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**Roche: Committed to Innovation**

*William M. Burns, CEO Roche Pharmaceuticals*



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## Roche: Committed to innovation

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### Growth opportunities

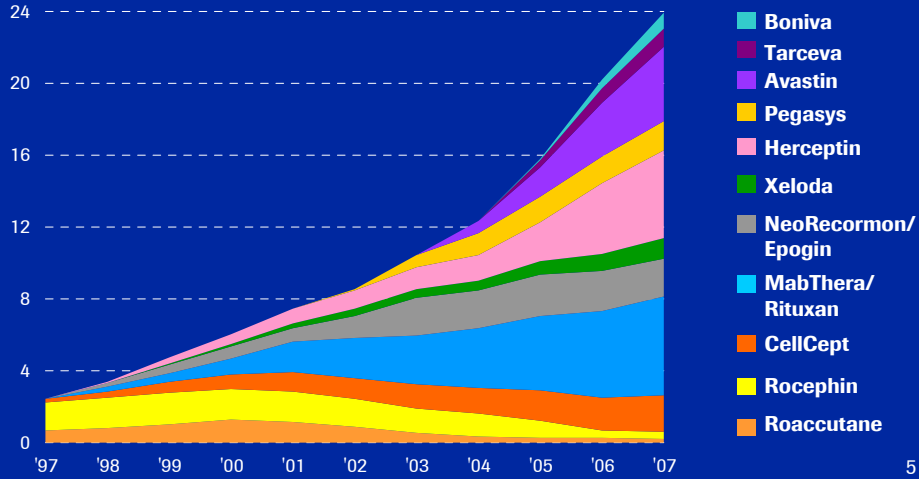
### Summary

## Differentiated and rejuvenated product portfolio

From 1 to 9 products with sales at or above CHF 1 billion



Pharmaceuticals key products (CHF billion)  
By year of market introduction

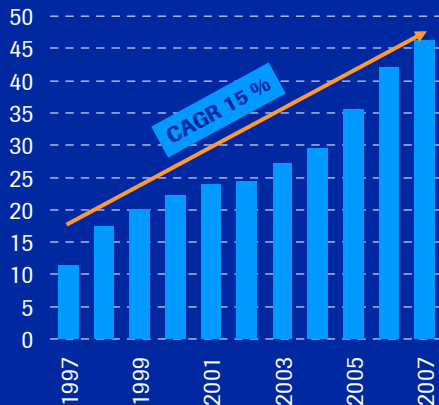


## Focus on differentiated products paying off

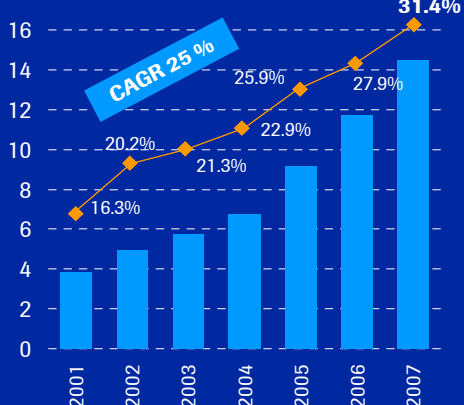
Outstanding long-term value creation



Group sales<sup>1</sup> (CHF billion)



Group operating profit<sup>2</sup> (CHF billion)



Continuing to focus on our core assets

<sup>1</sup> Prescription and Diagnostics

<sup>2</sup> Continuing businesses, before exceptional items

# New pharma model: Disease Biology Areas (DBAs)

*Alignment and focus*



## Disease Biology Areas

Idea ←————→ Market



- Clear focus
- More independent and flexible disease areas
- Faster and simpler decision processes

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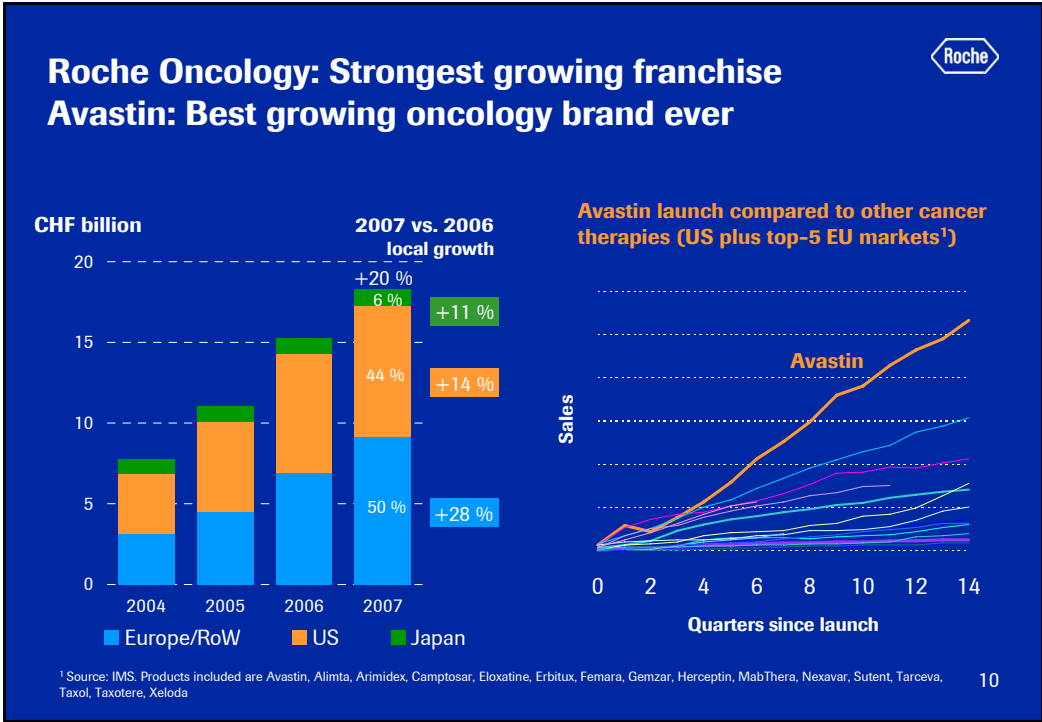
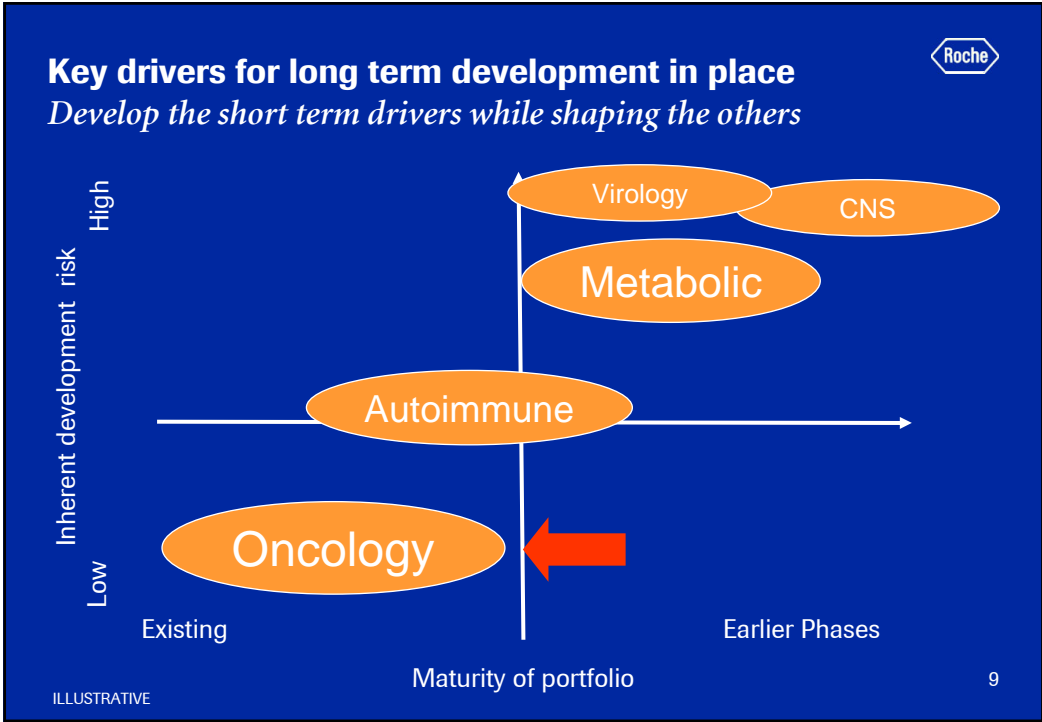
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Growth opportunities

Summary

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# Avastin still early in its journey

## Realizing full potential across tumor types



Tumour	Early/adjvant (Potential for cure)	Advanced/metastatic (Extending life)	
		1 <sup>st</sup> -line of treatment	2 <sup>nd</sup> -line of treatment
Colon/ rectal	Phase III (AVANT, NSABP C-08, E5202, E5204)	Launched [EU, US, JP; broad label in 1st and subsequent lines]	
Lung (NSCLC)	Phase III (E1505)	Launched [EU majority of chemos, US carboplatin/paclitaxel]	Phase III (BETA Lung w/Tarceva)
Breast (HER2-)	Phase III (BEATRICE, E5103)	Launched [EU paclitaxel] Phase III (AVADO, RIBBON-1)	Phase III (RIBBON-2, incl. w/Xeloda)
Breast (HER2+)	Phase III (BETH w/Herceptin)	Phase III (AVEREL w/Herceptin)	-
Kidney (RCC)	-	Launched [EU; with interferon]	

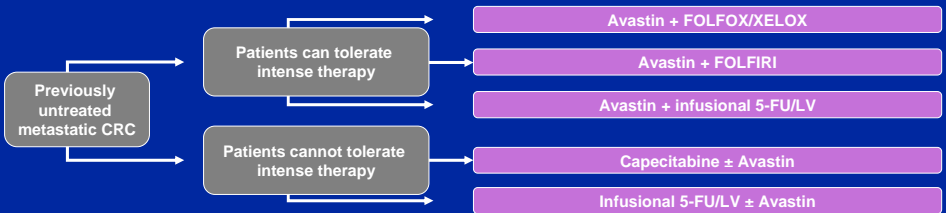
Avastin also trialed in gastric, ovarian, prostate, aNHL, and brain (GBM)

(Trial names) [Approval status]. More trials are ongoing than listed above.

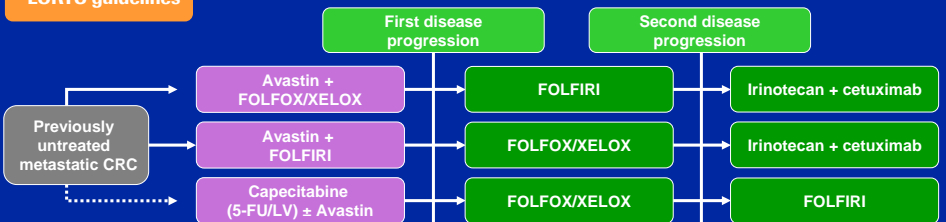
# Avastin recommended first line in major treatment guidelines



### NCCN guidelines



### EORTC guidelines



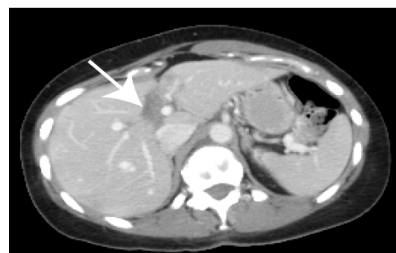
## New market opportunities: Trastuzumab-DM1

*Very promising early data*

- (24 pts evaluated, Phase 1)
- 6 objective responses, 4 responses on-going at the last data cut-off; the longest has persisted over 8 months
- In addition to trastuzumab, all responders previously received taxane &/or vinca alkaloid
- No unexpected cardiotoxicity has been observed so far

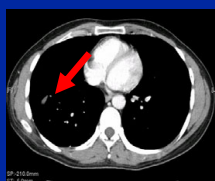
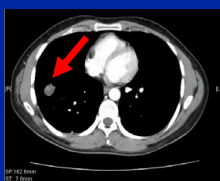
CT Liver Scan Image at Baseline

CT Liver Scan Image at End of Cycle 2



Krop et al, ECCO 2007

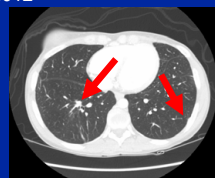
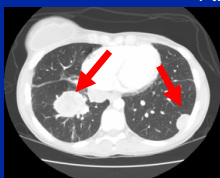
## New market opportunities: IGF1-R Inhibitor – Impressive results in Phase I- Eligibility as multi-tumor compound?



Baseline June 19, 2006

Restaging Week 25  
Dec 29, 2006

Pt. 8012



Baseline Dec 8, 2006

Restaging Week 6  
Jan 25, 2007

### Unique Features

- Selective to IGF pathway which is a key factor in tumor growth

### Drivers for Value

- IGF pathway linked to many tumor types
- Speed: Sarcoma collaborative groups to allow exclusive focused trial and rapid market access

## Oncology in 2008



### *Preparing for new market opportunities*

#### Breast cancer (BC)

- Phase III data (AVADO, RIBBON-1) to broaden Avastin label for combination with all major chemos
- HER2+ mBC
  - phase III for pertuzumab started in Q1 2008
  - opted-in to trastuzumab-DM1 (in phase II)
- Avastin adjuvant trials started: Large potential new market opportunity

#### Metastatic colorectal cancer (mCRC)

- Launch of Avastin and Xeloda in 1st and later lines – not restricted by chemo choice (incl. oxaliplatin) (EU)

#### Chronic lymphocytic leukemia (CLL)

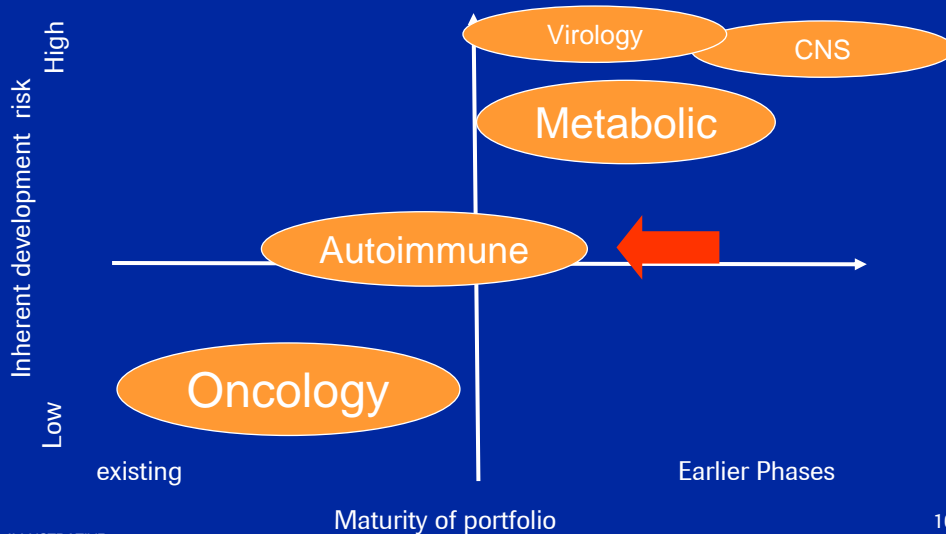
- Positive data in 1st line - MabThera to enter a new market in Europe

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## Key drivers for long term development in place



*Develop the short term drivers while shaping the others*



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## Overview of autoimmune disorders



INDICATION	PRIMARY PATHOLOGY	HIGH NEED?	PREVALENCE	PATIENTS	DISEASE PROGRESSION
<b>Rheumatoid Arthritis</b>	Joint/bone degeneration	Yes	1% global	+60 mln WW	Mild, moderate, severe
<b>Multiple Sclerosis</b>	Destruction of CNS myelin	Yes	<<1% global	0.6 - 1.1 mln WW	RRMS; SPMS; PPMS; PRMS *
<b>Systemic Lupus Erythematsps</b>	Tissue damage from autoantibodies	Yes	<0.5% global	0.5 mln US	Mild, intermittent, persistent, fulminant
<b>Lupus Nephritis</b>	Renal involvement in SLE	Yes	40 - 85% SLE patients	0.2 - 0.4 mln US	Class I - VI
<b>Psoriasis</b>	Autoimmune infiltration into skin	Yes	1 - 2% global	125 mln WW	Plaque-type: Mild, moderate, severe

\* RRMS: Relapsing-Remitting Multiple Sclerosis; SPMS: Secondary-Progressive MS; PPMS: Primary Progressive MS; PRMS: Progressive Relapsing MS<sup>17</sup>

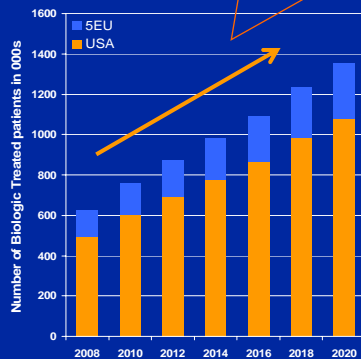
## Rheumatoid Arthritis: Continued Market Growth

*Driven by increased Biologic use, novel oral DMARDs & PHC*



### Short to mid-term growth driver

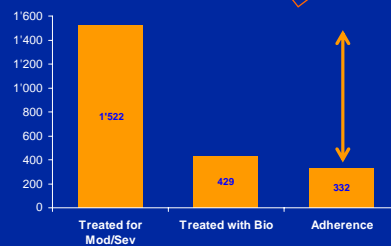
Biologic treated population projected to grow 6.7% CAGR (2008-2020)



### Mid to Long-Term Opportunities

- Low biologic penetration Mod/Sev RA
- Future growth drivers:
  - Novel orals: Potent & convenient
  - PHC defined differentiation = premium pricing
  - New brand Rx combination regimens

2007 US Patient Treatment



## Next generation anti-CD20s (MabThera follow-ons)

### Opportunities for improvement



#### 2nd gen. anti-CD20 (ocrelizumab)

Fully humanized

#### Potential clinical benefits

- Less immunogenicity
- Better tolerability
- Shorter infusion time

#### 3rd gen. anti-CD20 (R7159)

Fc engineered (glycosylations)

- Increased CD20 binding and apoptosis
- Increased ADCC (antibody dependent cell-mediated toxicity)
- Reduced CDC (complement dependent cell toxicity)

#### Potential clinical benefits

- Improved efficacy
- Less infusion reactions

In phase III in RA, and SLE  
LN to start soon  
Phase II in RRMS to start 2008

In phase I for oncology / hematology

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## Inflammation Portfolio

A rich pipeline in multiple AI indications



	LI	LO	P0	P1	P2	P3	Mkt
Rheumatology	nn	nn		nn	Actemra sJIA	Actemra RA	MabThera RA
	nn	nn		nn	Ocrelizumab SLE	Ocrelizumab RA	
	nn	nn				MabThera DMARD inad resp	
	nn	nn				MabThera RA MTX naive	
		nn					
	nn	nn					
Respiratory	nn	nn	nn	nn	RARy		
	nn	nn		hMAb OX40L			
	nn	nn		nn			
Transplant		nn		nn			CellCept
Pain		nn		nn			
Other		nn		S1P1 Psoriasis	ocrelizumab ANCA vasculitis	CellCept lupus nephritis	
					nn	CellCept pemphigus vulgaris	

Project origin: TPI Group Origin External Origin Line Extensions

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## Inflammation/Autoimmune in 2008

*Major market opportunities in focus*



### Rheumatoid Arthritis

- MabThera for DMARD-IRs (1st line biologic): expect data and filing
- Actemra: regulatory approval expected (first in US)

### Multiple Sclerosis

- Phase II/III data for MabThera in PPMS in H1 2008

### Lupus

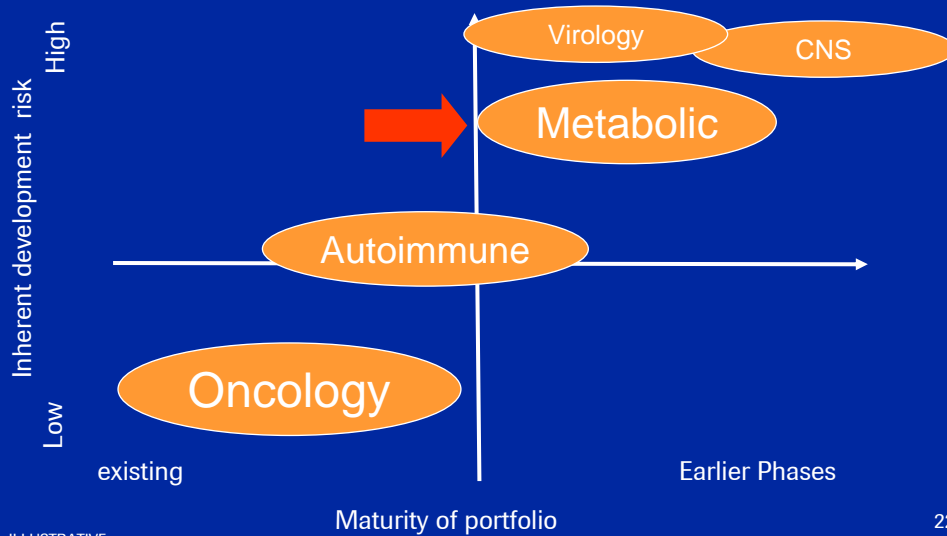
- Phase II / III data for MabThera in SLE in H1 2008

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## Key drivers for long term development in place



*Develop the short term drivers while shaping the others*

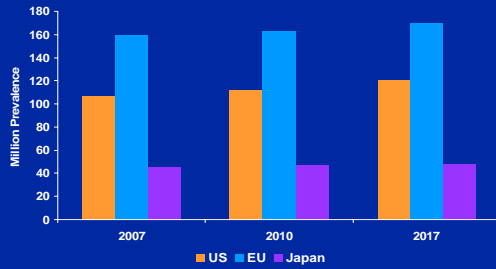


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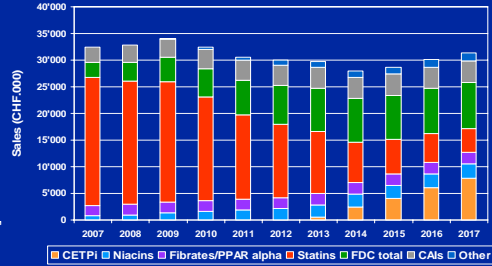
## Dyslipidemia: Future growth driven by HDL treatment



Dyslipidemia Prevalence will Continue to Grow



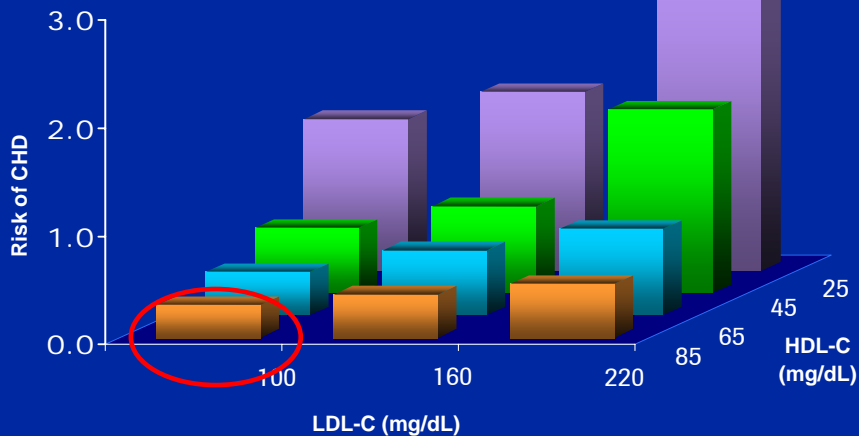
Revenue Development Assumptions (US)



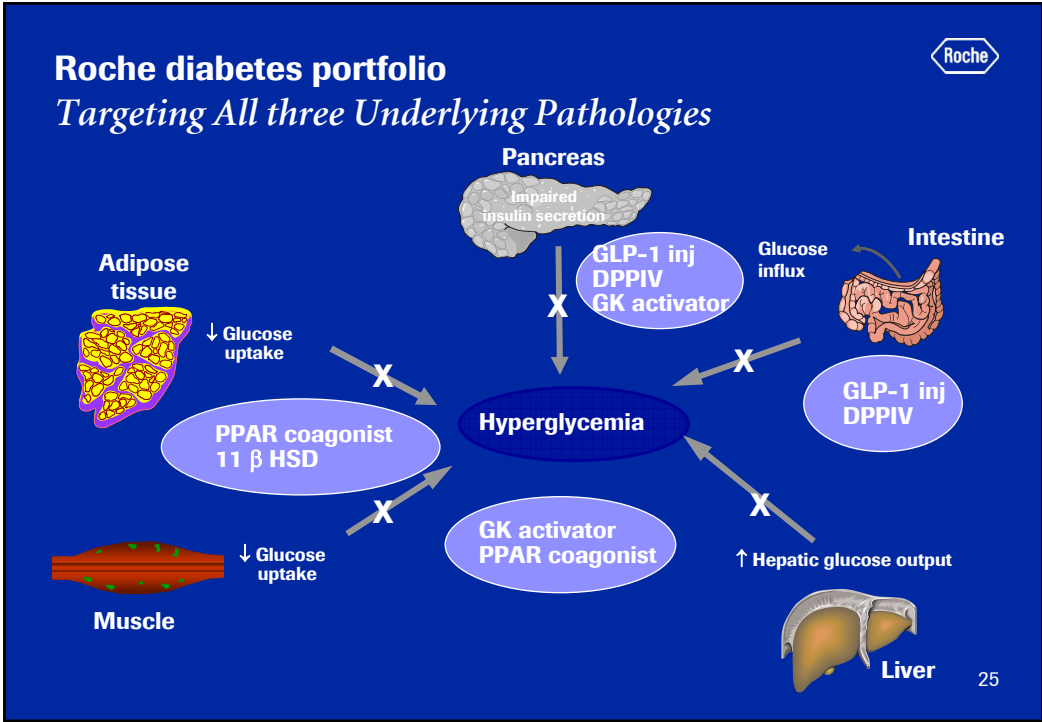
CETP inhibitors and other HDL raising drugs will be responsible for the majority of market value

Sources: Wood MacKenzie, IMS therapy forecaster, in-house assumptions  
Decision Resources, Cardium Study#4, 2007, Datamonitor Pipeline Insight dyslipidemia 2007

## Dyslipidemia: Low HDL-C is an Independent Factor of CHD Risk even when LDL-C is Low



Gordon T et al. Am J Med 1977;62:707-714



## Our metabolic pipeline: maturing quickly

*Committed to become a major player*

	NMP/TA	LI	LO	P0	P1	P2	P3	Reg
Roche	nn	nn	nn	nn	nn	PPAR-CO	CETP	
	nn	nn	nn		nn	GLP-1		
	nn	nn	nn		nn	DPP-IV (3)		
	nn	nn	nn		nn	SGLT2		
	nn	nn	nn		nn			
	nn	nn	nn		nn			
	nn	nn						
	nn	nn						
	nn	nn						
	nn	nn						
	nn	nn						
	nn	nn						
	nn	nn						
	nn	nn						
Chugai	nn	nn			nn			
	nn	nn			SGLT2			
	nn	nn						
	nn	nn						
	nn	nn						
	nn	nn						
Shanghai				nn				

February 04, 2008 26

## Metabolism/Diabetes in 2008

*Committed to become a major player*



### CETP Inhibitor to move into phase III

- Safety data to be presented at major medical meetings

### GLP-1 phase II data available

- Decision for phase III to be taken in H1 2008
- Publication planned for 2008

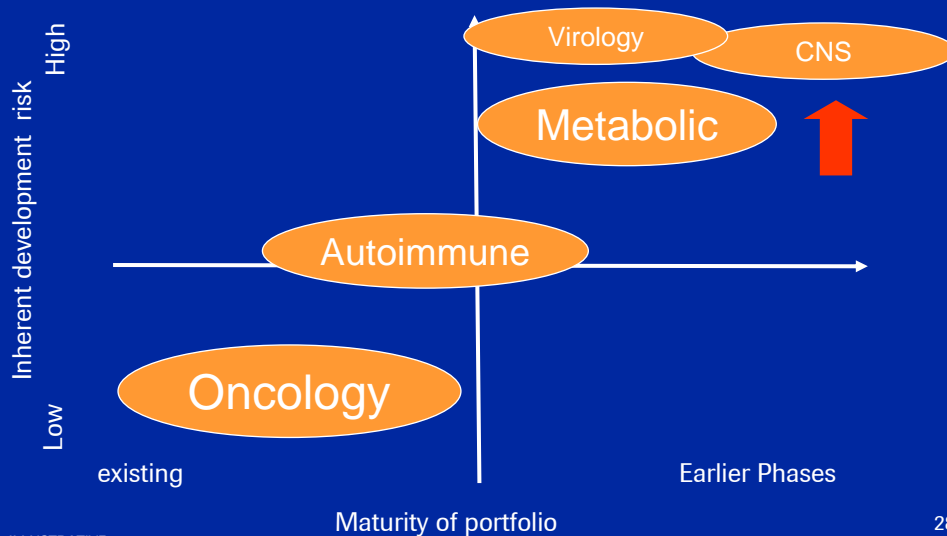
### Phase II data for DPP-IV (3) and PPAR $\alpha$ expected

- Looking for a differentiated profile (e.g. weight reduction for DPP-IV)
- Go/no-go decisions for phase III trials to be taken this year

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## Key drivers for long term development in place

*Develop the short term drivers while shaping the others*



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# CNS; One of the top 3 in the market



*Growth driven by elderly and adolescent patient populations*

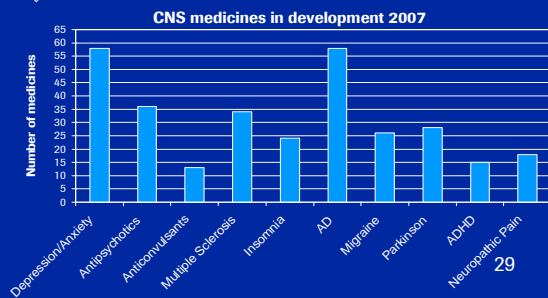
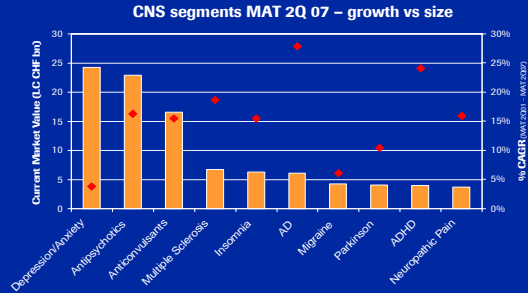
2018 US: Alzheimer expect to rank number 3 after cancer/diabetes

### High unmet medical needs

- Limited efficacy in various indications
- Long term patient benefit
- Need to improve patient functioning

### Diagnosis of limited benefit

- Based on behavior and not etiology
- Biological criteria will play major role for future diagnostic criteria



# CNS Portfolio 2008- A clear commitment



### Development Phase

	TA	LI	LO	P0	P1	P2	P3
Neurology	nn	nn	nn	nn	nn	Nic-a7 mem345	Rituximab
	nn	nn		nn	nn	Ocrelizumab	
					S1P1		
					nn		
Psychiatry	nn	nn	nn		nn		
	nn	nn	nn		GLYT1		
		nn	nn		GLYT1(2)		
		nn	nn				
		nn	nn				
			nn				

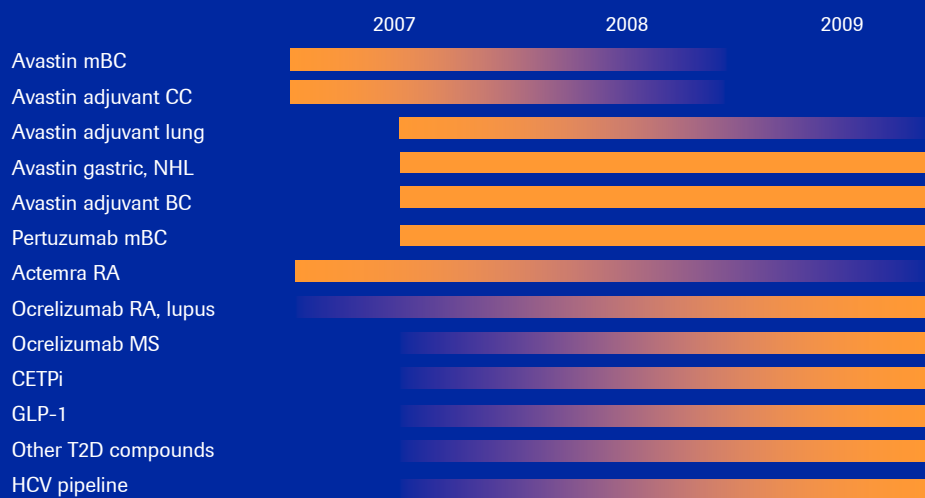
## Roche: Committed to innovation

### Growth opportunities

#### Summary

## Major phase III commitments – large investments

*Additional large phase III trials started or starting soon*

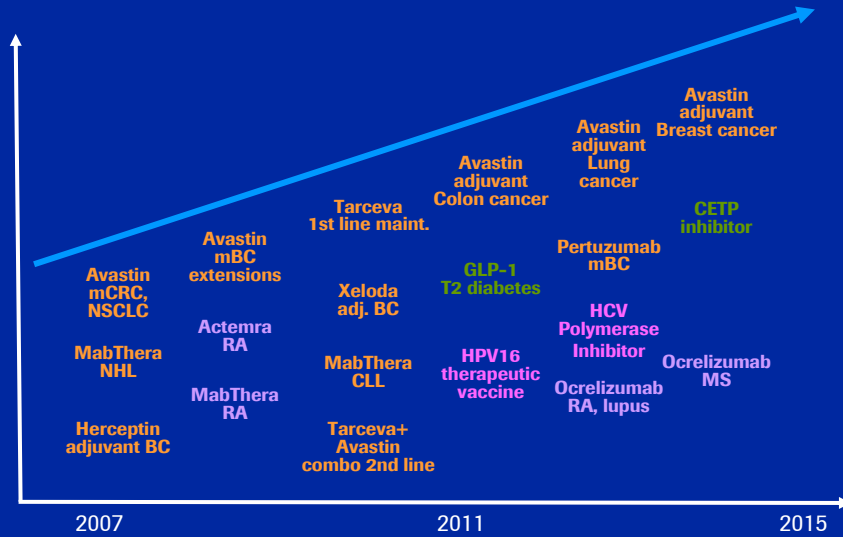




## Key drivers in place for sustainable growth



Current and projected sources of value creation (phase IIb and III)



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## Our objectives for 2008



### Sales

- High single-digit local currency sales increase for Roche Group (excl. Tamiflu pandemic<sup>1</sup>)
- Above-market sales growth<sup>1</sup> in both divisions

### Core EPS

- Core earnings per share target<sup>2</sup> at least at record 2007 level despite significant increase in R&D investment and considerably lower Tamiflu pandemic sales

### Shareholder return

- Continuous increase in dividend pay-out ratio over the next 3 years

<sup>1</sup> Excluding government and corporate stockpiling orders of Tamiflu for pandemic use  
<sup>2</sup> At constant exchange rates

Barring unforeseen events

## Major growth opportunities outside the US



*Herceptin leading the way*

