



Progressing our strategy

Second quarter results to June 30, 2013

Flemming Ornskov, MD
Chief Executive Officer

Graham Hetherington
Chief Financial Officer



Our purpose

We enable people with life-altering conditions to lead better lives.

THE “SAFE HARBOR” STATEMENT UNDER THE PRIVATE SECURITIES LITIGATION REFORM ACT OF 1995

Statements included herein that are not historical facts are forward-looking statements. Such forward-looking statements involve a number of risks and uncertainties and are subject to change at any time. In the event such risks or uncertainties materialize, Shire's results could be materially adversely affected. The risks and uncertainties include, but are not limited to, that:

- Shire's products may not be a commercial success;
- revenues from ADDERALL XR® are subject to generic erosion;
- the failure to obtain and maintain reimbursement, or an adequate level of reimbursement, by third-party payors in a timely manner for Shire's products may impact future revenues and earnings;
- Shire relies on a single source for manufacture of certain of its products and a disruption to the supply chain for those products may result in Shire being unable to continue marketing or developing a product or may result in Shire being unable to do so on a commercially viable basis;
- Shire uses third party manufacturers to manufacture many of its products and is reliant upon third party contractors for certain goods and services, and any inability of these third party manufacturers to manufacture products, or any failure of these third party contractors to provide these goods and services, in each case in accordance with its respective contractual obligations, could adversely affect Shire's ability to manage its manufacturing processes or to operate its business;
- the development, approval and manufacturing of Shire's products is subject to extensive oversight by various regulatory agencies and regulatory approvals or interventions associated with changes to manufacturing sites, ingredients or manufacturing processes could lead to significant delays, increase in operating costs, lost product sales, an interruption of research activities or the delay of new product launches;
- the actions of certain customers could affect Shire's ability to sell or market products profitably and fluctuations in buying or distribution patterns by such customers could adversely impact Shire's revenues, financial conditions or results of operations;
- investigations or enforcement action by regulatory authorities or law enforcement agencies relating to Shire's activities in the highly regulated markets in which it operates may result in the distraction of senior management, significant legal costs and the payment of substantial compensation or fines;
- adverse outcomes in legal matters and other disputes, including Shire's ability to obtain, maintain, enforce and defend patents and other intellectual property rights required for its business, could have a material adverse effect on Shire's revenues, financial condition or results of operations;

and other risks and uncertainties detailed from time to time in Shire's filings with the U.S. Securities and Exchange Commission, including those risks outlined in “Item 1A: Risk Factors” in Shire's Form 10-K for the year ended December 31, 2012.



To be as brave as the people we help.

Agenda

Good performance and strategy progression



**Fleming
Ornskov, MD**

Financial review and 2013 outlook



**Graham
Hetherington**

Summary



**Fleming
Ornskov, MD**

Q & A

All

- **Good performance**
- **Strategy progression**
- **Deeper insight into pipeline**
 - Dry eye disease
 - Binge eating disorder

Flemming Ornskov, MD
Chief Executive Officer



Our purpose

We enable people with life-altering conditions to lead better lives.

Accelerating product sales growth through 2013



We anticipate delivering full year double digit
Non GAAP earnings growth



To be as brave as the people we help.

Two strategic priorities

In-line

Drive **optimum performance** from our currently marketed products

Pipeline

Build our **future assets** through both R&D and Business Development

ADHD: initiatives to fuel VYVANSE® growth

- Renewed emphasis on pediatric market in time for Back-to-School period
- Added 50 incremental reps to specifically detail prescribers
 - Comprising 30 new reps hired and 20 reassigned from educational roles
- Increased coverage of the faster growing adult market
- Refocused all field activities on prescribers

Back-to-School (BTS) Opportunity

- There is a significant market opportunity during BTS (*Aug-Oct*)
 - 30% of pediatric market is available during BTS months
- Primary focus on pediatric and adolescent market share growth supported by:
 - Enhanced incentive compensation plan
 - New marketing messages and materials
 - New patient trial and access resources
 - Increased investment in online promotion for Health Care Providers and Consumers



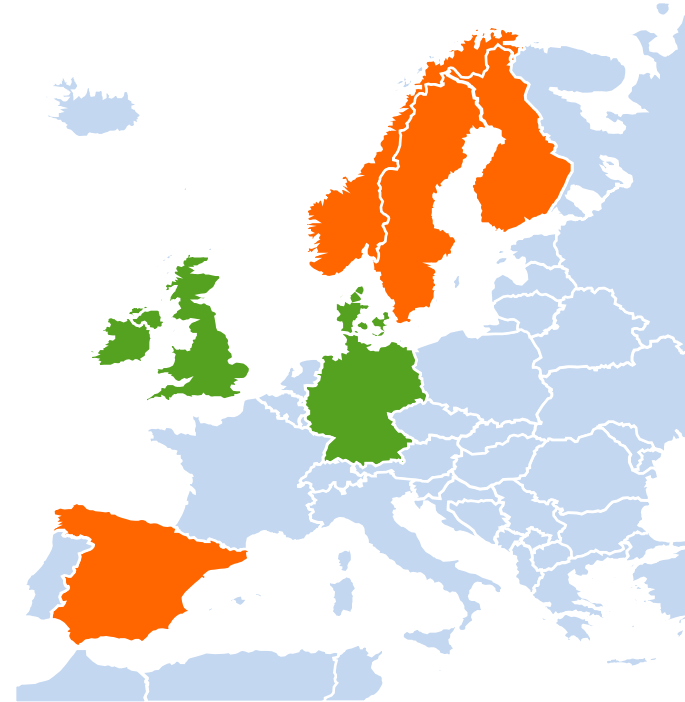
Please see full prescribing information



To be as brave as the people we help.

ELVANSE®: international expansion on track

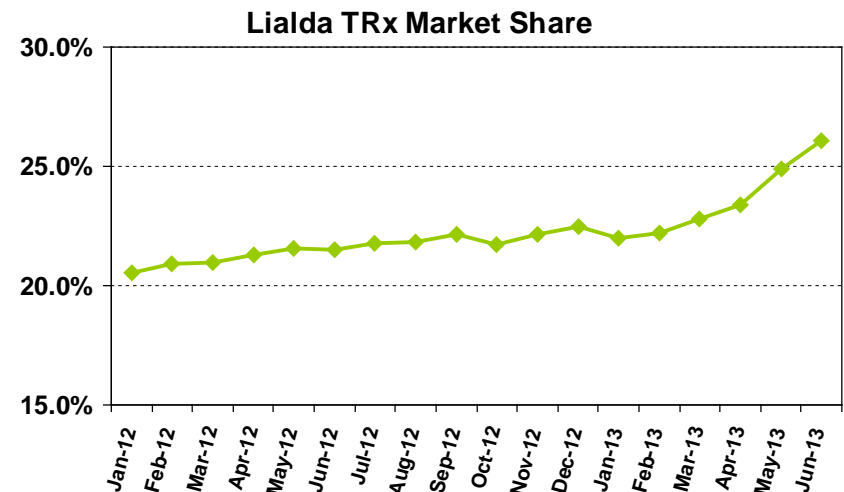
- Launches underway in UK, Denmark, Germany, Ireland
- 2014 launches planned for Spain, Sweden, Norway and Finland
- Eight target countries represent ~ 75% of the EU ADHD market



■ 2013 launch ■ 2014 launch

LIALDA®/MEZAVANT®: Capitalizing on growth opportunities

- 46% Growth in Q2 2013 Sales vs. prior year and 37% versus Q1 2013
 - US Lialda TRx growth is the main driver, strong growth ex-US in Australia, Canada & Spain
 - Favorable stocking trends & growth in US non-retail vs. Q1
- 5-ASA market leadership in the US and worldwide
 - ASACOL manufacturing cessation in the US provided opportunity
 - Lialda is now the market leading 5-ASA by Rx volume in the US
 - Shire 5-ASA franchise* is the global market leader in sales and Rx
- Favorable MCO formulary positioning
 - 9 out of 10 UC patients can now access Lialda in the US without managed care access restrictions
- Further, moderate share growth expected
 - Usage continues to grow among prescribers



*Lialda/Mezavant & Pentasa (US only)



To be as brave as the people we help.

FIRAZYR®: strong uptake in US continues

- Firazyr now has a high and growing share of all treated US acute attacks, according to our market research
- Physician and patient interest and feedback continue to be highly positive



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REPLAGAL[®]: strong leader in our markets

- Retaining a majority of patients that switched to Replagal and strong growth in naïve patients
- More commercial resources added
- We expect Replagal revenues to increase in second half*

Competing effectively: growing patient numbers

* Compared to the first half of 2013.



To be as brave as the people we help.

Two strategic priorities

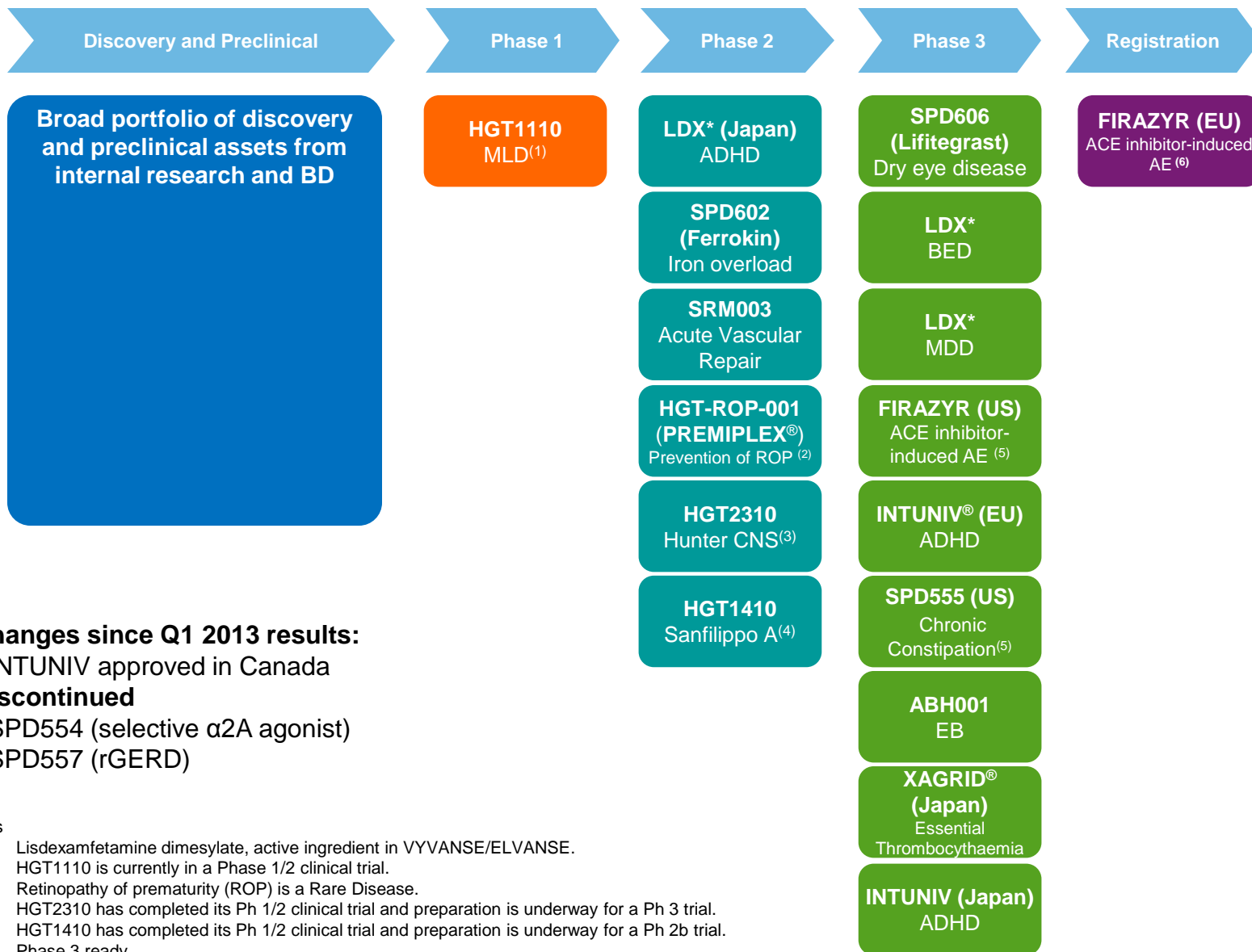
In-line

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Pipeline

Build our **future assets** through both R&D and Business Development

Building an innovative pipeline to deliver future growth



Changes since Q1 2013 results:

- INTUNIV approved in Canada
- Discontinued**
- SPD554 (selective α 2A agonist)
- SPD557 (rGERD)

Notes

- * Lisdexamfetamine dimesylate, active ingredient in VYVANSE/ELVANSE.
- (1) HGT1110 is currently in a Phase 1/2 clinical trial.
- (2) Retinopathy of prematurity (ROP) is a Rare Disease.
- (3) HGT2310 has completed its Ph 1/2 clinical trial and preparation is underway for a Ph 3 trial.
- (4) HGT1410 has completed its Ph 1/2 clinical trial and preparation is underway for a Ph 2b trial.
- (5) Phase 3 ready.
- (6) Application for EU label change, based on an investigator sponsored trial was, filed in December 2012.

Lifitegrast – Dry eye is a significant commercial opportunity



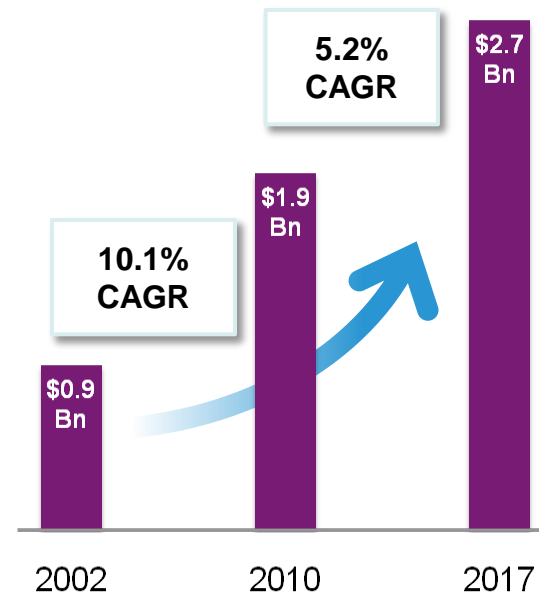
- One of the most common complaints to eye care specialists
- Inflammatory ocular surface disease giving rise to discomfort, dryness, gritty sensation, blurred vision

25M people affected
in the US

9M moderate to
severe sufferers

<10% patients currently
on prescription therapy

Potential global market of \$2.7 Billion



Source: 2011 Marketscope, LLC Comprehensive Report on the Global Market for Dry eye Products

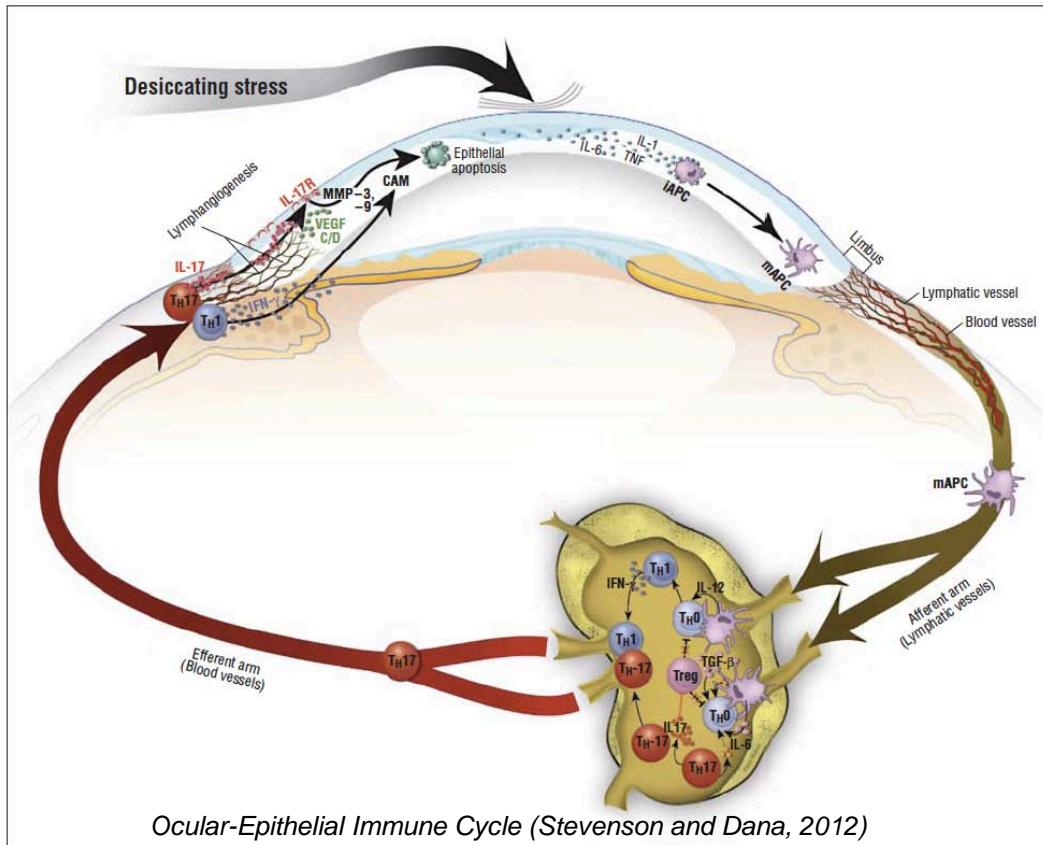
Source: GBI Research Oct 2011



To be as brave as the people we help.

Potential next generation therapy

Lifitegrast reduces chronic inflammation by inhibiting LFA-1/ICAM-1 binding that influences T-cell activation, mobility, and cytokine release

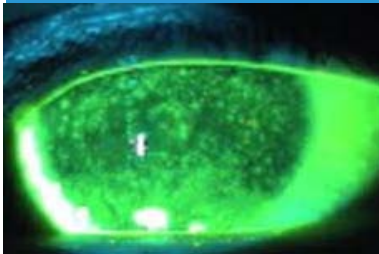


Hypothesis

Break cycle of chronic T-cell mediated inflammation

How is Dry eye efficacy assessed?

Signs



Objective measures of damage to the ocular surface

- Fluorescein – corneal surface
- Lissamine – conjunctival surface
- Staining = cell death / injury

Symptoms



Patient reported symptom score (0 – 100) based on questionnaire

- Discomfort, dryness
- Gritty sensation
- Sensitivity to light/ blurred vision

Lifitegrast: development program on track

Patient population	18 Years and older with history of dry eye in both eyes	18 Years and older with history of dry eye in both eyes	18 Years and older with history of dry eye in both eyes
Phase/study	Ph 3 OPUS-1	Ph 3 OPUS-2	Ph 3 SONATA
# of patients	N=588	N~700	N~300
Design	Multicenter, Randomized, Double-Masked and Placebo-Controlled Study	Multicenter, Randomized, Double-Masked and Placebo-Controlled Study <ul style="list-style-type: none"> • Patients must have used Artificial Tears within 30 days 	Multicenter, Randomized, Double-Masked and Placebo-Controlled Study <ul style="list-style-type: none"> • Long-term safety • 1-year duration
Primary endpoint	<ul style="list-style-type: none"> • Inferior Corneal Fluorescein Staining • Visual-related function subscale of OSDI • Safety and tolerability of lifitegrast Ophthalmic Solution (5.0%) compared to placebo, including incidence and severity of ocular and non-ocular adverse events. 	<ul style="list-style-type: none"> • Corneal staining score • Patient-reported dryness score • Safety and tolerability of lifitegrast Ophthalmic Solution (5.0%) compared to placebo, including incidence and severity of ocular and non-ocular adverse events 	<ul style="list-style-type: none"> • The Safety of Lifitegrast as assessed by ocular and Non-Ocular AEs
Status	<ul style="list-style-type: none"> • Results reported Oct 2012 	<ul style="list-style-type: none"> • Fully enrolled • Pivotal headline data expected 1Q 2014 	<ul style="list-style-type: none"> • Fully enrolled

Binge eating disorder: an emerging treatment area with high potential

Emerging science

- Significant increase in the science surrounding BED
- Increased disease awareness leading to a larger number of articles in print

New DSM classification

- Newly recognized as a disorder in the main section in DSM-5* (May 2013)

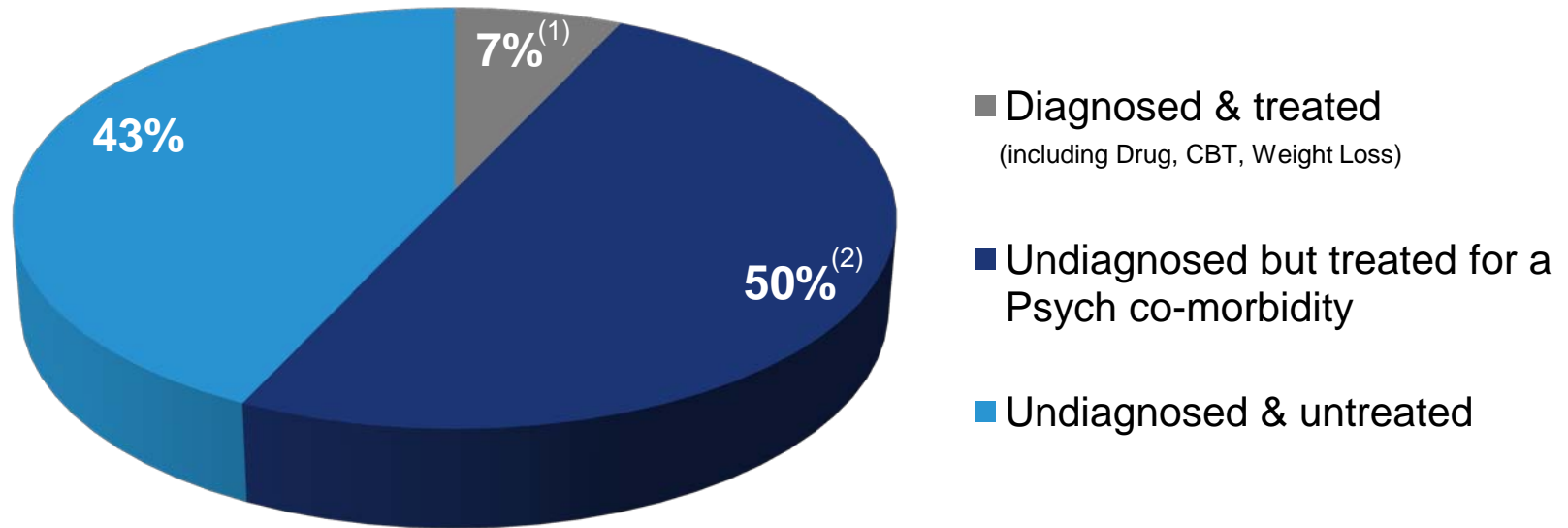
Market ready for education on diagnosis and treatment



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BED is a prevalent condition but is often not diagnosed and treated in the US

3 Million US adults suffer from BED (prevalence 1.2%)



Typical BED patient: Not easily noticeable - obesity rates similar to national average

Who are they? ⁽¹⁾

- Average age of onset for BED 25.4yrs, Average Duration: 8.1 yrs
- 67% are 18-44 years old
- 70% female
- 79% have life time psych co-morbidities

What do they do?

- Consume large amount of food - short duration
- Feel great guilt and shame

What do they think?

- Bingeing is something I do vs. something I have
- It is a character flaw - cannot control my binge episodes on my own

What do they want?

- To stop the binge and more importantly the urge to binge
- Don't want to live with this shame any longer

What is their environment?

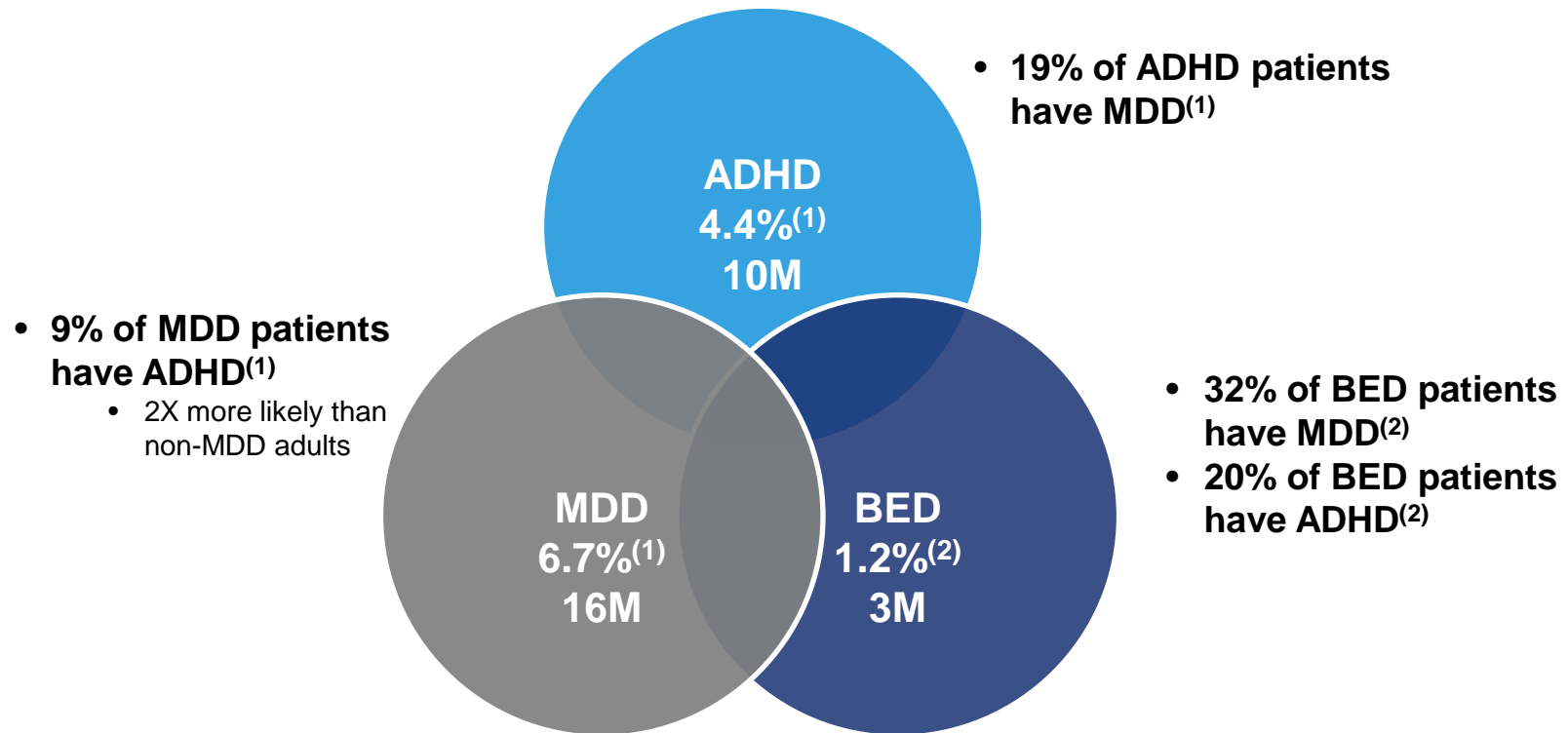
- Within the neighborhood - friend, or colleague at work
- Bingeing occurs in private - disorder is hidden from community



To be as brave as the people we help.

Patients that suffer from ADHD, MDD, or BED are at higher risk for a second condition

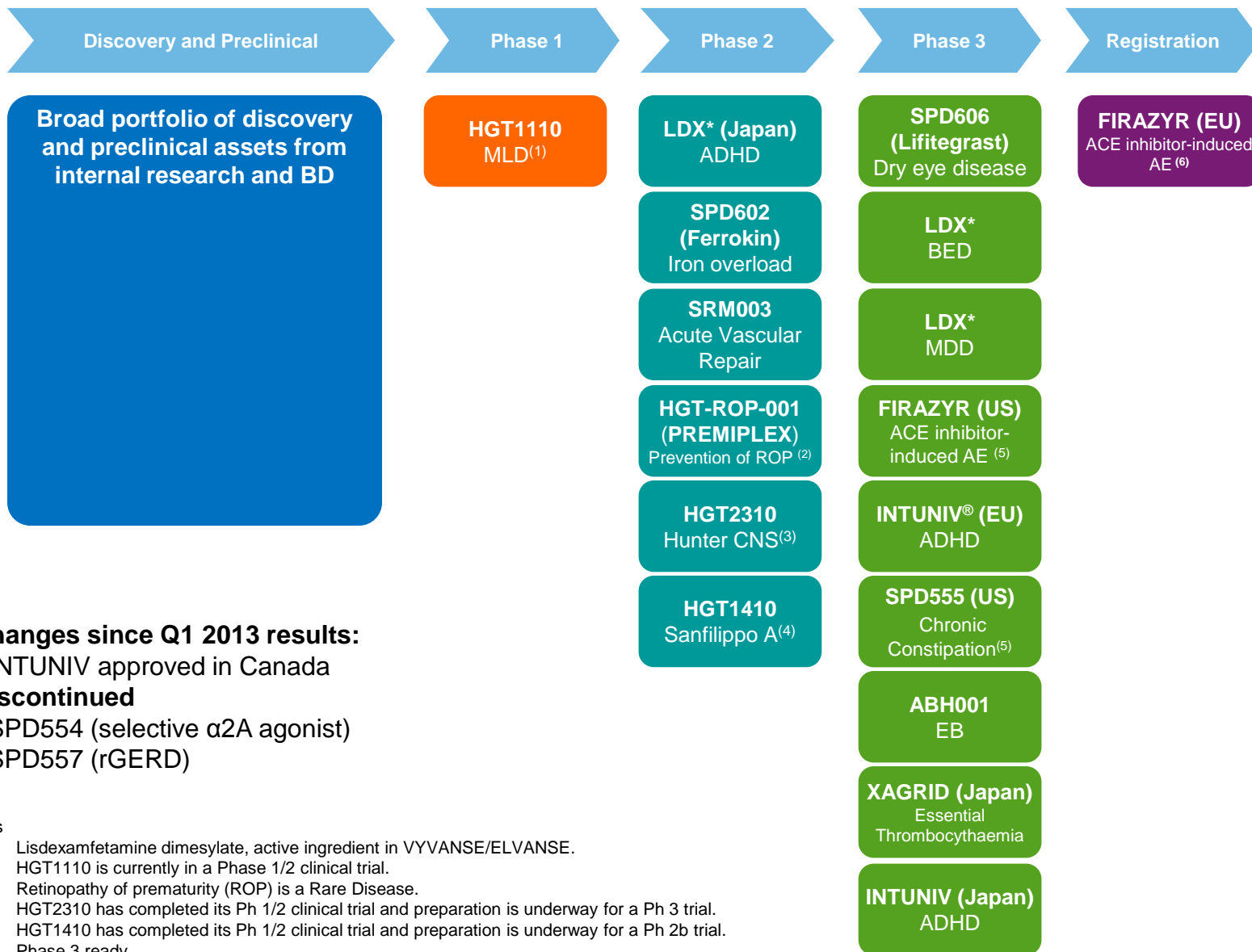
12-Month Prevalence of ADHD, MDD, BED in Adults



LDX BED program update: ahead of schedule

Patient population	Adults Aged 18-55 Years With Moderate to Severe Binge eating disorder	Adults Aged 18-55 Years With Moderate to Severe Binge eating disorder	Adults Aged 18-55 Years With Moderate to Severe Binge eating disorder
Phase/study	Ph 3 '343' study	Ph 3 '344' study	Ph 3 '345' study Open label extension from patients in studies 208, 343, 344
# of patients	N~356	N~356	N~530
Design	<ul style="list-style-type: none"> • Multicenter, Randomized, Double-blind, Parallel-group, Placebo-controlled, Dose-optimization Study • 50-70mg LDX once daily • 12 weeks 	<ul style="list-style-type: none"> • Multicenter, Randomized, Double-blind, Parallel-group, Placebo-controlled, Dose-optimization Study • 50-70mg LDX once daily • 12 weeks 	<ul style="list-style-type: none"> • Multicenter, Open-label, 12 Month Extension Safety and Tolerability Study • Start date Aug -12 • 50-70mg LDX once daily • 52 weeks
Primary endpoint	<ul style="list-style-type: none"> • Binge days per week 	<ul style="list-style-type: none"> • Binge days per week 	<ul style="list-style-type: none"> • Occurrence of treatment-emergent adverse events (TEAEs) as a measure of safety • Columbia Suicide Severity Rating Scale (C-SSRS)
Status	Pivotal study results expected 1Q 2014	Pivotal study results expected 1Q 2014	Pivotal study results expected 1Q 2014

Building an innovative pipeline to deliver future growth



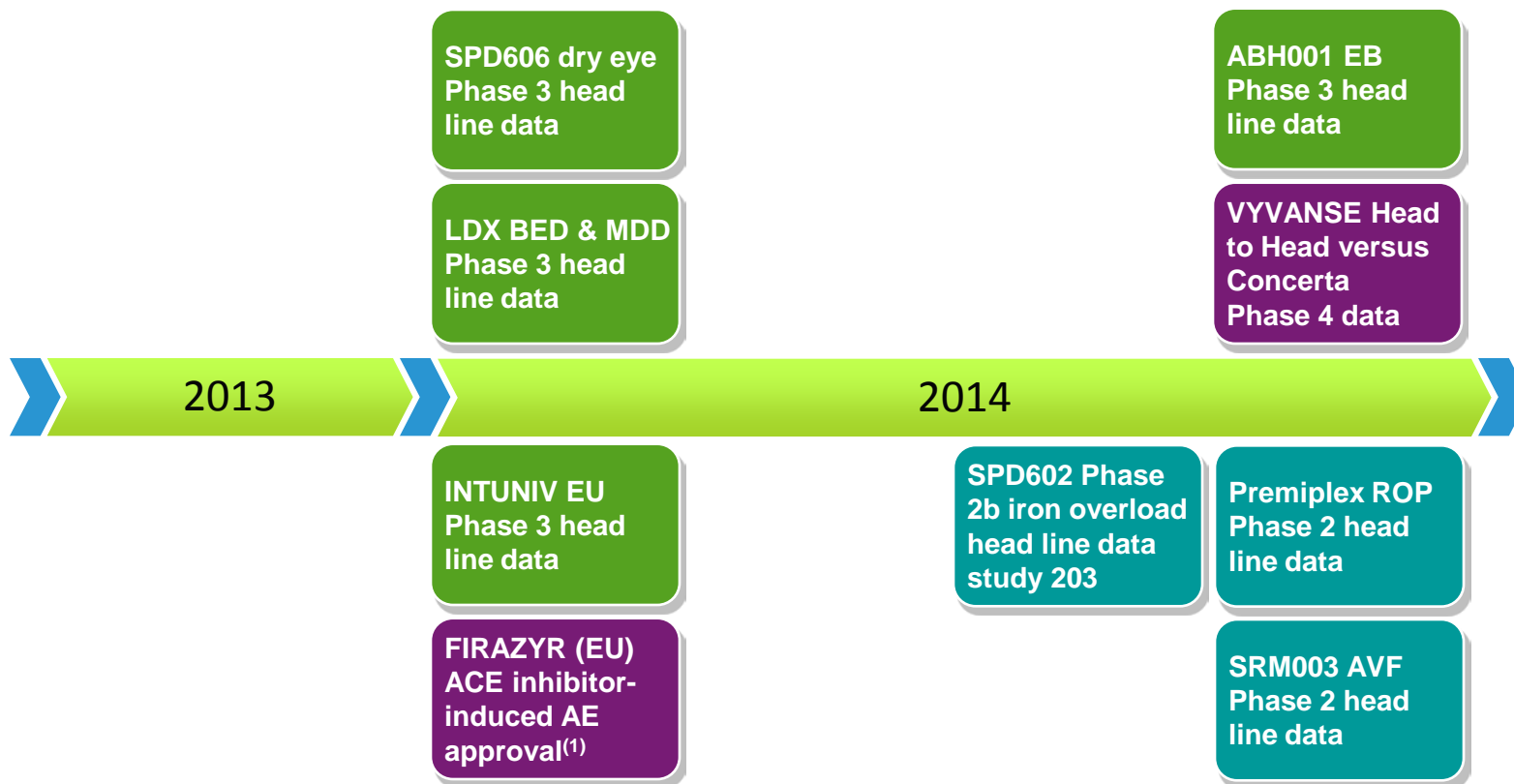
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Significant clinical milestones



- Registration and Phase 4
- Phase 3
- Phase 2

(1) Application for EU label change, based on an investigator sponsored trial was, filed in December 2012.

Financial review and 2013 outlook

Graham Hetherington
Chief Financial Officer



Our purpose

We enable people with life-altering conditions to lead better lives.

Improved product sales growth drives Q2 performance

	Q2 2013 \$m	Q2 2012 \$m	Reported Growth	Like for Like Growth ⁽¹⁾
Product sales	1,230	1,148	+7%	+8%
Product sales excluding ADDERALL XR	1,118	1,014	+10%	+11%
Royalties and other revenues	45	60	-26%	-26%
Total revenues	1,275	1,208	+6%	+6%
EBITDA⁽¹⁾	483	448	+8%	+7%
EBITDA % of product sales⁽¹⁾⁽²⁾	36%	34%	187bp	
EPS - ADS⁽¹⁾	\$1.79	\$1.68	+6%	
Cash generation⁽¹⁾	374	520	-28%	

(1) These are Non GAAP financial measures. See appendix for a list of items excluded from the US GAAP equivalent used to calculate these measures.

(2) Excluding royalties and other revenues.



To be as brave as the people we help.

Majority of top ten products deliver double digit growth

	Q2 2013 \$m	Q2 2012 \$m	Reported Growth		CER Growth
			\$m	%	%
VYVANSE	300	266	34	+13%	+13%
ELAPRASE	149	122	27	+22%	+25%
LIALDA / MEZAVANT	138	94	44	+46%	+46%
REPLAGAL	114	123	(9)	-7%	-5%
INTUNIV	90	69	21	+31%	+31%
VPRIV	83	83	-	-	+1%
PENTASA	74	64	10	+15%	+15%
FIRAZYR	50	32	18	+56%	+56%
DERMAGRAFT	22	52	(30)	-57%	-57%
OTHER	98	109	(11)	-9%	-8%
Product sales excluding ADDERALL XR	1,118	1,014	104	+10%	+11%
ADDERALL XR	112	134	(22)	-16%	-16%
Product sales	1,230	1,148	82	+7%	+8%

Continuing delivery of operating leverage

Year on Year:	2013 YTD	2012 YTD
Product sales	+4%	+20%
R&D⁽¹⁾	+15%	+15%
SG&A⁽¹⁾	-10%	+15%
Combined R&D and SG&A⁽¹⁾	-2%	+15%

Ratios:

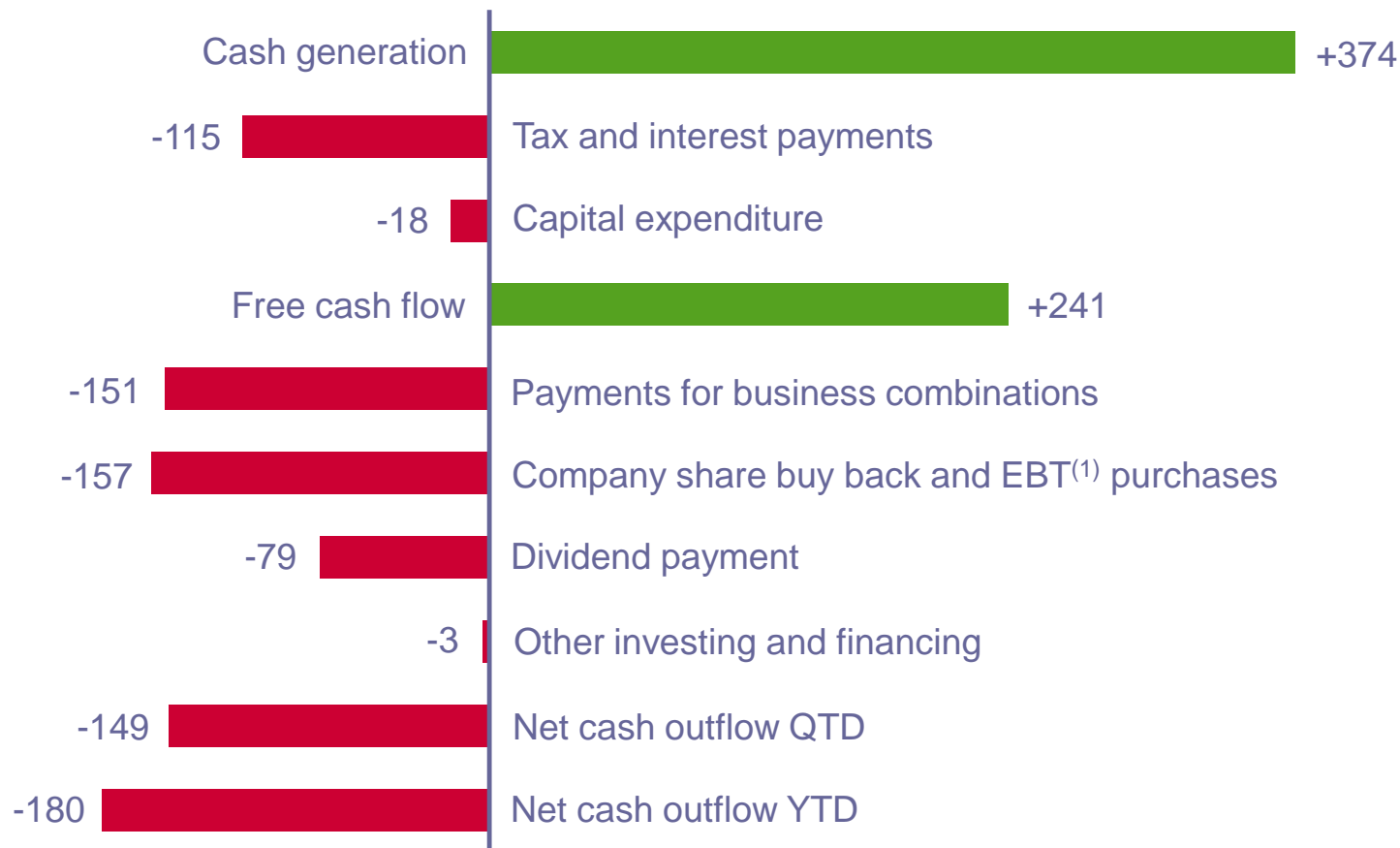
% of product sales		
Gross margin⁽¹⁾	87%	87%
R&D⁽¹⁾	19%	18%
SG&A⁽¹⁾	32%	38%
EBITDA^{(1) (2)}	35%	32%

(1) These are Non GAAP financial measures. See appendix for a list of items excluded from the US GAAP equivalents used to calculate these measures.

(2) Excluding royalties and other revenues.

Good cash generation in Q2

Millions of USD



Note: Shire has a revolving 5 year credit facility of \$1.2bn signed in November 2010 which remained undrawn as at June 30, 2013.

(1) Employee Benefit Trust ("EBT").



To be as brave as the people we help.

Anticipate delivering double digit Non GAAP earnings growth in 2013

Full year 2013 dynamics

	Direction v. FY 2012	Latest guidance	Previous guidance
Product sales	↑	Growth in the mid-to-high single digits	
Royalties and Other revenues	↓	Combined royalties & other revenues down 35-40%	Combined royalties & other revenues down 30-40%
Gross margins	≈	At a similar level to 2012	
R&D	↑	Low double digit growth	Low-to-mid teens growth
SG&A	↓	2-4% lower than 2012	Marginally lower than 2012
Combined R&D and SG&A	↑	Only marginally higher than 2012	Low single digit growth
Tax rate	≈	Core effective tax rate of 18-20%	
Reported EPS-ADS	↑	Anticipate delivering full year double digit Non GAAP earnings growth	In line with current consensus earnings expectations ⁽¹⁾

(1) Based on the consensus update at the time of the Q1 earnings announcement (compiled by Consensus Forecast Ltd as of May 2, 2013) of \$6.67 Non GAAP diluted earnings per ADS for the year ending December 31, 2013.

Summary

Flemming Ornskov, MD
Chief Executive Officer



Our purpose

We enable people with life-altering conditions to lead better lives.

Good performance and progressing our strategy

Strong and sustainable business that meets patients' needs today

Optimising our structure and ways of working, generating a healthy profit and cash flow

Investing in our pipeline to meet the unmet needs of tomorrow

Anticipate delivering full year double digit Non GAAP earnings growth

Questions and Answers



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Appendix



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Operating leverage – Key financial ratios

Year on Year:	Q2 2013	Q2 2012
Product sales	+7%	+16%
R&D⁽¹⁾	+15%	+20%
SG&A⁽¹⁾	-5%	+5%
Combined R&D and SG&A⁽¹⁾	+2%	+10%

Ratios:

% of product sales		
Gross margin⁽¹⁾	87%	87%
R&D⁽¹⁾	19%	18%
SG&A⁽¹⁾	32%	36%
EBITDA^{(1) (2)}	36%	34%

(1) These are Non GAAP financial measures. See appendix for a list of items excluded from the US GAAP equivalents used to calculate these measures.

(2) Excluding royalties and other revenues.

Product sales – regional analysis

	US \$m	Europe \$m	LATAM \$m	Other \$m	Total \$m
Q2 2013 product sales	829	246	69	86	1,230
% of Product sales	67%	20%	6%	7%	
YoY growth	7%	-3%	91%	6%	7%
YTD 2013 product sales	1,589	487	101	170	2,347
% of Product sales	68%	21%	4%	7%	
YoY growth	5%	-2%	12%	10%	4%
FY 2012 product sales	2,929	983	171	324	4,407
% of Product sales	67%	22%	4%	7%	
YoY growth	14%	0%	33%	19%	12%

Royalties & Other revenues

	Q2 2013 \$m	Q2 2012 \$m	Reported Growth
3TC and ZEFFIX	11	10	+7%
FOSRENOL	11	13	-17%
ADDERALL XR	5	26	-81%
REMINYL & Other	9	7	+33%
Royalties	36	56	-36%
Other revenues	9	4	+111%
Royalties & Other revenues	45	60	-26%

Shire income statement growth analysis

	2012 Q1	2012 Q2	2012 Q3	2012 Q4	2012 FY	2013 Q1	2013 Q2	FY 2013 Dynamics	
								Direction v. FY 12	Explanations
Total Product Sales	\$1,107m	\$1,148m	\$1,055m	\$1,097m	\$4,407m	\$1,117m	\$1,230m	↑	Growth in the mid-to-high single digits
versus prior year	+24%	+16%	+4%	+5%	+12%	+1%	+7%		
Royalties & Other revenues	\$65m	\$60m	\$45m	\$104m	\$274m	\$45m	\$45m	↓	Combined royalties & other revenues down 35-40%
versus prior year	-22%	-14%	-32%	+11%	-12%	-31%	-26%		
Total Revenues	\$1,172m	\$1,208m	\$1,100m	\$1,201m	\$4,681m	\$1,162m	\$1,275m	≈	At a similar level to 2012
versus prior year	+21%	+14%	+1%	+5%	+10%	-1%	+6%		
Gross Margin ^{(1) (2)}	86%	87%	85%	86%	86%	87%	87%	≈	At a similar level to 2012
Combined R&D and SG&A ⁽²⁾	\$632m	\$615m	\$588m	\$645m	\$2,480m	\$592m	\$626m	↑	Only marginally higher than 2012
versus prior year	+20%	+10%	+3%	+8%	+10%	-6%	+2%		
Tax Rate ⁽²⁾	20%	20%	18%	15%	18%	19%	+23%	≈	Core effective tax rate of 18-20%
EPS – ADS ⁽²⁾	\$1.48	\$1.68	\$1.36	\$1.58	\$6.10	\$1.63	\$1.79	↑	Anticipate delivering full year double digit Non GAAP earnings growth
versus prior year	+20%	+26%	+6%	+4%	+14%	+10%	+6%		

(1) Gross margin calculated as a percentage of product sales.

(2) These are Non GAAP financial measures. See appendix for a list of items excluded from the US GAAP equivalents used to calculate these measures.



To be as brave as the people we help.

Non GAAP cash flow measures

Non GAAP cash generation and free cash flow reconciliation	Q2 2013 \$m	Q2 2012 \$m
Non GAAP cash generation⁽¹⁾	374	520
Tax and interest payments, net	(115)	(54)
US GAAP net cash provided by operating activities	259	466
Capital expenditure	(18)	(33)
Non GAAP free cash flow⁽²⁾	241	433

- (1) Non GAAP cash generation represents net cash provided by operating activities, excluding up-front and milestone payments for in-licensed and acquired products, tax and interest payments.
- (2) Non GAAP free cash flow represents net cash provided by operating activities, excluding up-front and milestone payments for in-licensed and acquired products, but including capital expenditure in the ordinary course of business.

Non GAAP net cash

	June 30, 2013 \$m	December 31, 2012 \$m
Cash and cash equivalents	1,302	1,482
Convertible bonds	(1,100)	(1,100)
Other	(9)	(9)
Net cash	193	373

Non GAAP measures

- This presentation contains financial measures not prepared in accordance with US GAAP.
- These Non GAAP financial measures are used by Shire's management to make operating decisions because they facilitate internal comparisons of the Company's performance to historical results and to competitors' results. They should not be considered in isolation from, as substitutes for, or superior to financial measures prepared in accordance with US GAAP.
- The following items are excluded from these non-GAAP financial measures:
 - Amortization and asset impairments:**
 - Intangible asset amortization and impairment charges; and
 - Other than temporary impairment of investments.
 - Acquisitions and integration activities:**
 - Up-front payments and milestones in respect of in-licensed and acquired products;
 - Costs associated with acquisitions, including transaction costs, and fair value adjustments on contingent consideration and acquired inventory;
 - Costs associated with the integration of companies; and
 - Noncontrolling interest in consolidated variable interest entities.
 - Divestments, re-organizations and discontinued operations:**
 - Gains and losses on the sale of non-core assets;
 - Costs associated with restructuring and re-organization activities;
 - Termination costs; and
 - Income / (losses) from discontinued operations.
 - Legal and litigation costs:**
 - Net legal costs related to the settlement of litigation, government investigations and other disputes (excluding internal legal team costs).