

## Worldwide Healthcare Trust PLC

Annual General Meeting 18 July 2023

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## Portfolio Managers: Sven Borho & Trevor Polischuk

#### **Sven Borho, CFA**

Managing Partner



Sven H. Borho, CFA, is a founding Partner of OrbiMed. Mr. Borho heads the public equity team and he is the portfolio manager for OrbiMed's public equity and hedge funds.

Mr. Borho has played an integral role in the growth of OrbiMed's asset management activities. Mr. Borho started his career in 1991 when he joined OrbiMed's predecessor firm as a Senior Analyst covering European pharmaceutical firms and biotechnology companies worldwide. He has been a portfolio manager since 1993.

Mr. Borho studied business administration at Bayreuth University in Germany and received a M.Sc. (Econs.), Accounting and Finance, from The London School of Economics.

#### **Trevor Polischuk, PhD**

Partner



Trevor M. Polischuk, Ph.D., is a Partner on the public equity team, focused on the global pharmaceutical industry.

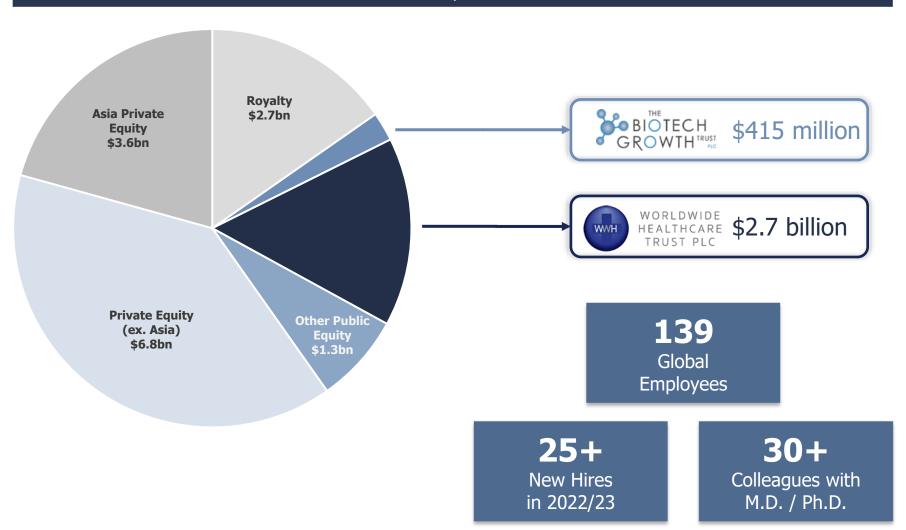
Previously, he worked at Lehman Brothers as a Senior Research Analyst covering the U.S. pharmaceutical industry. Dr. Polischuk began his career at Warner Lambert as a member of the Pharmaceutical Global Marketing Planning team. In this role, he helped coordinate marketing activities for the second generation gabapentinoid blockbuster product, Lyrica (pregabalin).

Dr. Polischuk holds a Doctorate in Neuropharmacology & Gross Human Anatomy from Queen's University and an M.B.A. from Queen's University.



## OrbiMed Corporate Update

#### **Total AUM: \$17.6 billion**



OrbiMed
Healthcare Fund Management

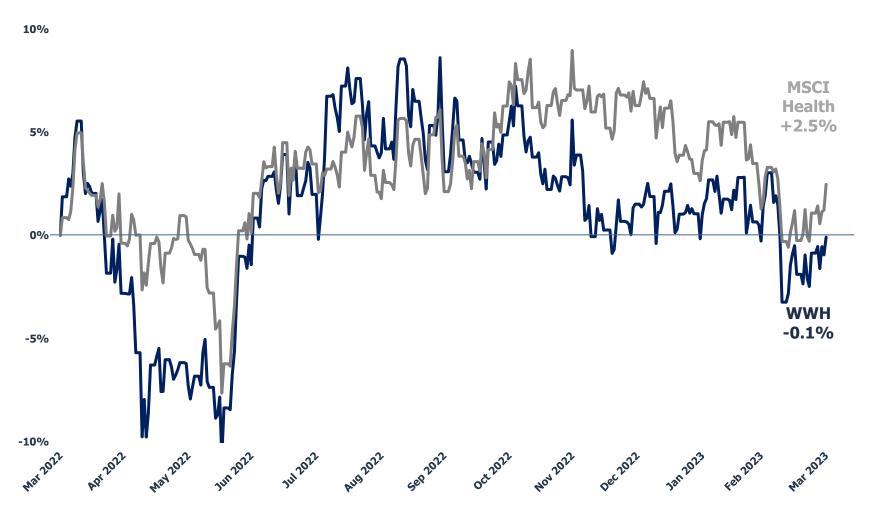


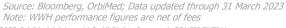
FY2022 Performance (31 March 2023)



## Performance

#### FY2022 (31 March 2023)

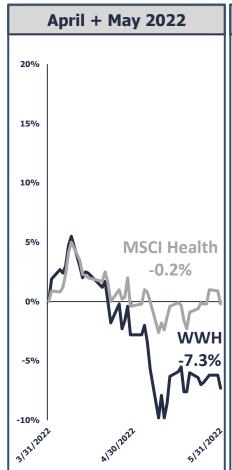


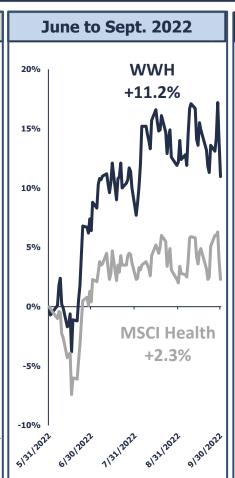


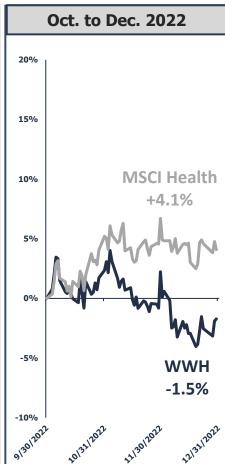


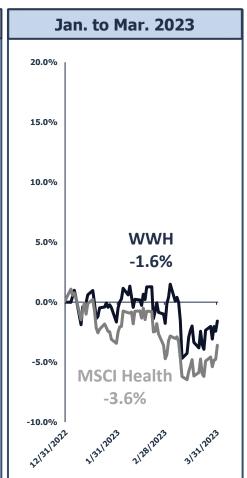
## Performance Segments

#### FY2022 (31 March 2023)





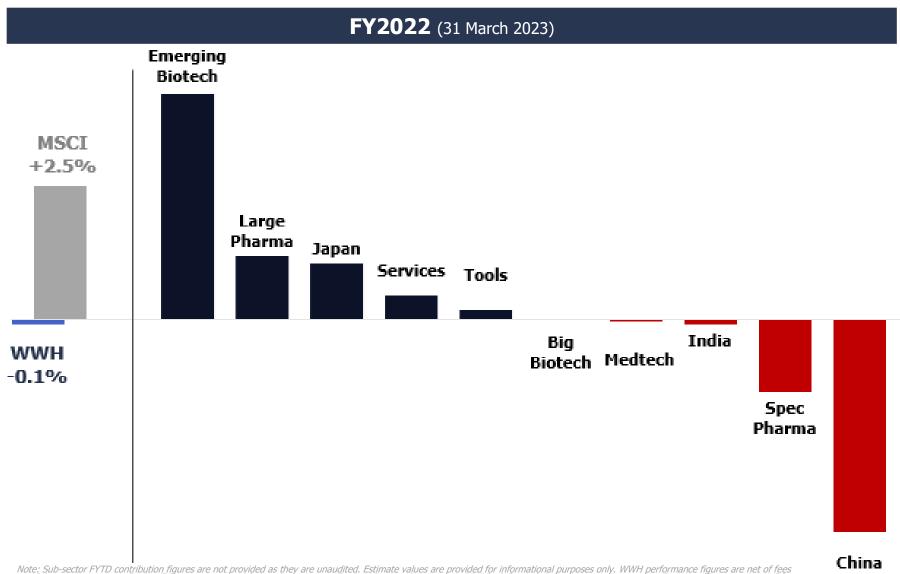




Source: Bloomberg, OrbiMed; Data updated through 31 March 2023

Note: WWH performance figures are net of fees

## Performance: Sources of Contribution (Absolute)



Note: Sub-sector FYTD contribution figures are not provided as they are unaudited. Estimate values are provided for informational purposes only. WWH performance figures are net of fe Source: Bloomberg, OrbiMed



## FY2022: Top 5 Contributors / Detractors

#### **Top 5 Contributors**

	Country	Sector	Contribution £'000	Contribution per share* £
Global Blood Therapeutics**	USA	Biotechnology	30,805	0.5
Seagen**	USA	Biotechnology	28,289	0.4
Daiichi Sankyo	Japan	Pharmaceutical	23,488	0.4
BioMarin Pharmaceutical	USA	Biotechnology	21,598	0.3
Sarepta Therapeutics	USA	Biotechnology	20,665	0.3

#### **Top 5 Detractors**

Edwards Lifesciences	USA	Healthcare Equipment & Supplies	(18,551)	(0.3)
Roche	Switzerland	Pharmaceutical	(24,481)	(0.4)
Horizon Therapeutics**	USA	Biotechnology	(29,324)	(0.5)
Mirati Therapeutics	USA	Biotechnology	(33,332)	(0.5)
Shanghai Bio-heart Biological Technology	China	Healthcare Equipment & Supplies	(42,324)	(0.7)

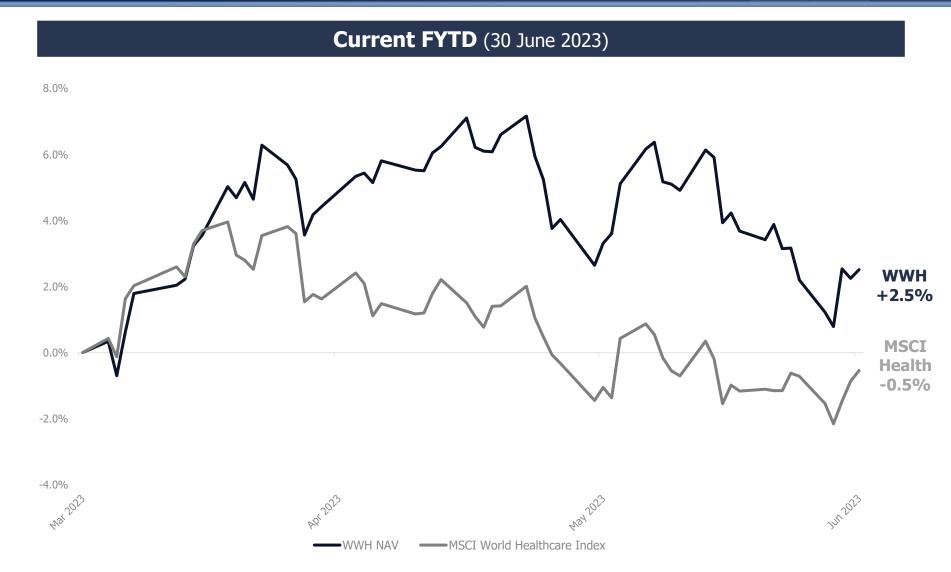
<sup>\*</sup> Calculation based on 64,474,422 shares being the weighted average number of shares in issue during the year ended 31 March 2023



<sup>\*\*</sup> Not held at 31 March 2023

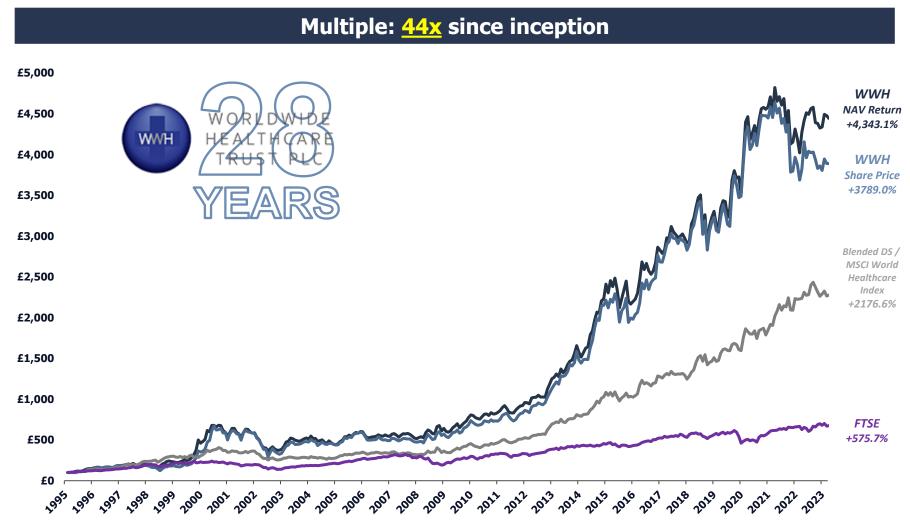
Past performance is no guarantee of future results. See Endnotes for more information. Excludes the effect of Structured Finance and Options and represents fund performance before fees and costs. Total return numbers are estimated and based on an unaudited holdings-based attribution model that excludes trading costs, fees, and expenses.

### Performance Current FYTD





## Performance Since Inception



<sup>\*</sup>The "Blended DS/MSCI Benchmark" uses the Datastream World Pharma/Biotech TR Index from inception to September 30, 2010, the MSCI World Health Care Gross Index from September 30, 2010 to December 31, 2011, and the MSCI World Health Care Net Index from December 31, 2011. Past performance is no guarantee of future results. See Endnotes for additional information regarding the performance presented in this slide and those that follow. Updated through 30 June 2023.





## **2023 Investment Themes**



### 2023 Investment Themes

2023

4

Overhang Lifted: Rx Pricing 2

M&A Accelerating 3

Innovation:
A New Era
of MegaBlockbusters

4

Biotech Under-Performance



## U.S. Drug Price Reform: Overhang Lifted

## Three Main Components to the IRA 2022 re: Drug Price Reform



## **Drug Price Inflation Cap**



Requires drug co.s to pay rebates to Medicare if they increase drug prices faster than inflation



- Price is no longer a revenue driver for pharma
- LIST prices do rise today, but < levels of inflation
- NET prices are cut even further due to rebating
- Curbs bad actors = positive for industry image

#### Medicare Part D Redesign



Removes the "donut hole" and replaces it with an "out-of-pocket cap" of \$2,000 per patient



#### **POSITIVE**

- Lowers out of pocket expenses for patients
- Increases affordability of medicines
- Increases adherence / treatment duration of times
- Increases patient access
- Should result in increased volumes

## **Medicare Price Negotiations**



Enables Health & Human Services to set the prices of certain costly drugs within the Medicare program



- Not in effect until 2026
- Only 10 drugs per year are negotiated
- Targets medicines that are near the end of their life cycles but do not yet have generic competition (9 yrs – small molecules / 13 yrs – biologics)
- Only a potential \$40 billion incremental hit to U.S. Rx revenues to 2032 (out of \$660 billion)

The IRA will have an impact but the net of it is manageable. Perhaps the biggest benefit is that the headline risk is now behind us.



## A Conversation with Dave Ricks – CEO, Eli Lilly & Co.

## Three Main Components to the IRA 2022 re: Drug Price Reform





#### **Drug Price Inflation Cap**

"It is not a major issue...it's really kind of a non-event...it's kind of a **nothing burger** from our perspective."

#### **Medicare Part D Redesign**

"There are (many) changes to part D design, that we saw as **positive** on a net basis."

#### **Medicare Price Negotiations**

"Honestly, it was pretty disappointing..."

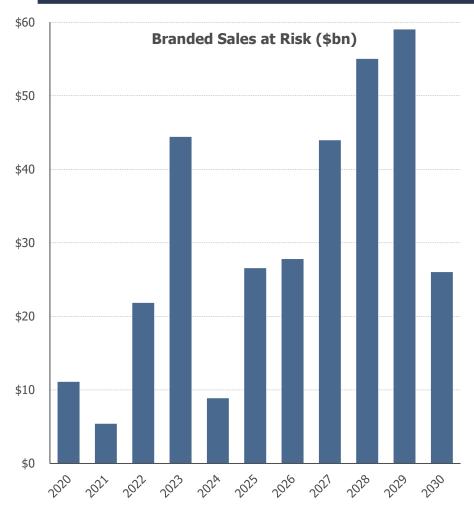
Source: JP Morgan June 2023

The Pharma industry has commenced legal proceedings against HHS



## Large Cap Pharma: Patent Cliff Approaching

### Nearly \$250 billion in branded sales are at risk (2025-2030)



Source: Evaluate	. "The Patent	Winter is	Coming",	May 25, 2022.
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Co.	Drug	US Loss of Exclusivity (Projected)	'22 Global Sales* (\$bn)
abbvie	HUMIRA	2023	\$21.2
MERCK	KEÝTRUDA* (pembrolizumab) <sub>krjecton 100 reg</sub>	2028	\$20.9
Bristol Myers Squibb	Eliquis. (apixaban) tablets 52%,	2026	\$18.2
abbyie Johnson Johnson	imbruvic <mark>ă</mark> *	2027	\$9.8*
Johnson Johnson	Stelara (ustekinumab)	2023	\$9.7
ر <sup>ااا</sup> Bristol Myers Squibb˚	OPDIVO. (nivolumab)	2028	\$8.3
<b>₹</b> Pfizer	IBRANCE palbociclib	2027	\$5.4*
<b>♦</b> MERCK	Januvia	2023	\$5.3*

Source: S&P Global report, data as of February 17, 2022; Company Reports \*2022 sales unless otherwise noted, 2021 sales



## M&A Activity Surging

2022 Acquisitions						
Date	Target	Acquirer	% Premium			
11/29/22	HORIZON	AMGEN	48%			
11/29/22	neo gene	AstraZeneca 🕏	\$320m (private)			
11/21/22	Imago <b>() j</b>	MERCK	107%			
10/23/22	MYOVANT SCIENCES	Sumitomo	50%			
10/18/22	AVEO ONCOLOGY	<b>LG</b> Chem	43%			
10/18/22	VKOUOS	Lilly	121%			
10/03/22	LogicBio	AstraZeneca	667%			
09/01/22	forma THERAPEUTICS.	novo portiek	49%			
08/08/22	GBT <sup>™</sup>	<b>₹</b> Pfizer	102%			
08/04/22	CHEMOCENTRYX	<b>AMGEN</b>	116%			
08/04/22	mır <del>ö</del> bio	<b>GILEAD</b>	\$405m (private)			
07/11/22	<b>⋄</b> VIACYTE	VERTEX	\$320m (private)			
07/11/22	La Jolla Pharmaceutical	INNOVIVA	84%			
07/05/22	TeneoTwo	AstraZeneca 🕏	\$1,270 (private)			
06/17/22	<b>Seagen</b>	MERCK	Rumored			
06/03/22	Turning Point	ر <sup>ااا</sup> ا Bristol Myers Squibb ّ	122%			
05/31/22	<b>♦</b> Affinivax	GSK	\$2,650 (private)			
05/10/22	biohaven pharmaceuticals	<b>₹</b> Pfizer	79%			
04/13/22	SIERRA	GSK	39%			

2023 Acquisitions						
Date	Target	Acquirer	% Premium			
6/29/23	sigilon therapeutics	Lilly	~380%			
6/20/23	DICE	Lilly	~40%			
6/12/23	CHINOOK" THERAPEUTICS	U NOVARTIS	83%*			
5/22/23	VectivBio <sup>▽</sup>	<b>Ironwood</b> °	80%			
5/10/23	CTÎ	SODI	89%			
04/30/23	IVERIC	astellas	23%			
04/25/23	SPECTRUM*	-ASSERTIO=	~94%*			
04/18/23	Bellus HEALTH	GSK	>100%			
04/16/23	Prometheus Biosciences	MERCK	75%			
03/27/23	Jounce	Concentra Biosciences	75%			
03/13/23	<b>Seagen</b>	<b>Pfizer</b>	42%			
03/13/23	proventionbio	sanofi	278%			
01/19/23	CoNCERT Pharmaceuticals Inc.	SUN PHARMA	~67%*			
01/09/23	CINCOR	AstraZeneca	>200%*			
01/09/23	Albireo	SIPSEN Innovation for patient care	>100%			
01/08/23	AMRYT	<b>⊕</b> Chiesi	>100% * With CVR			

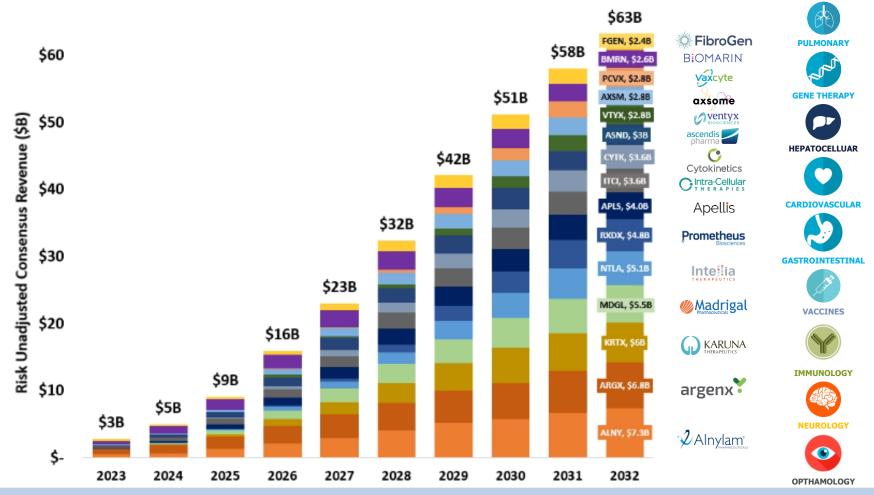
M&A: Material Inflection YOY				
	# of Deals	\$ Value of Deals		
FY 2021	17	\$31 billion		
FY 2022	30	\$113 billion		
YOY	+76%	+265%		

Source: FactSet, Company Websites Note: Public companies shown here are not necessarily representative of portfolio holdings

<sup>\*</sup>Represent announced transactions where the target was a publicly traded company. Source: FactSet, data as of June 30, 2023

## M&A: "Filling the Gap"

#### 15 Novel Biotech-Sourced Launches with >\$60 billion in Cumulative Revenues

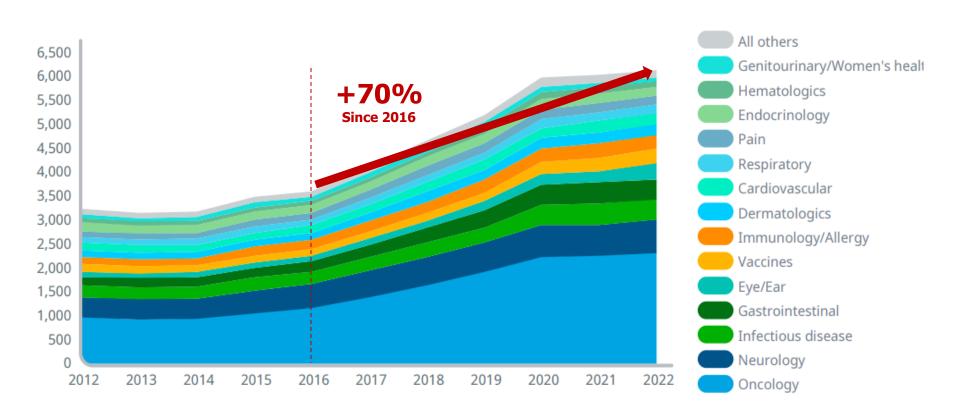


Stretching the list to 25 products could fill an additional >\$20 billion in revenues.



## Innovation: Pipelines are Full

#### **Record Number of Molecules Under Development**



Source: IQVIA Pipeline Intelligence, Dec 2022; IQVIA Institute, Jan 2023

**Pipeline Products:** (Phase I  $\rightarrow$  Filing / Therapeutic Class / 2012-2022)

## Innovation: The Next Big Thing?

Oncology



**ADCs** 



Bristol Myers Squibb"

Approved

**TIGIT** 

Roche)

Awaiting the final analysis in lung cancer, expected 3Q23

AstraZeneca 2

ADAURA - ASCO '23

argeted Therapy

51% reduction in risk of death in Adj. lung cancer

orals

**Obesity** 



tirzepatide

New Standard of Care in

HER2+ "low" metastatic BC



AstraZeneca

GGG

Approved

semaglutide

wegovy\* semaglutide injection 2.4 mg

SELECT NEXT GENERATION

Approved







Positive data across multiple oral tx's at ADA 2023

donanemab

**Alzheimer's** 



antenerumab

**GRADUATE 1/2 CLINICAL TRIALS** 





Patients on drug improved; but outcome was not SS



lecanemab



Met all endpoints & reduced cognitive decline by -27%

**TRAILBLAZER** 



Met all endpoints & reduced cognitive decline by -36%

Note: Examples may not be representative of portfolio holdings.



## New Era of Mega-Blockbuster Drugs?

2021					
Revenue	Drug	Company			
>\$30bn	COMIRNATY* (CDVID-19 Vaccine, mRNA)	<b>P</b> fizer			
	HUMIRA adalimumab	abbvie			
	KEYTRUDA° (pembrolizumab) vijection 100 mg	MERCK			
>\$20bn	spikevax~ COVID-19 Vaccine, mRNA	moderna			
	PAXLOVID™ (nirmatrelvir tablets   ritonavir tablets)	<b>P</b> fizer			
	Eliquis。 (apixaban) tablets ﷺ	الله Bristol Myers Squibb			
>\$10bn	Revilmid' (lenalidomide).cocobs	( Bristol Myers Squibb			
	imbruvica. (ibrutinib)	abbvie			
	Stelara® (ustekinumab)	Johnson Johnson			

#### 2030 (OrbiMed Estimates)

Drug Company





















milvexian





















## Biotech Underperformance

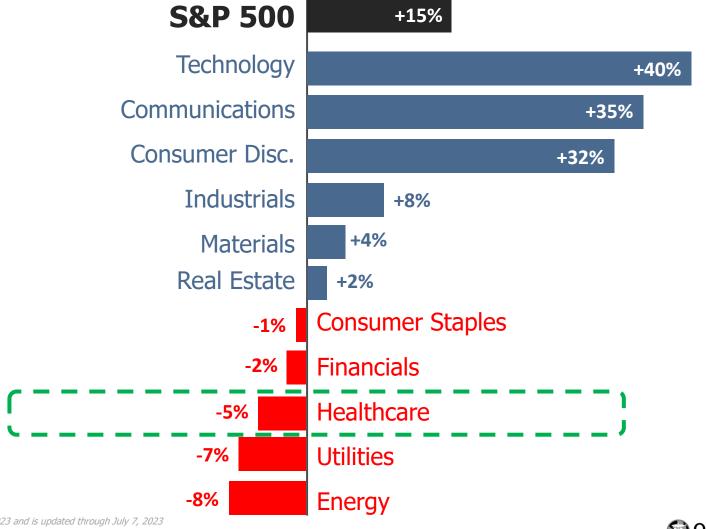


Historic underperformance of Biotechnology stocks has continued in 2023



### S&P 500 Performance CYTD

#### **S&P Constituents by Industry** (7 July 2023)



Source: Bloomberg Note: Chart begins on Jan 1, 2023 and is updated through July 7, 2023

OrbiMed



## **Portfolio & Positioning**



# Worldwide Healthcare Trust Holdings 30 June 2023

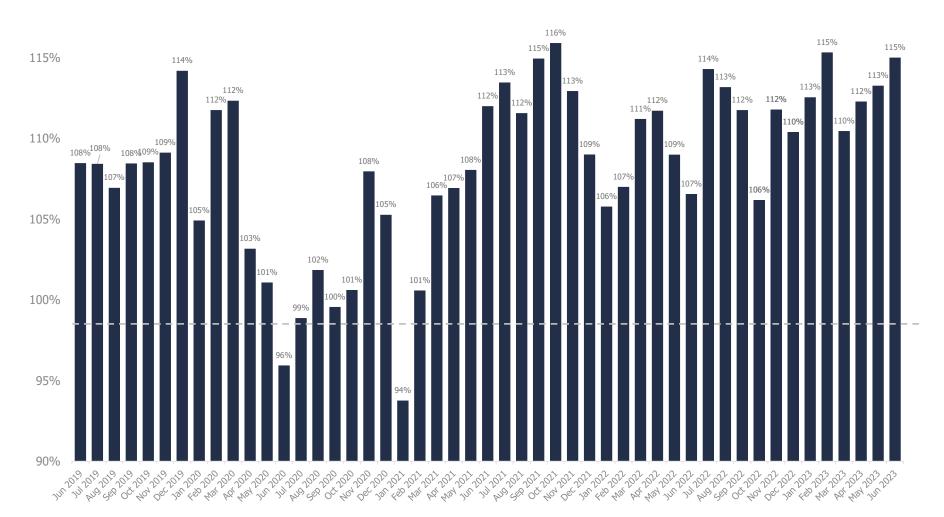
Subsector	WWH %NAV	MSCI World HC	Delta
Pharmaceuticals	26.2	43.5	(17.3)
Big Pharma	25.9	40.3	(14.5)
Spec Pharma/Generics	0.3	0.3	3.2
Biotechnology	28.2	8.7	19.5
Big Biotech	3.2	4.6	(1.4)
Emerging Biotech	25.0	4.1	20.9
Life Science Tools	3.6	11.4	(7.8)
Medtech/Devices	22.6	17.2	5.4
Healthcare Services	15.7	15.0	0.7
Japan	6.3	4.2	2.1
Emerging Markets	5.7	0.0	5.7
Structured Finance	0.0	0.0	0.0
Privates	6.7	0.0	6.7
Total	115.0	100.0	15.0

<sup>\*</sup>Excludes Options. Basket constituent positions are allocated to their respective subsectors. Future weightings may differ.



# WWH Leverage over Time As of 30 June 2023

120%



Source: OrbiMed, Bloomberg PORT

# Worldwide Healthcare Trust Holdings 30 June 2023

	Market Price \$ Millions	Pct. Value		Market Price \$ Millions	Pct. Value		Market Price \$ Millions	Pct. Value
Biotechnology	¥		Healthcare Services / Distributors	¥		Medtech	¥	
Apellis Pharmaceuticals Inc	55.1	2.0	Cigna Group/The	36.9	1.4	Baxter International Inc	120.2	4.4
Biogen Inc	81.0	3.0	Evolent Health Inc	60.3	2.2	Boston Scientific Corp	139.4	5.1
BioMarin Pharmaceutical Inc	83.8	3.1	Humana Inc	119.0	4.4	Edwards Lifesciences Corp	42.4	1.6
Catalyst Basket	31.6	1.2	R1 RCM Inc	46.4	1.7	Intuitive Surgical Inc	171.7	6.3
Ionis Pharmaceuticals Inc	51.6	1.9	Tenet Healthcare Corp	61.0	2.2	SI-BONE Inc	40.5	1.5
Iovance Biotherapeutics Inc	20.1	0.7	UnitedHealth Group Inc	101.9	3.8	Stryker Corp	72.2	2.7
Madrigal Pharmaceuticals Inc	48.7	1.8		425.6	15.7	Zimmer Biomet Holdings Inc	27.8	1.0
M&A Basket	140.3	5.2	Emerging Markets				614.4	22.6
Mirati Therapeutics Inc	22.8	0.8	Apollo Hospitals Enterprise Lt	17.1	0.6	Privates		
Neurocrine Biosciences Inc	35.9	1.3	Jiangsu Yuyue Medical Equipmen	28.9	1.1	API Holdings	17.6	0.6
Sarepta Therapeutics Inc	44.5	1.6	New Horizon Health Ltd	26.7	1.0	Caris	47.5	1.8
uniQure NV	15.4	0.6	Shanghai Kindly Medical Instru	24.4	0.9	Crossover	21.4	0.8
United Therapeutics Corp	49.4	1.8	WuXi AppTec Co Ltd	16.6	0.6	Edda	18.9	0.7
Vaxcyte Inc	24.0	0.9	WuXi AppTec Co Ltd	16.5	0.6	RiMAG	14.2	0.5
Vertex Pharmaceuticals Inc	39.1	1.4	Wuxi Biologics Cayman Inc	14.1	0.5	Ruipeng Pet Group	14.7	0.5
Xenon Pharmaceuticals Inc	13.8	0.5		155.2	5.7	Visen	15.8	0.6
	774.6	28.5	Life Science Tools & Services			Yuanxin Tech	18.5	0.7
Pharmaceuticals			Natera Inc	31.4	1.2		182.1	6.7
AstraZeneca PLC	160.2	5.9	Thermo Fisher Scientific Inc	66.5	2.4			
Bristol-Myers Squibb Co	81.0	3.0		97.9	3.6	Structured Finance & Privates	182.1	6.7
Daiichi Sankyo Co Ltd	82.0	3.0						
Eisai Co Ltd	89.1	3.3				Cash	-407.7	-15.0
Eli Lilly & Co	76.0	2.8						
GSK PLC	27.1	1.0				Total Portfolio	2715.5	100.0
Novo Nordisk A/S	124.5	4.6						
Roche Holding AG	112.5	4.1						
Sanofi	121.2	4.5				<u>wwh %nav</u>	WWH Active Share%	

Note: position summaries above excludes positions <0.50% of NAV

873.6

32.2

	<u>WWH %NAV</u>	WWH Active Share%
Top 10	48%	70%
Top 15	64%	7070
Top 20	76%	WWH % Equity Book [Companies with
Top 30	93%	Positive Net Income]
		78%



## Playbook for 2023

1	Continue to Invest in Innovation	
2	Key Overhangs Lifted	
3	Continued High Expectations for M&A	
4	Re-ignite our Catalyst Strategy	
5	Great Entry Point for Biotech	ENTER HERE
6	Valuations Undemanding	\$
7	Overall Bullish Outlook	A



### **Endnotes**

#### **General Notes**

- 1. The information presented herein relates to Worldwide Healthcare Trust PLC (the "Fund"). OrbiMed Capital LLC ("OrbiMed") is an investment adviser registered with the U.S. Securities and Exchange Commission (the "SEC") that specializes in the investment of clients' assets, including the Fund's assets, in healthcare and life sciences companies, including the biotechnology and pharmaceutical sectors, across a number of products and strategies. This presentation includes information specifically relating to the Fund, and potential OrbiMed clients or fund investors should be aware that such information may not be applicable to other OrbiMed funds, products or strategies. The information contained in this presentation is not intended to supplement or replace the disclosures made in Part 2 of OrbiMed's Form ADV filed with the SEC or in the prospectus or other offering document for any investment fund sponsored and/or managed by OrbiMed or its affiliates. SEC registration does not imply a certain level of skill or training.
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- 6. An investment in the Fund is speculative and involves a high degree of risk. The Fund is an investment trust listed on the London Stock Exchange. Investment trusts have the ability to borrow to invest, which is known as gearing. Companies with higher gearing are subject to higher risks, and therefore the investment value may change substantially. The value and the performance of the net asset value of the Company may not be the same as its share price. In considering an investment in the Fund, prospective investors should also be aware of certain other special considerations and risk factors, including those related to: the risk of deterioration in the financial markets in general and the healthcare sector in particular; the risk that the Fund could incur losses due to the failure of third parties to perform their contractual commitments to the Fund or financial difficulties on the part of brokerage firms, banks or other financial institutions that hold assets of the Fund; risks arising from the operation of pooled investment vehicles like the Fund; and tax-related risks arising from the status and operation of the Fund. See the Fund Documents for more detail on those and other risks related to investing in Shares.
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## **Endnotes** (continued)

#### **Indices**

Information about indices is provided to allow for comparison of the performance of the Shares to the Fund's benchmark and certain other recognized indices. Investors cannot invest directly in an index, which also does not take into account trading commissions and costs. The indices shown are unmanaged, do not charge fees or expenses and do not employ special techniques such as leveraging or short selling. The volatility of indices may be materially different from the performance of the Fund. In addition, the Fund's portfolio holdings may differ significantly from the securities that comprise such indices.

The MSCI World Health Care Index is a market-value weighted index of health care sector equities within the MSCI World Index, a free float-adjusted market capitalization weighted index that is designed to measure the equity market performance of developed markets. Index data for the MSCI World Health Care Index is available since December 30, 1994. For the period from the Fund's inception through December 30, 1994, returns of the MSCI World Health and Personal Care Index are used as a proxy for the MSCI World Health Care Index. Return numbers for these indices include reinvestment of dividends net of withholding taxes, as calculated by the index provider.

