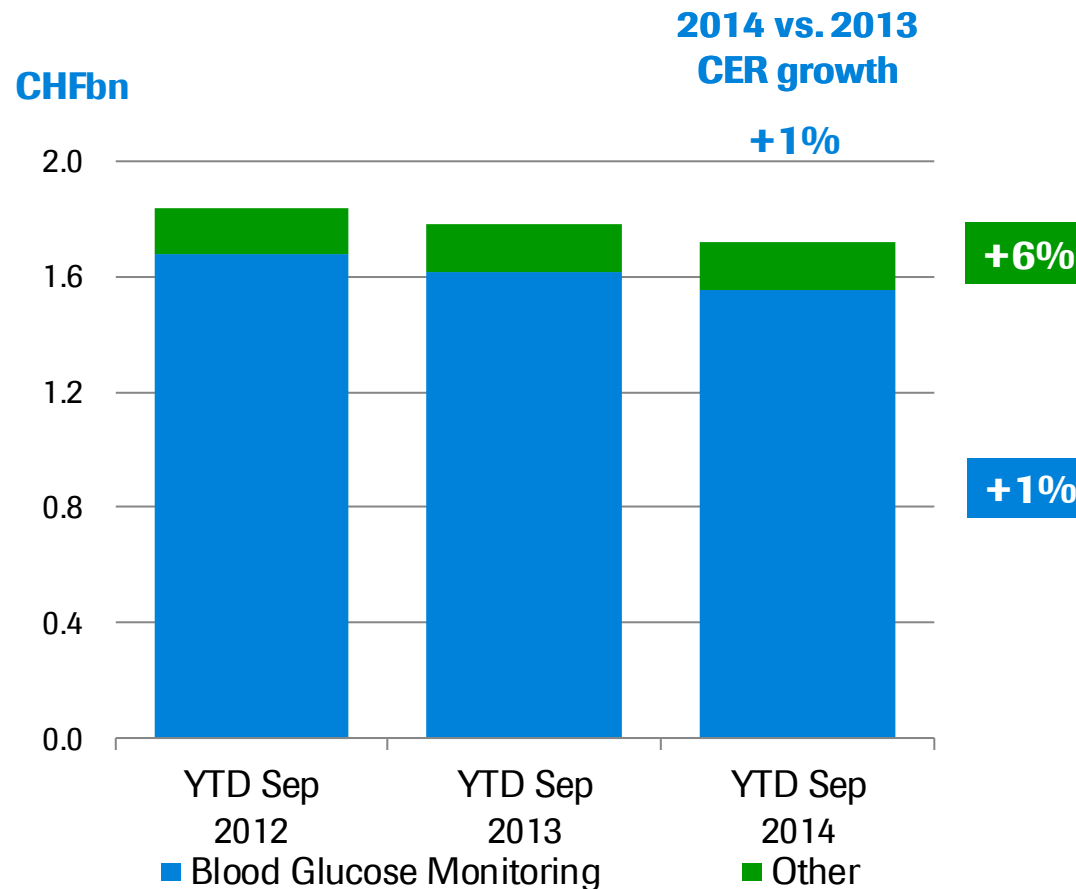
Professional Diagnostics

Strong growth driven by Immunodiagnostics



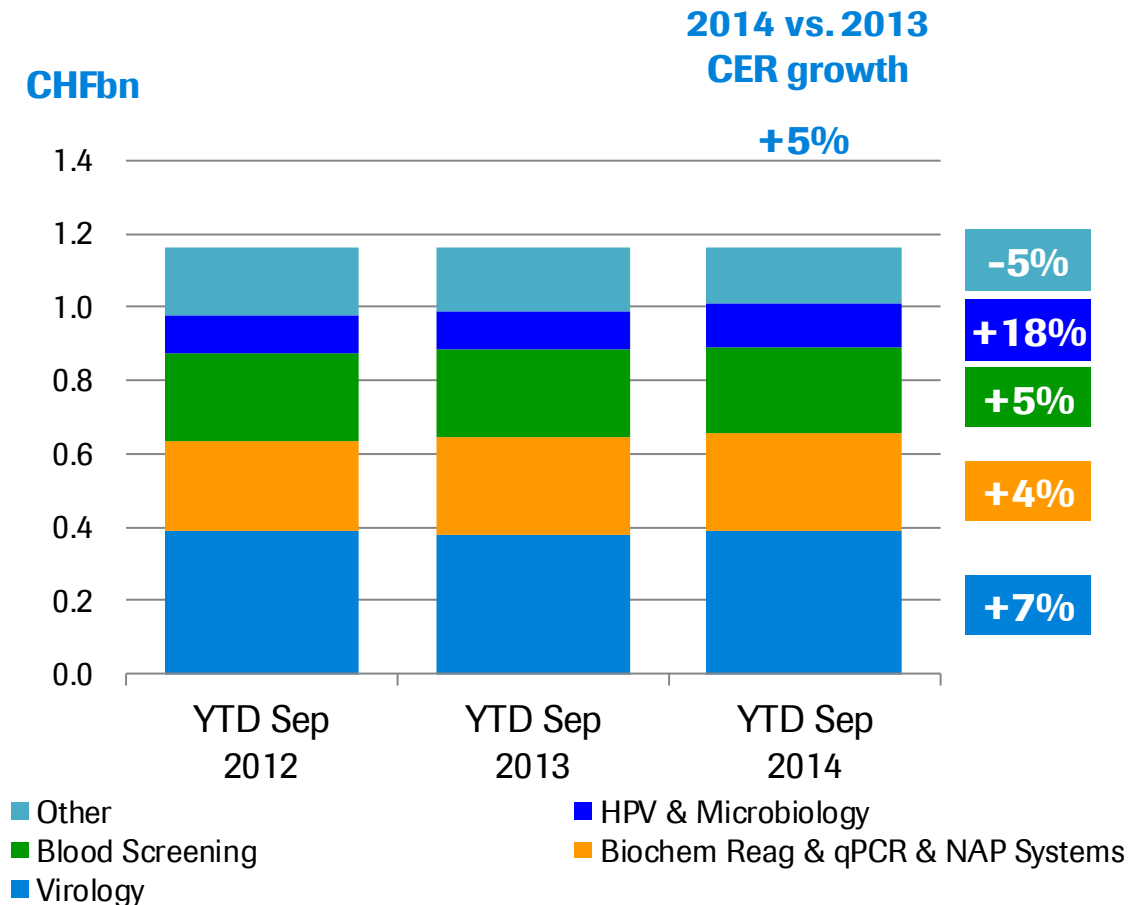
Diabetes Care

Adapting to a challenging market environment



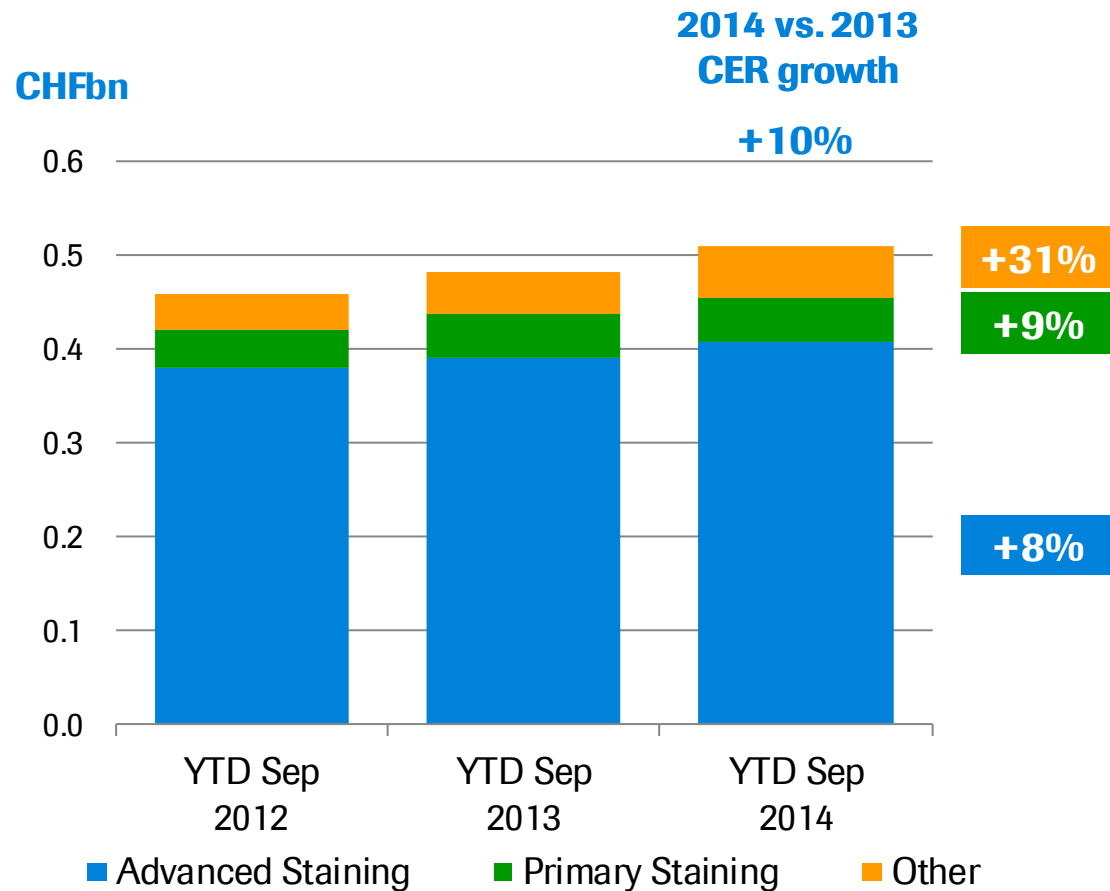
Molecular Diagnostics

Growth driven by Virology and Blood Screening



Tissue Diagnostics

Strong growth in EMEA¹ and North America



2014: Key product launches

Professional Diagnostics

Product	Description	Region
cobas m 511	Fully integrated and automated Hematology system	EU
cobas 6500 (u 701)	Automated urinalysis work area platform including u701 microscopy analyzer	EU ✓
Syphilis	Immunoassay for the detection of <i>Treponema pallidum</i>	EU ✓
PE Prognosis	Claim extension for short-term prediction, rule in/out of Preeclampsia in pregnancy	EU
Anti Mullerian Hormone	Fully automated test for the assessment of ovarian reserve for fertility	EU ✓

2014: Key product launches

Diabetes Care

Product	Description	Region
Accu-Chek Connect	bG meter that connects wirelessly via Bluetooth to a smartphone app and cloud to transmit bG values	EU ✓
Accu-Chek Insight	Next generation insulin delivery system combining an insulin pump and a blood glucose meter that functions as a pump remote control	EU ✓

2014: Key product launches

Molecular Diagnostics

Product	Description	Region
cobas 6800/8800	Next generation PCR platform for molecular testing in virology and blood screening, serving mid to high volumes	WW* ✓
MPX 2.0	Next generation multiplex test for blood screening for HIV, HCV and HBV	US
HSV 1 and 2 test	Detection of Herpes Simplex Virus on cobas 4800 platform	EU ✓
MRSA/SA test	Detection of MRSA/SA on cobas 4800	EU ✓
C-difficile test	Detection of C-difficile on cobas 4800	US ✓

* excluding US

Planned launches may be delayed or not occur as a result of adverse regulatory decisions or other factors

2014: Key product launches

Tissue Diagnostics

Product	Description	Region
Connect-V	Middleware providing connectivity for RTD instruments to simplify interfacing and connectivity to laboratory and hospital information systems	WW

Pipeline summary

Marketed products additional indications

Global Development late-stage trials

pRED (Roche Pharma Research & Early Development)

gRED (Genentech Research & Early Development)

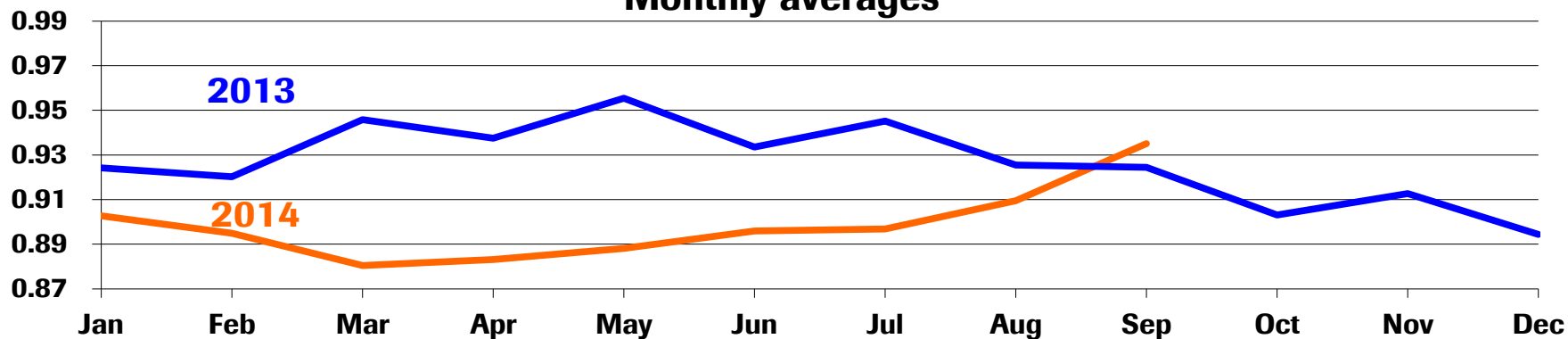
Roche Group YTD Sept 2014 sales

Diagnostics

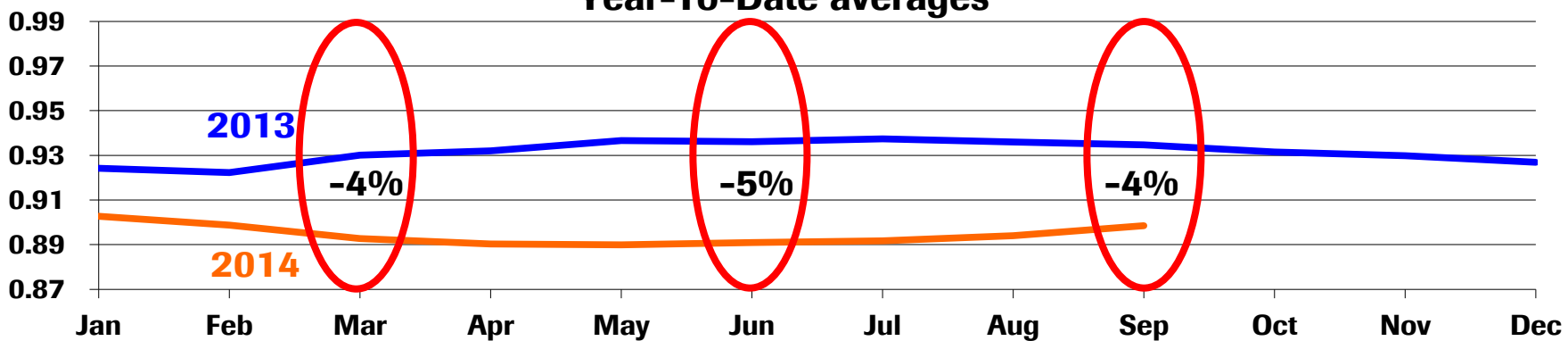
Foreign exchange rate information

CHF / USD

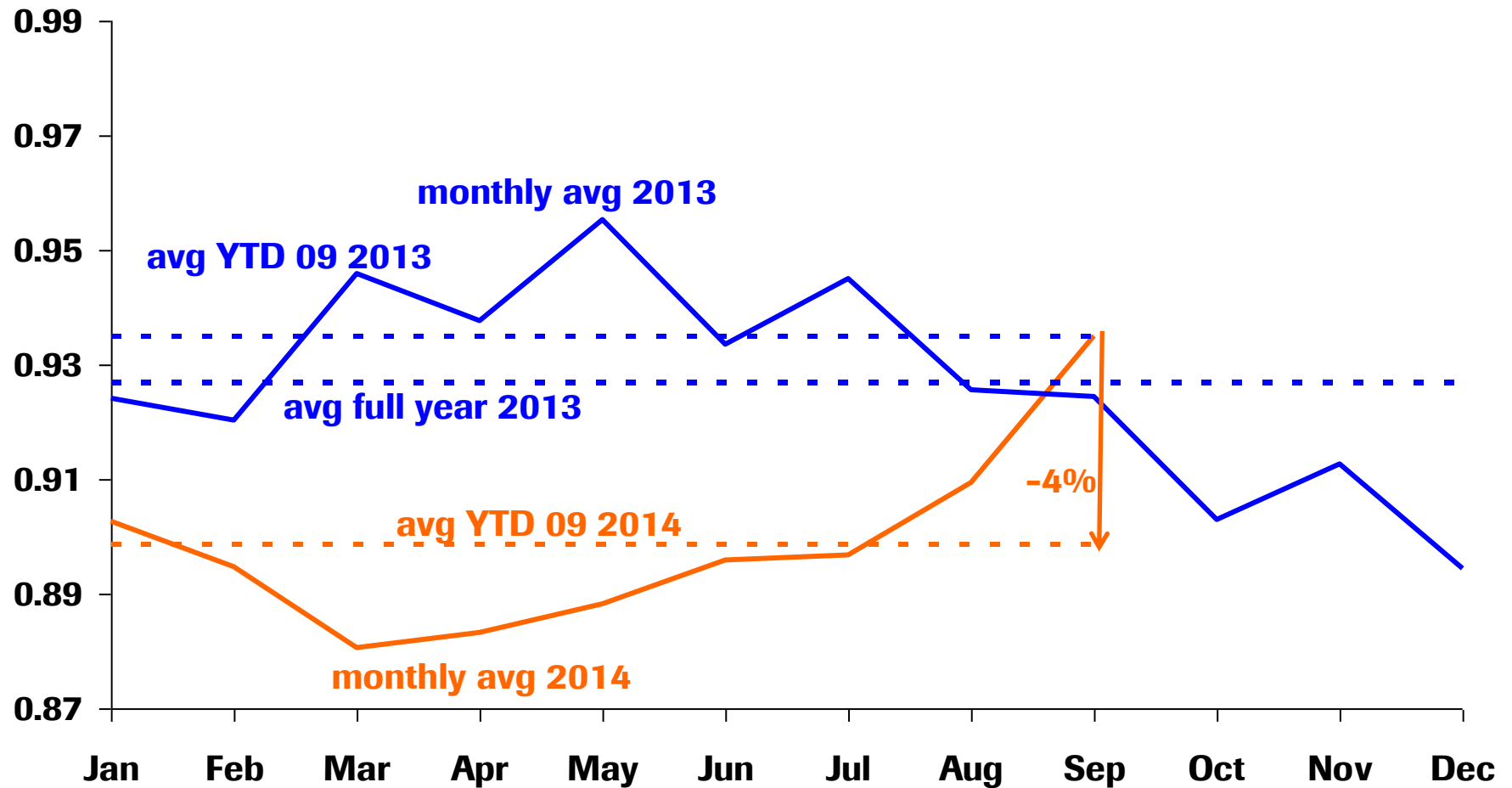
Monthly averages



Year-To-Date averages

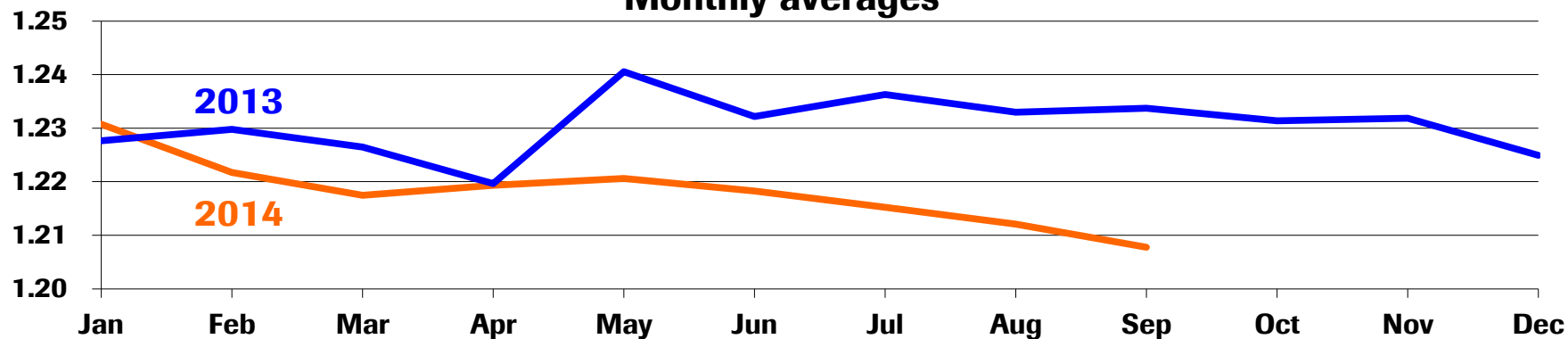


CHF / USD

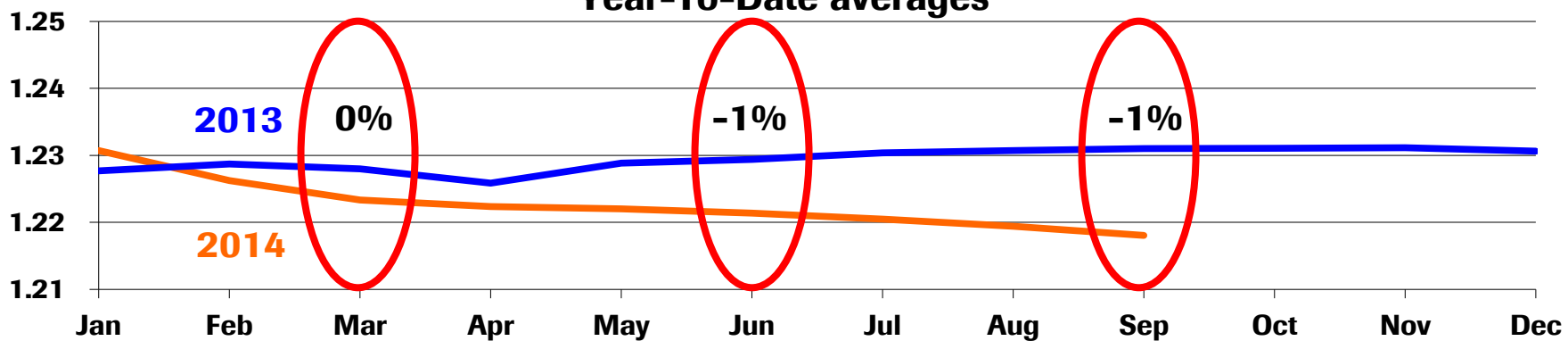


CHF / EUR

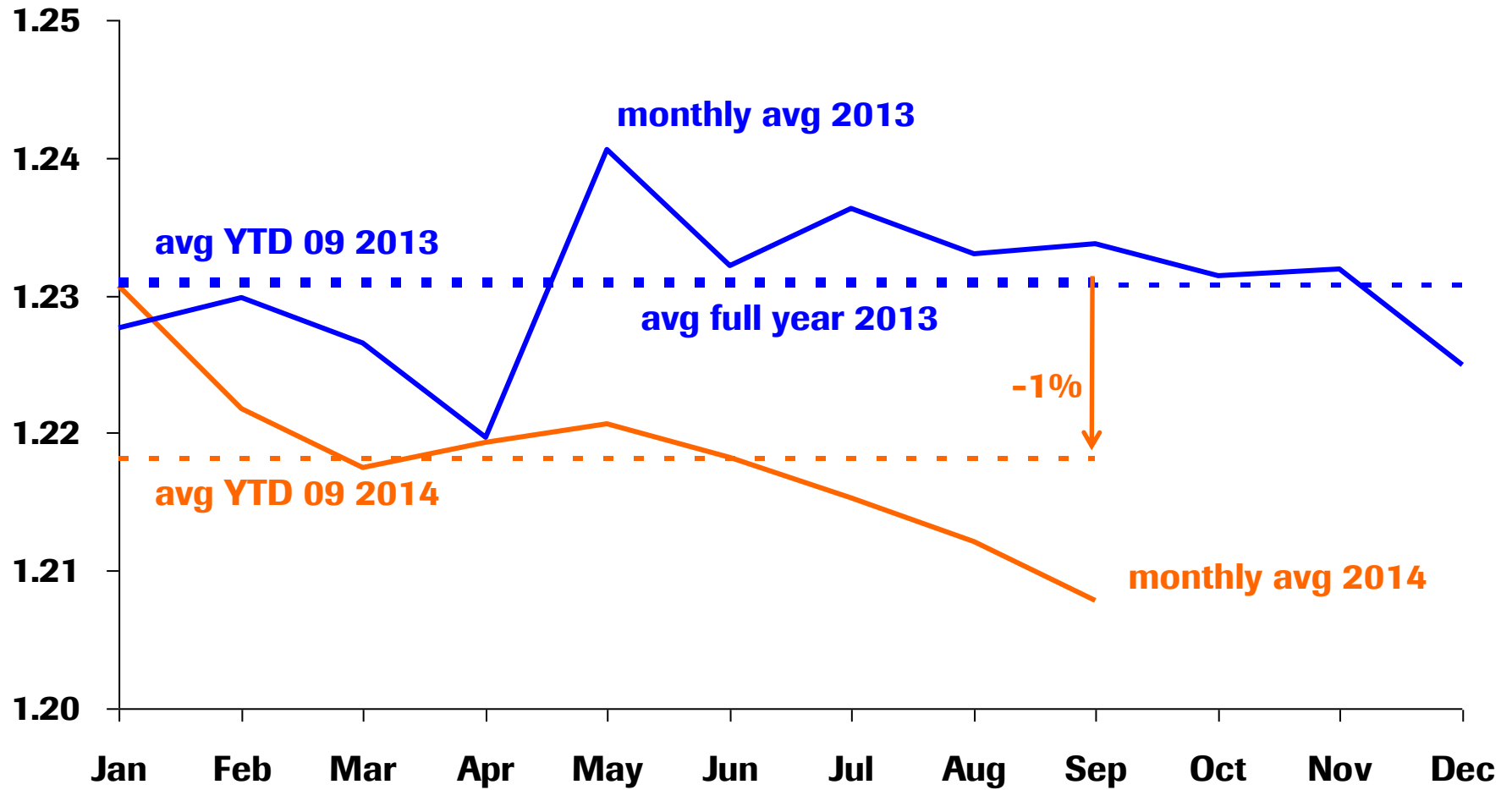
Monthly averages



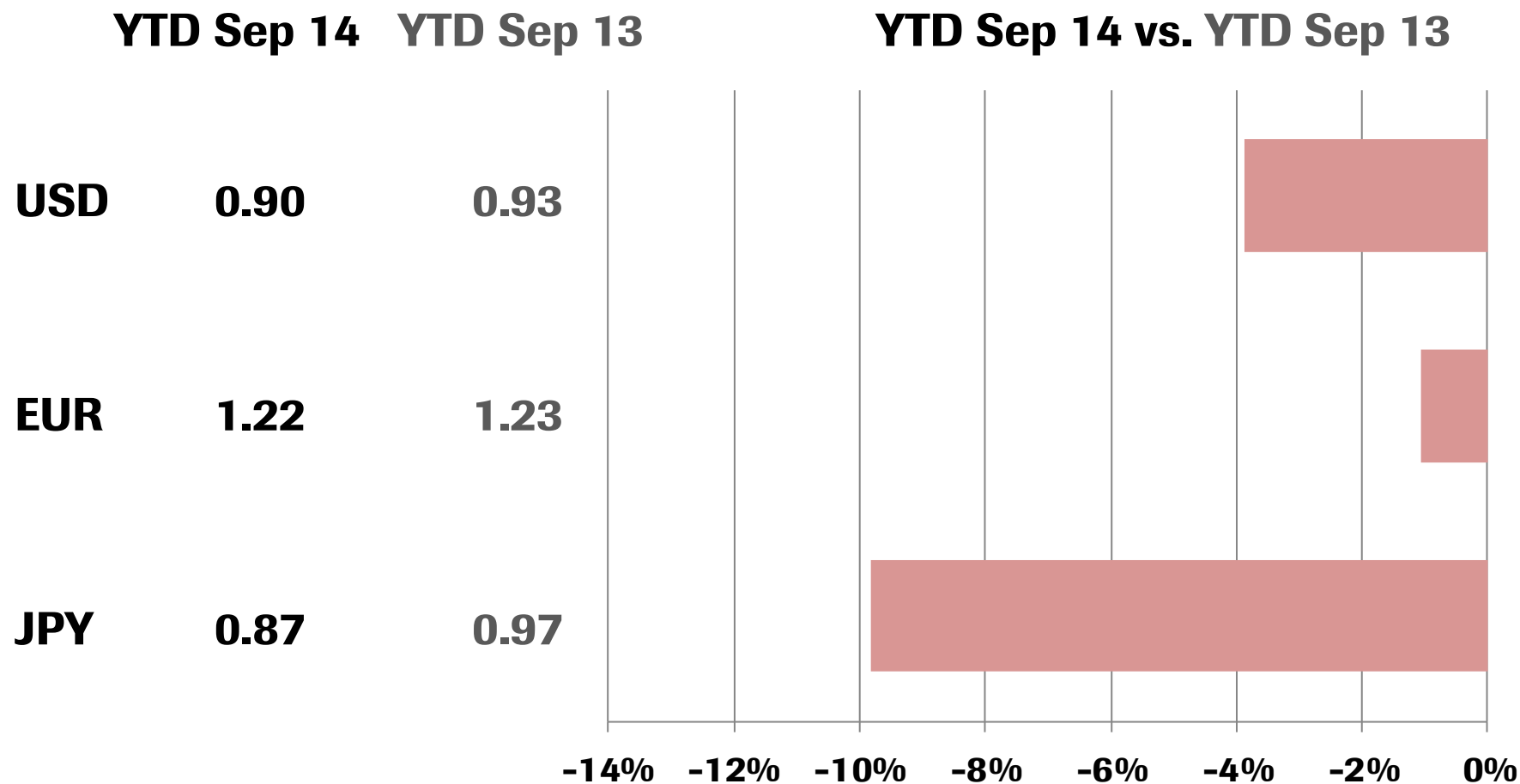
Year-To-Date averages



CHF / EUR



Average exchange rates



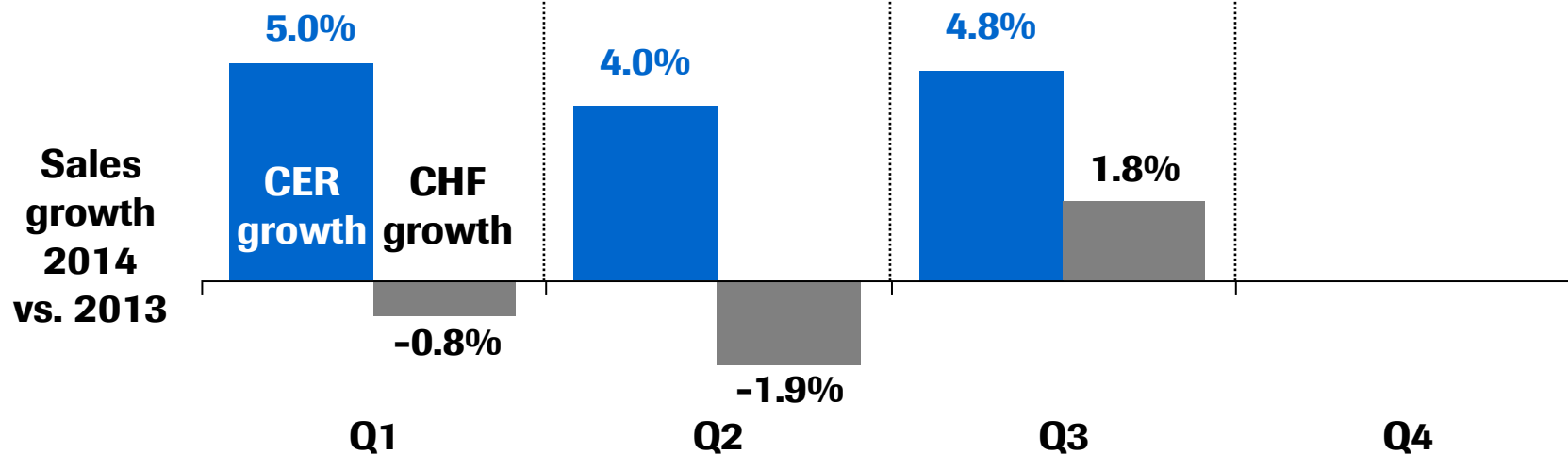
Exchange rate impact on sales growth

In Q3 2014 negative impact from USD, Latin American currencies and JPY

Average exchange rates versus prior year period

CHF / EUR	-0.4%	-0.9%	-1.9%
CHF / USD	-4.0%	-5.6%	-1.9%
CHF / JPY	-13.9%	-8.9%	-6.6%

Difference
in CHF / CER
growth



CER = Constant Exchange Rates

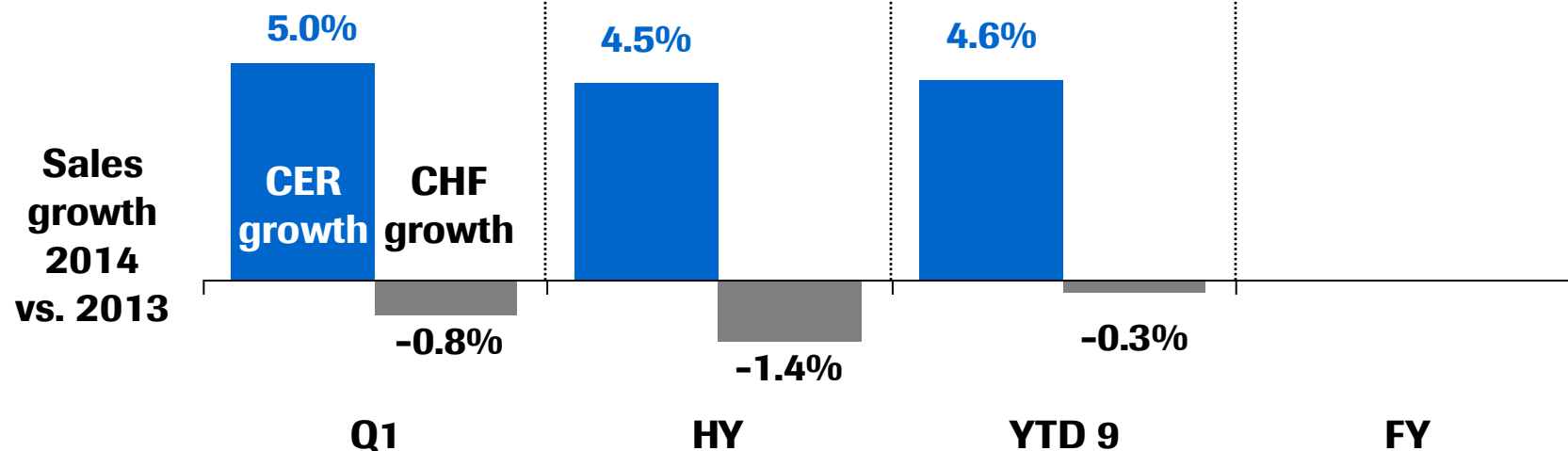
Exchange rate impact on sales growth

In YTD September 2014 negative impact from USD, Latin American currencies and JPY

Average exchange rates versus prior year period

CHF / EUR	-0.4%	-0.7%	-1.1%
CHF / USD	-4.0%	-4.8%	-3.9%
CHF / JPY	-13.9%	-11.4%	-9.8%

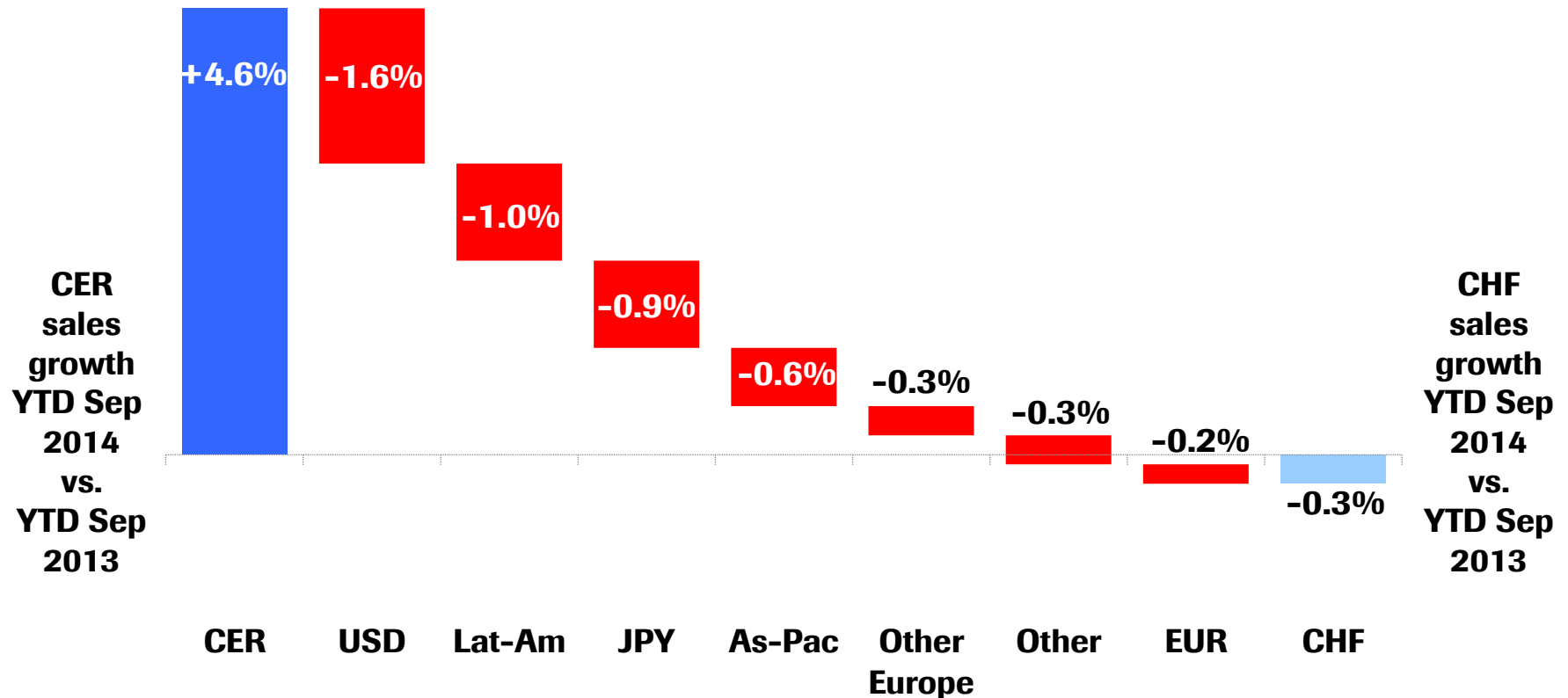
Difference
in CHF / CER
growth



CER = Constant Exchange Rates (avg full year 2013)

Exchange rate impact on sales growth

Negative impact from USD, Latin American currencies and JPY



Doing now what patients need next