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R&D Strategy: Reliable fill & flow with greater novelty and improved return on investment



Accelerate Discovery output

- Now have 30 DPUs, of which two thirds are from the original 2009 set. Average 20% turnover every 3 year cycle
- 65% of NMEs* in the clinic were either discovered or worked on by the DPUs
- Average of 60-65 publications annually in world class journals across pharma and vaccines

Focus where science is innovative

- 80% of NMEs*, biologicals and vaccines have potential to be 1st in class
- Competitive advantage through epigenetics, cell & gene technology, adjuvants, self amplifying RNA, inhaled technology, chimp adenovector

Improve balance internal vs external

- 60% of NMEs* in the clinic are home-grown, 40% partnered or in-licensed
- >1,500 collaborations inclusive of academic, public-private partnerships, biotech and pharma

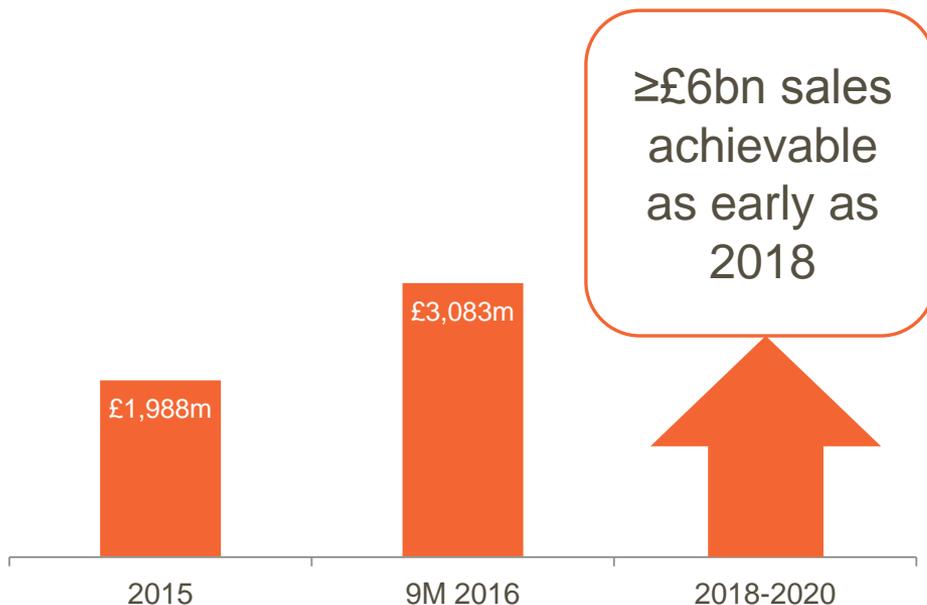
Reduce fixed cost and improve ROI

- 20% faster study execution times[^]
- Pharma R&D headcount reduced from 12,000 to 8,500 since 2008, reduced to two global pharma R&D hubs
- Balance discovery and development R&D spend (pharma split ~40% Discovery; ~60% Development)

R&D driving growth and returns to shareholders



Annual sales from 11 new products*



25% of pharmaceutical sales from new pharma products⁺ in Q3 2016

2016 pipeline progress:

Filed 4 assets for regulatory approval

Closed triple

Shingrix

Benlysta SC

sirukumab RA

Started 5 Phase III studies

Started 5 Phase II studies

*11 new products defined as: Breo, Anoro, Incruse, Arnuity, Nucala, Tanzeum, Tivicay, Triumeq, Menveo, Bexsero and Shingrix. All expectations and targets regarding future performance should be read together with the "Assumptions related to 2016-2020 outlook" on page 35 of the Group's third quarter earnings release dated 26 October 2016.

⁺New products refers to pharma only excluding vaccines

R&D Strategy: focused on 6 therapy areas



**HIV / Infectious
Diseases**

Respiratory

Vaccines

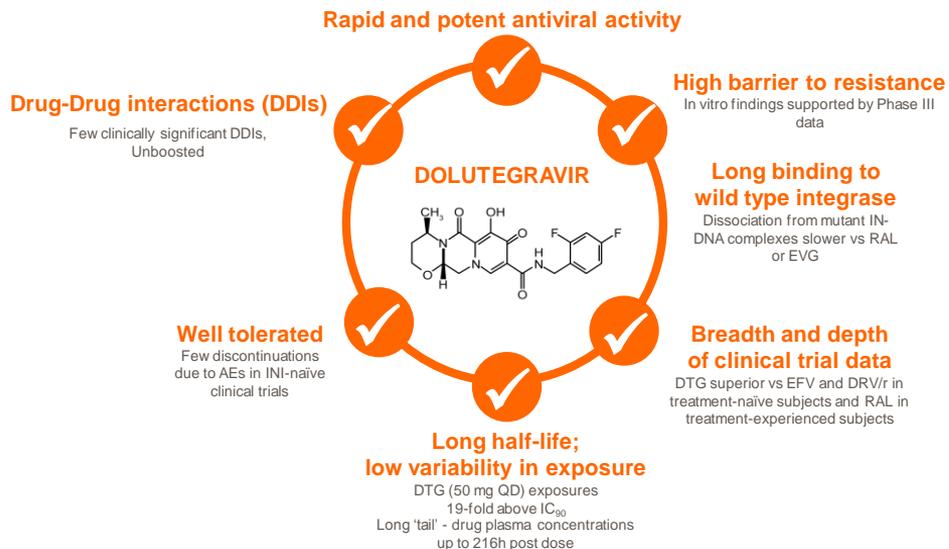
**Immuno-
Inflammation**

Oncology

**Rare
Diseases**

Amongst integrase inhibitors, dolutegravir stands out

Unique product characteristics



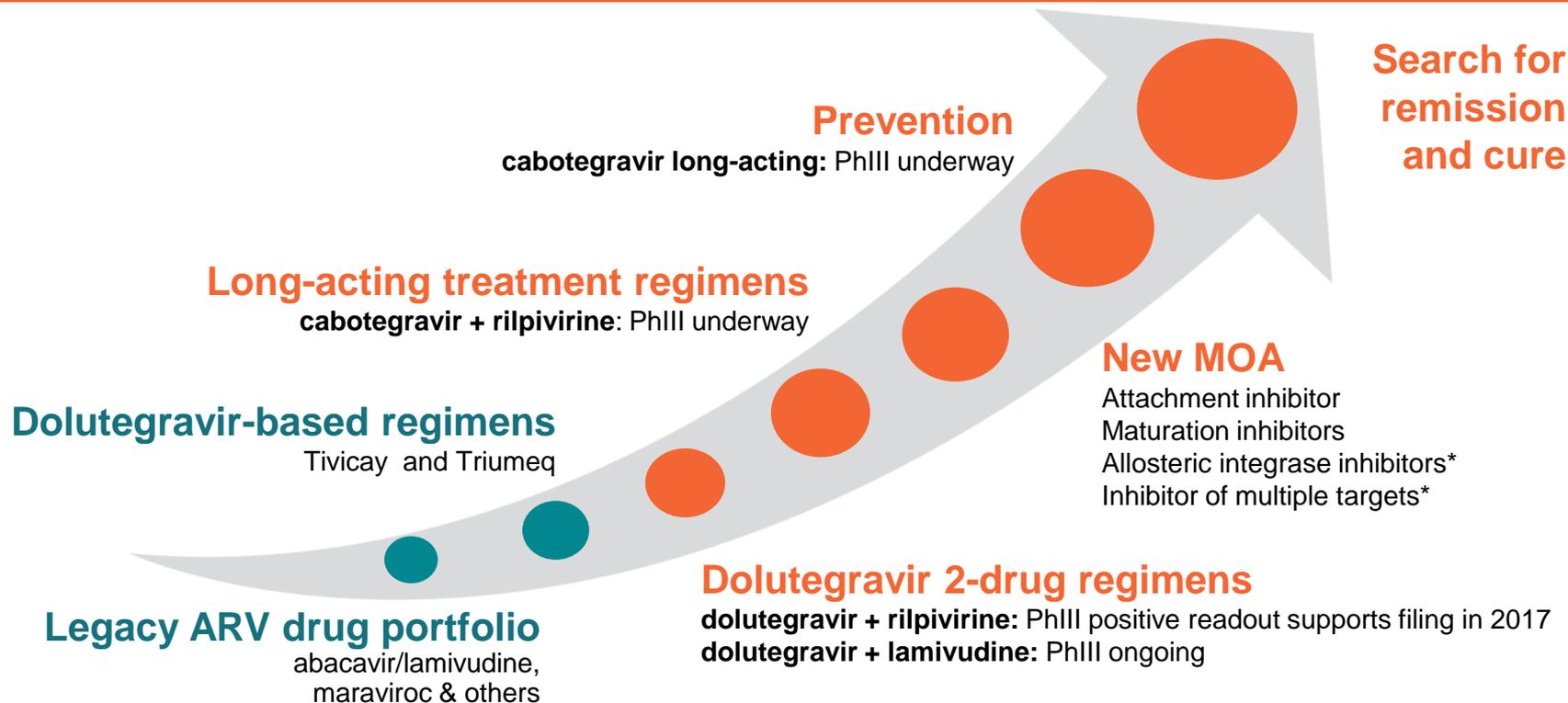
Unprecedented and unmatched clinical trial results in HIV

Vs. efavirenz	Vs. raltegravir	Vs. darunavir	Vs. atazanavir
SUPERIOR (naïve)	SUPERIOR (experienced)	SUPERIOR (naïve)	SUPERIOR (women/naïve)
	 NON INFERIOR (naïve)		

SINGLE, FLAMINGO, SPRING 2, SAILING and ARIA were non-inferiority studies with a pre-specified analysis for superiority. Chart shows primary endpoint outcomes

Positive headline results from dolutegravir + rilpivirine two drug regimen Phase III study, supports filing in 2017

Innovative pipeline addressing unmet patient needs



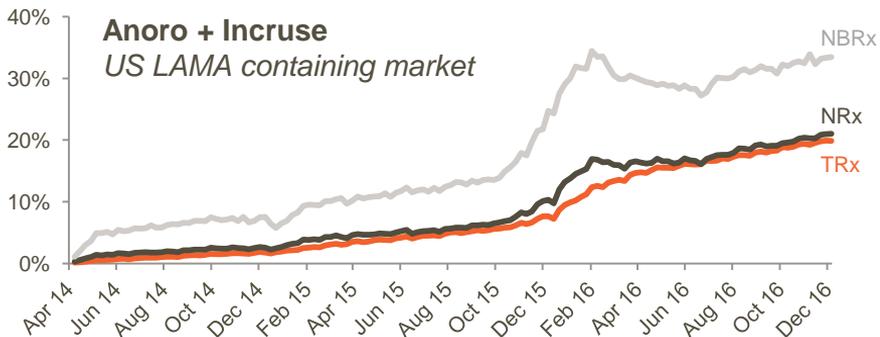
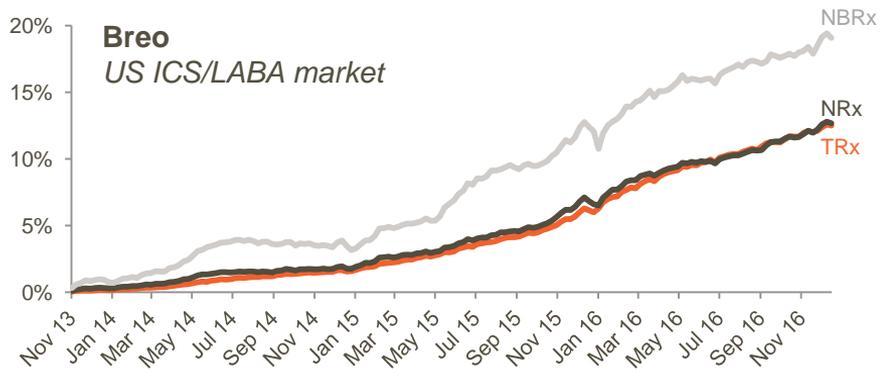
*Denotes preclinical asset

Ongoing studies: DTG+3TC GEMINI studies started Aug 2016; CAB+RPV ATLAS and FLAIR studies started Nov 2016; CAB monotherapy HPTN083 study started Dec 2016

Portfolio of once-a-day, easy-to-use Ellipta inhalers



Strong commercial performance; closed triple filed



Closed triple:

- Filed in US and EU for COPD in Q4 2016
 - 10 month review expected in US
- FULFIL data demonstrated superiority vs Symbicort in lung function presented at ERS Sept 2016
- IMPACT COPD exacerbation data expected H2 2017
- Started Phase III for asthma Q4 2016

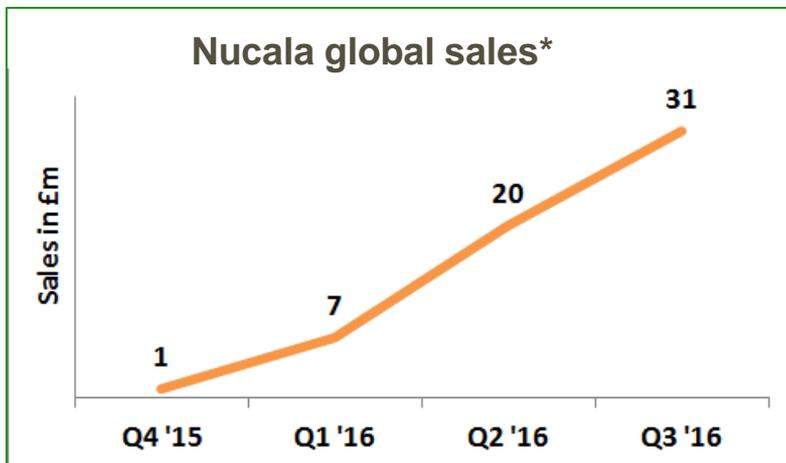
Completes Ellipta inhaler portfolio



Nucala launch off to a strong start



Additional data and indications expected to drive further growth



Launched in US, Europe, Japan

US J code available Jan 2017



Pipeline update:

- COSMOS study† on positive long term safety and efficacy of Nucala presented at AAAAI
- JACI publication^ showing hospitalisations and ER visits halved with Nucala
- MUSCA study showing QoL and lung function to be presented at AAAAI, March 2017
- Phase III COPD data expected 2017
- In development for:
 - Eosinophilic granulomatosis with polyangiitis (EGPA)
 - Atopic dermatitis
 - Hyper eosinophilic syndrome (HES)
 - Nasal polyposis

*Source: GSK company results

†Long-term Efficacy and Safety of Mepolizumab in Patients With Severe Eosinophilic Asthma: A Multi-center, Open-label, Phase IIIb Study Njira Lugogo, MD; Christian Domingo, MD; Pascal Chanez, MD, PhD; Richard Leigh, MBChB; Martyn J. Gilson, MSc; Robert G. Price, MSc; Steven W. Yancey, MSc; and Hector G. Ortega, MD. Clinical Therapeutics/Volume 38, Number 9, 2016

^Meta-analysis of asthma-related hospitalization in mepolizumab studies of severe eosinophilic asthma. Yancey S, Ortega H, Keene O, Mayer B, Gunsoy N, Brightling C, Bleecker ER, Haldar P, Pavord I. Journal of Allergy and Clinical Immunology, 2016

Mepolizumab data in eosinophilic lung disease (EGPA)



Phase III data supports 2017 filing

Co-primary endpoints

	mepolizumab	placebo	p value
Accrued duration of remission	19/68 (28%)	2/68 (3%)	p<0.001
Remission at wk 36 and 48	22/68 (32%)	2/68 (3%)	p<0.001

Secondary endpoints

	mepolizumab	placebo	p value
Average OCS dose during last 4 wks \leq 4mg/day	30/68 (44%)	5/68 (7%)	p<0.001
Remission within first 24 wks and maintained to study end	13/68 (19%)	1/68 (1%)	p=0.007
Time to first EGPA relapse	Hazard ratio = 0.32; 95% CI: (0.21, 0.50)		p<0.001

Full results from the study, including data from the secondary endpoints, will be submitted for presentation at an upcoming scientific congress and for publication in a peer-reviewed journal. The pivotal phase III study, MEA115921, was a randomised, double-blind study with the purpose to investigate the efficacy and safety of mepolizumab 300mg (administered subcutaneously every 4 weeks) compared with placebo over a 52-week study treatment period in 136 patients with relapsing or refractory EGPA receiving standard of care therapy including background corticosteroid therapy with or without immunosuppressive therapy.

Next generation respiratory medicines



Subdivision of severe asthma patients



Phenotypically distinct patients: anti-IL33r – PhIIa start 2017
Extended pharmacology: anti-IL5 mAb – PhIII start 2018

Disease modification in COPD



Inhaled PI3K δ inhibitor – PhIIb start 2017
mepolizumab – file 2017
danirixin – PhIIb start 2017

Potential in additional disease areas



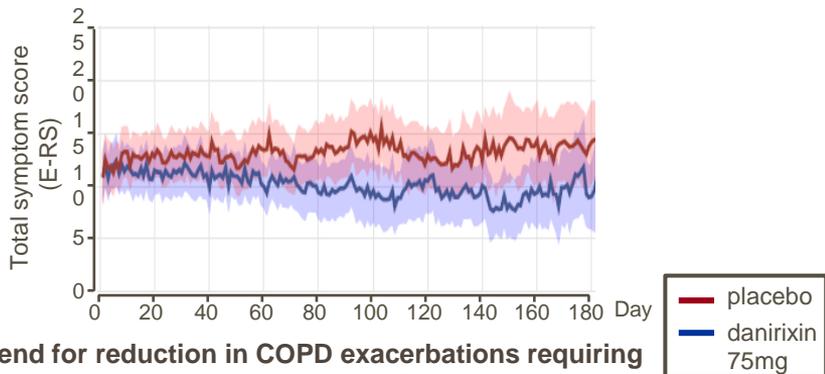
IPF: Inhaled $\alpha\beta 6$ inhibitor – PhIIa start 2018
ALI: TNFR1 antagonist dAb – PhII data 2017

Pipeline progression in two promising new mechanisms of action

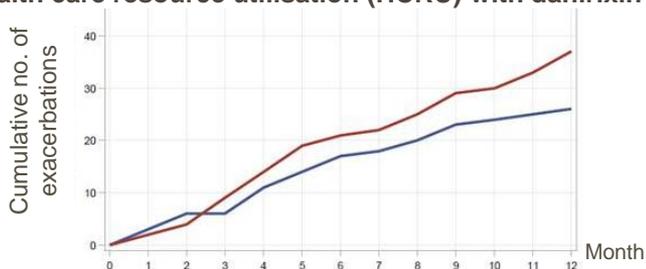


Oral danirixin

Real-time data demonstrate improvement of symptoms with danirixin in symptomatic COPD (frequent exacerbators)



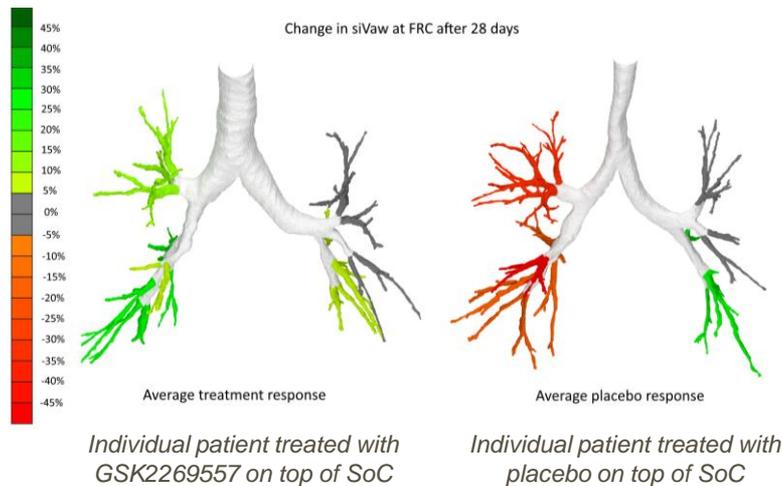
Trend for reduction in COPD exacerbations requiring health care resource utilisation (HCRU) with danirixin*



GSK, data on file (study 200163). *Interim PhII data in symptomatic COPD

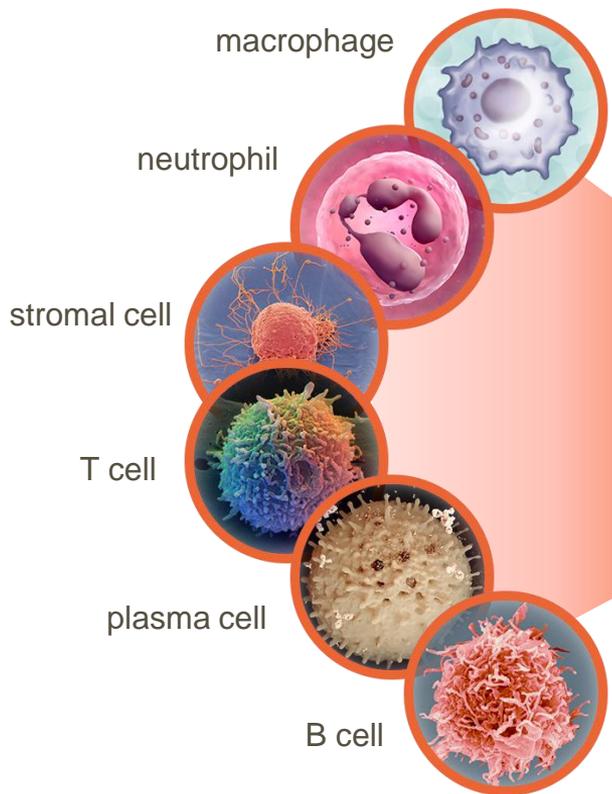
Inhaled PI3K δ inhibitor

Directionality of neutrophil migration is aberrant in COPD patients and corrected by PI3K inhibition - *in vitro*



Sapey et al, *AJRCCM* 2011; 183: 1176
Burrowes et al. *Interface Focus* 2013;3:20120057 (Fluidita)

Deep pipeline in Immuno-Inflammation



GSK Pipeline

Targeted Biologicals

- Benlysta
- sirukumab
- Anti-GM-CSF
- Anti-IL-7
- Anti-CCL20
- Anti-LAG3
- Anti-OSM

Targeted Small Molecules

- RIP1
- I-BET

Targeting Resistant Disease

- RIP1
- I-BET
- Anti-IL-7
- Anti-CCL20
- Anti-LAG3
- Anti-OSM

Early Intervention & Remission Induction

- Anti-GM-CSF
- RIP1
- Anti-CCL20
- Anti-LAG3

Benlysta: extensive ongoing development



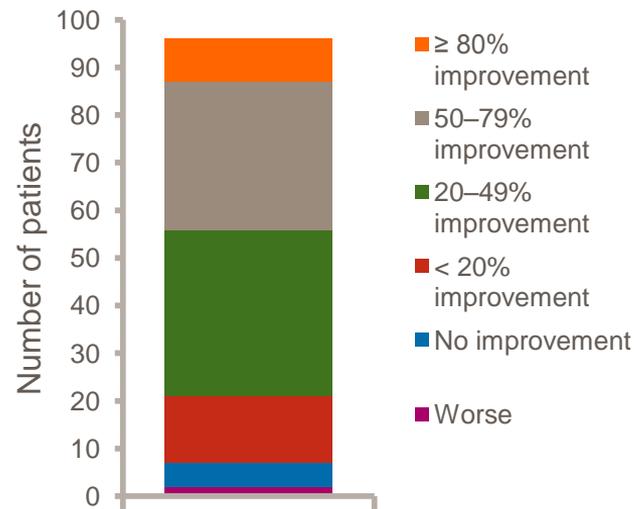
The only medicine approved to treat SLE in over 50 years

- Only medicine to treat SLE* to have succeeded in PhIII
 - Three other medicines have recently failed
- 4th consecutive positive pivotal study
 - Improvement in time to first severe flare
 - Trend for reduction in corticosteroid use
 - Further filings Japan (Dec 2016); China (2017)
- Multiple ongoing studies, including subgroups in SLE, lupus nephritis, long-term remission pre-treatment with rituximab and other indications

Sub-cutaneous formulation
filed in US & EU Sept 2016



Real world studies reinforce effectiveness
through strong patient response



Real world studies observed an overall clinical improvement of at least 20% in 78% of patients

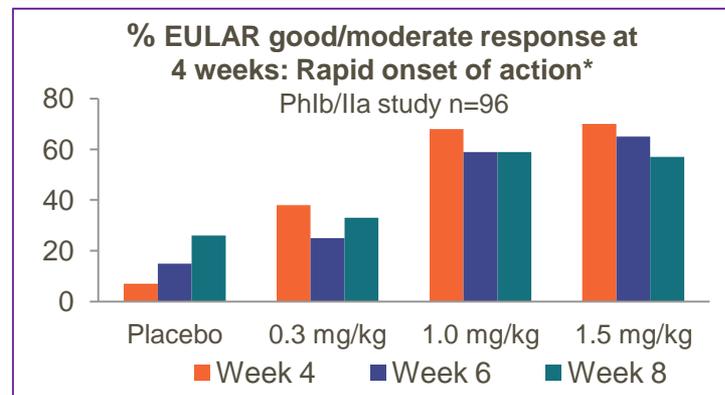
Schwartz A, et al. *Rheumatol Ther* 2016

GSK'165: potential first in class anti-GM-CSF



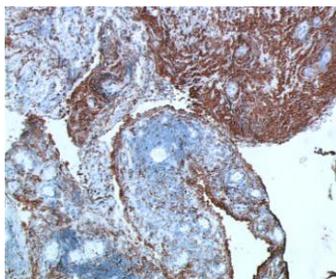
Granulocyte-macrophage colony stimulating factor (GM-CSF) for RA and hand OA

- Activated macrophages abundantly expressed in early RA synovial tissue
- Reduction in macrophage infiltration correlates with improvement in disease activity scores^{1,2}
- Important in macrophage production and infiltration in the tissues
- Macrophage related markers may facilitate a precision medicine approach
- Potential to target a number of immuno-inflammatory diseases



'165 potential as first in class aGM-CSF

- Phase IIb study ongoing in RA
- Global programme, including US
- High bar fertility hurdle achieved
- Data expected H2 2017



Further studies for hand osteoarthritis underway, Phase II data expected H2 2017

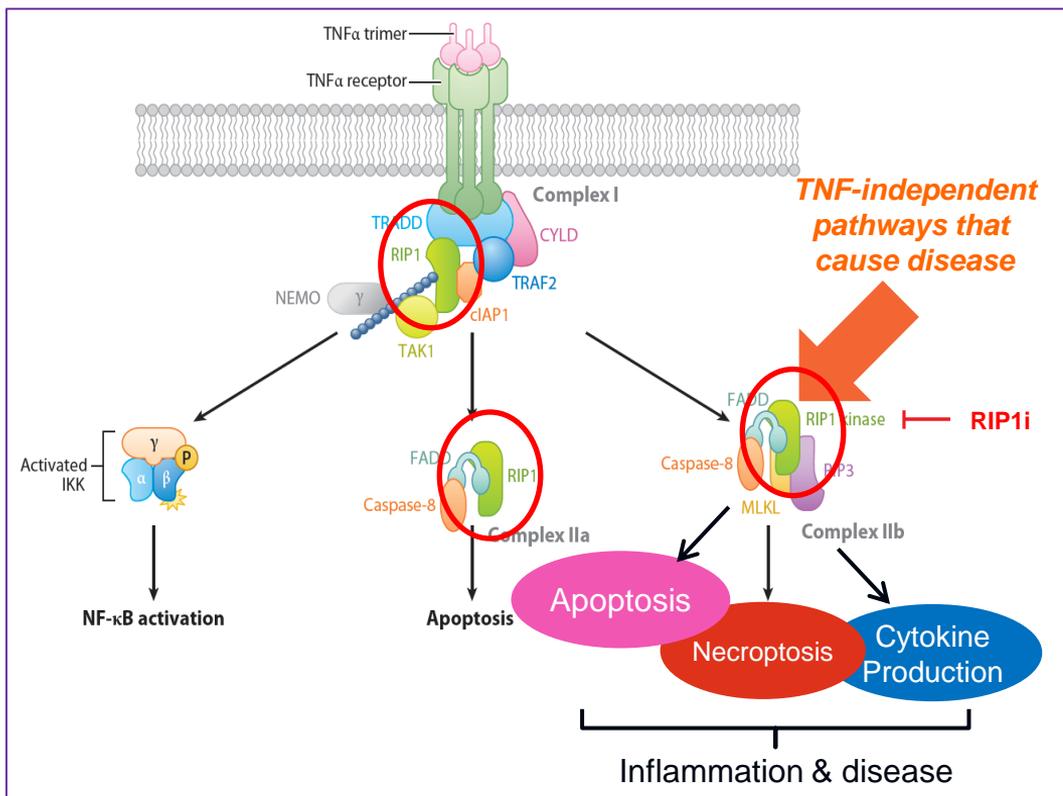


¹ Boumans MJ, *et al.* Arthritis Rheum. 2011;63:3187-94.

² Bresnahan B, *et al.* J Rheumatol 2009;36:1800-2.

*Behrens, *et al.* Ann Rheum Dis. 2015;74:1058-64

GSK'772: oral anti-inflammatory RIP-1 kinase inhibitor with potential for psoriasis, RA and ulcerative colitis

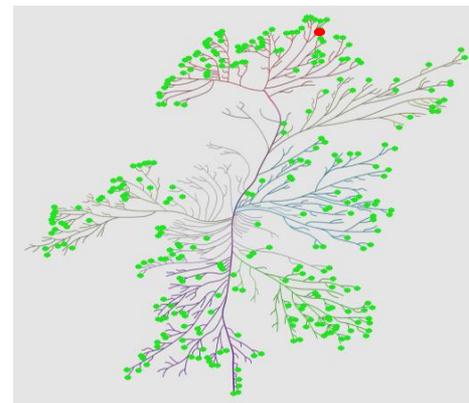


Derived from Christofferson et al., Ann.Rev. Physiol. (2014) 76:129

Phlla readouts 2017/18:

- Psoriasis
- RA

Kinome plot



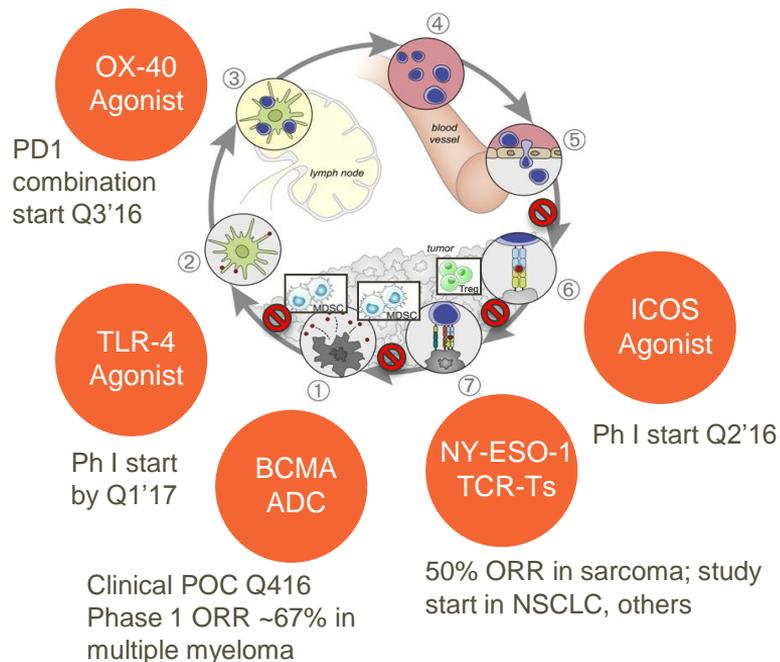
GSK2982772 - most selective ATP competitive kinase inhibitor to advance into man

ATP = adenosine triphosphate

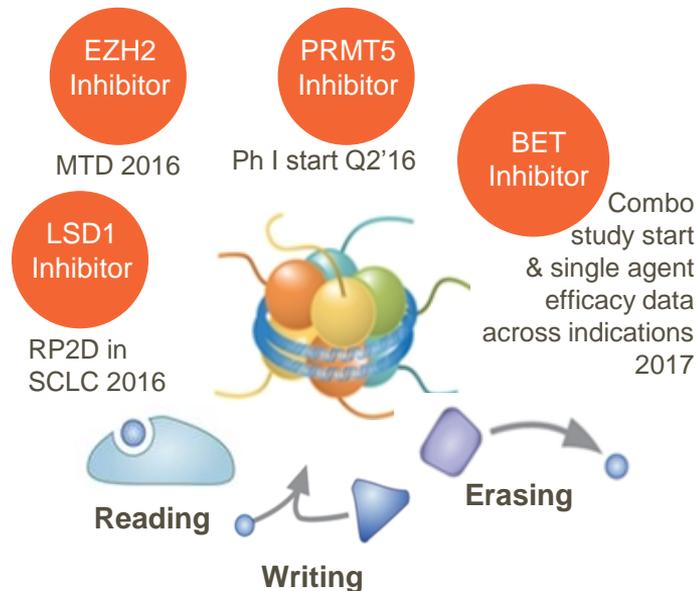
Multiple pipeline opportunities in oncology



Immuno-Oncology



Epigenetics

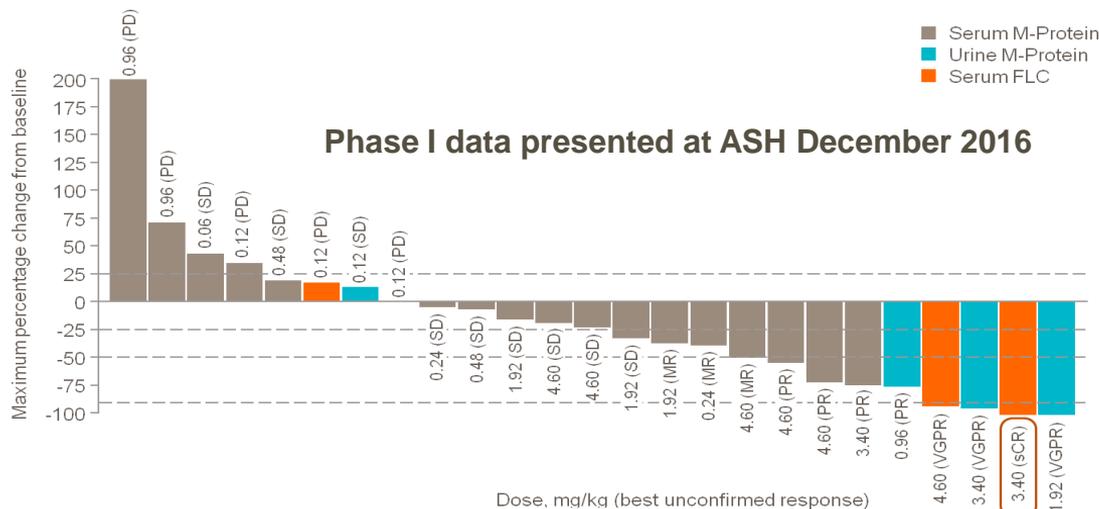


GSK'916: Anti-BCMA-ADC, potential first-in-class next generation therapy for multiple myeloma



- Cell Maturation Antigen
- Antibody Drug Conjugate (ADC) with MMAF (auristatin derivative)
- High-expression target in multiple myeloma
- Immunogenic cell death inducer
- Excellent Phase I efficacy in tough to treat population: ~67% at \geq Phase II dose

All doses: ORR = 8/30* (27%; 95% CI: 12.3%, 45.9%)
 At \geq Ph2 dose 3.4 mg/kg: ORR= 6/9 (66.7%; 95% CI: 0.29, 0.92%)



Safety observations:

Thrombocytopenia, transient

Corneal toxicity: dry eye, blurry vision, reversible

MMAF = Monomethyl auristatin F ; ASH: American Society of Hematology.

*30 patients have been enrolled and included in the denominator of Part 1; only 25 response are shown in the graph as some patients did not have response assessment (missing), or did not have data entered at the time of data cut.

Building capabilities in diseases with clear unmet need



Cell and Gene Therapy

Strimvelis for ADA SCID – first approved ex-vivo stem cell gene therapy

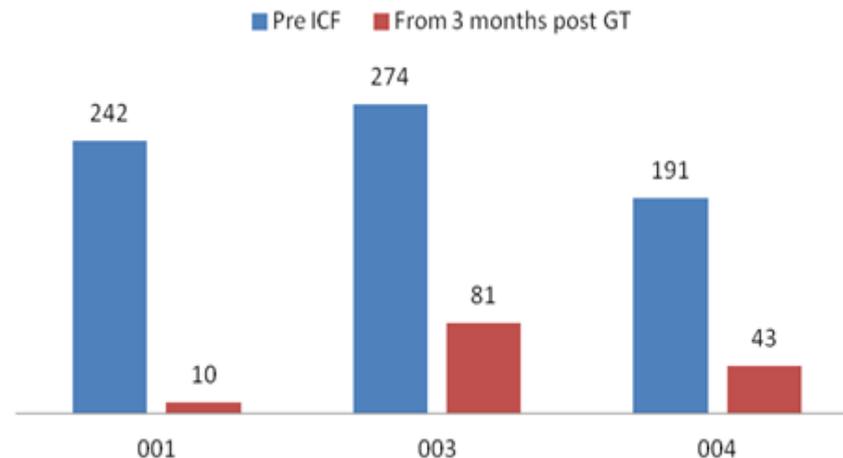
Pipeline of diseases and approaches:

- MLD
- WAS
- Beta thalassaemia**
- NY ESO
- Next gen CAR-T
- TCR

Preliminary data from ongoing study in beta thalassaemia**

- All patients severe genotype, β^0/β^+ (Cod39 / IVS1-110)
- Patients of this genotype make virtually no endogenous beta-globin and require frequent transfusions
- Reduction in beta-globin transfusions of 96%, 70% and 77% observed at data cut off (Nov 2016)

Beta-globin transfusion requirement Yearly ml/kg for each patient



Based on follow up of 1.1 year (patient 001), 6 months (patient 003) and 9 months (patient 004). ICF = Informed Consent Form. GT = gene therapy

**GSK has an exclusive option to in-license the Beta-Thal program from the Hospital San Raffaele (OSR) and the Telethon Foundation (Telethon); Data provided with consent of OSR/Telethon.

Intense period of R&D activity with multiple milestones



Expected Phase III starts by end 2018 include:

daprodustat for anemia (started Q4 16)
cabotegravir + rilpivirine in HIV treatment (started Q4 16)
closed triple asthma (started Q4 16)
cabotegravir for prophylaxis in HIV (started Q4 16)
Long-acting anti-IL5 for severe asthma
Anti-GM-CSF in early/established RA and hand OA
NY ESO-1 in sarcoma
OX40 in a solid tumour

Expected filings by end 2018 include:

dolutegravir + rilpivirine for HIV treatment
dolutegravir + lamivudine for HIV treatment
mepolizumab for EGPA, HES and nasal polyps
Closed triple exacerbation indication
Closed triple for asthma
'728 (TTR) for FAP
MLD (Metachromatic leukodystrophy)

Important clinical readouts by end 2018 inc:

Between 20-30 assets inc oncology & immuno-inflam.
mepolizumab: Phase III for EGPA (Q4 16)
Phase III for COPD
Phase III for HES and nasal polyps
Closed triple: Phase III COPD exacerbations
Phase III asthma
dolutegravir: Phase III dual combo with rilpivirine (Q4 16)
Phase III dual combo with lamivudine
Attachment inhibitor in HIV Phase III
MLD (Metachromatic leukodystrophy)
Shingrix: immunocompromised and revaccination

Expected approvals in 2017:

Shingrix
Closed triple for COPD
Benlysta SC
sirukumab for RA