

The Biotech Growth Trust PLC

Annual General Meeting

17 July 2025

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Portfolio Managers: Geoffrey Hsu & Josh Golomb

Geoffrey Hsu, CFA

General Partner



Geoffrey C. Hsu, CFA, is a General Partner at OrbiMed and has been portfolio manager of The Biotech Growth Trust since May 2005.

Mr. Hsu joined OrbiMed in 2002 as a biotechnology analyst. Prior to joining OrbiMed, he worked as a financial analyst in the healthcare investment banking group at Lehman Brothers. Mr. Hsu received his A.B. degree summa cum laude from Harvard University and holds an M.B.A. from Harvard Business School. Prior to business school, he spent two years studying medicine at Harvard Medical School. Josh Golomb Partner



Josh Golomb is a Partner at OrbiMed focused on biotechnology. He joined the firm in 2021 and became a portfolio manager of The Biotech Growth Trust in 2023.

Mr. Golomb has covered the biotechnology and pharmaceutical sectors at investment firms since 2004. He was most recently employed at HealthCor, a healthcare-dedicated hedge fund, from 2016 to 2021 as a senior biotechnology and pharmaceutical analyst. Previously, he covered biotech at Millennium, Citadel, Diamondback and ProMed Partners. Mr. Golomb received a B.S. in Operations Research with a concentration in Engineering Management Systems from the Fu Foundation School of Engineering and Applied Science at Columbia University.

OrbiMed Corporate Update

Global Healthcare-Dedicated Investment Firm with Over 25 Years of Experience



Source: OrbiMed AUM data estimate as of 30 June 2025. Employee data as of 30 June 2025.

OrbiMed: Extensive Resources to Invest in Biotech

Differentiated platform capable of delivering strong returns

OrbiMed's capabilities are well-suited to achieving the fund objective of generating capital appreciation by investing in biotech companies worldwide:

- Global footprint
 - 12 locations worldwide including New York, San Francisco, Hong Kong, Shanghai, and London
 - can invest in the U.S., Europe, and Asia; provides geographic diversification
- Large research team
 - 6 dedicated biotech analysts (including 2 PMs), with 60+ years of collective experience
 - additional expertise from 7 other public equity analysts covering other healthcare sectors
- Preferential access to deal flow
 - firm gets wall-crossed on 3-4 deals per week (not all investors have access to this deal flow)
 - potential opportunity to invest based on confidential information before public release
 - in some cases, warrants and other deal sweeteners can be present to enhance returns
- Private equity business
 - dedicated team of 50 professionals provides preferential access to crossover opportunities and IPOs in OrbiMed companies; BIOG has the option, but not the obligation, to participate
 - private equity team aids in diligence for both IPOs and other publicly-traded names
 - regular interactions with large pharma provide insight into areas of M&A interest
- Extensive experience
 - healthcare specialist firm with over 25 years of experience investing in biotech

BIOG

Performance

BIOG Performance for Fiscal 2025 and Beyond



Source: Frostrow, Bloomberg. Note: See Endnotes for additional information, including with regard to the calculation of these results and the index shown above.

Market Cap Performance Divergence in Biotech

Small and Mid Cap stocks underperformed during the fiscal year



Note: Chart shows equal-weighted performance of NBI stocks in their respective market cap buckets, using market cap classifications as of 31 March 2024. Updated as of 31 March 2025, performance calculated in USD.

Commercial vs Development Stage Biotech

Pre-revenue companies have significantly underperformed their commercial peers



Clinical Stage Biotech Basket

Commercial Biotech Basket

Biotech Underperformance vs S&P 500

(31 March 2021 – 30 June 2025)



Biotech Valuations At Unprecedented Lows



2001 2002 2003 2004 2005 2006 2007 2008 2009 2010 2011 2012 2013 2014 2015 2016 2017 2018 2019 2020 2021 2022 2023 2024 2025

Note: Monthly chart of all U.S. GICs defined biotechnology greater than \$10mm, using historical cash and debt sourced from Bloomberg, with annual GICS biotechnology universe refreshes. Updated through 30 June 2025.

Biotech Valuations At Unprecedented Lows

Valuations have yet to recover meaningfully from historical lows



Note: Monthly chart of all U.S. GICs defined biotechnology greater than \$10mm, using historical cash and debt sourced from Bloomberg, with monthly GICS biotechnology universe refreshes. Updated through 30 June 2025

BIOG Performance since Inception

18 May 2005 through 30 June 2025



Note: See Endnotes for additional information, including with regard to the calculation of these results and the index shown above. Source: Frostrow, Bloomberg. BIOG

Investment Themes

Key Breakthroughs in Biotech since Last AGM

Company	Indication	Product (MOA)	Highlights	
Summit Akesobio	Non-Small Cell Lung Cancer	Ivonescimab (novel PD1/VEGF bispecific antibody)	Reported positive HARMONI-2 trial results with a 49% reduction in disease progression relative to Keytruda, the standard of care	
GILEAD Pre-Exposure Prevention of HIV		Lenacapavir	Prevented 100% and 99.9% of HIV infections in trials with individuals engaging in high-risk behavior.	
insmed	Bronchiectasis	Brensocatib (DPP1 inhibitor)	Positive Phase 3 data with 20% reduction in exacerbations over 52 weeks. First ever successful bronchiectasis study.	
Scholar Spinal Muscular Atrophy (SMA)		Apitegromab (monoclonal antibody)	First drug to show a benefit in SMA patients on top of standard care in its Phase 3 trial	
Metabolic Dysfunction-Associated Steatohepatitis (MASH)		Efruxifermin	Announced Phase 2b data that showed a reversal of compensated cirrhosis due to MASH. No other drug tested to date has shown an improvement in late- stage liver disease.	
	Facioscapulohumeral Muscular Dystrophy (FSHD)		First drug to ever demonstrate DUX4 knockdown and muscle function improvement in a Phase 1/2 trial	
axsome	Alzheimer's Disease Agitation	AXS-05	First non-antipsychotic drug to show a benefit in this hard-to-treat patient population in a Phase 3 trial	

Note: Public companies shown here are not necessarily representative of portfolio holdings. Source: Company websites

Significant FDA Biotech Approvals Since Last AGM

Company	Indication	Product (MOA)	Highlights	
UroGen [.] Pharma	Low-grade intermediate-risk non- muscle invasive bladder cancer (LGIR NMIBC)	Zusduri	First and only medication for recurrent LGIR NMIBC	
VERTEX	Moderate to severe acute pain	Journavx	First and only non-opioid oral pain signal inhibitor	
ascendis pharma	Hypoparathyroidism	Yorvipath	First and only FDA-approved treatment for low parathyroid hormone in adults	
Syndax 🌮	Acute leukemia with a lysine methyltransferase 2A gene (KMT2A)	Revuforj	First menin inhibitor approved to treat KMT2A AML and ALL	
EXEVRA	Niemann-Pick disease type C (NPC)	Miplyffa	First and only therapy that can be combined with miglustat for NPC	
S NEUROCRINE® BIOSCIENCES	NEUROCRINE® Congenital adrenal hyperplasia (CAH)		First and only non-steroidal treatment to control androgen levels in classic CAH	
IONIS	Familial chylomicronemia syndrome (FCS)		First and only approved medicine for FCS	
PTC THERAPEUTICS	Aromatic L-amino acid decarboxylase deficiency (AADC)		First and only gene therapy for AADC	
X Adaptimmune	Synovial sarcoma	Tecelra	First and only cell therapy therapy for synovial sarcoma	

Note: Public companies shown here are not necessarily representative of portfolio holdings. Source: Company websites

Oncology

Cancer remains #1 therapeutic area for biotech drug development given unmet need



Multiple innovative approaches generating clinical benefit across tumor types, giving physicians more treatment options to extend the lives of cancer patients.

Note: Public companies shown here are not necessarily representative of portfolio holdings. Source: Company websites

Outlook for Oncology

Moving cancer to the "chronic disease" category and advancing more cures



...so much that "One Size Fits One"

Outlook for Gene and Cell Therapy

An explosion of new gene therapy medicines – all with curative applications

Today there are only 30 gene and cell therapy products approved by U.S. FDA*...



Gene Therapy Cell Therapy Gene Editing CRISPR Technologies

Safety Efficacy Administration



All with curative applications

...this could increase by 10x over the next 30 years.

*excluding blood products Source: https://www.fda.gov/vaccines-blood-biologics/cellular-gene-therapy-products/approved-cellular-and-gene-therapy-products

Innovation Well Reflected in BIOG

Portfolio has exposure to a wide swath of novel technologies, as shown below:



Source: OrbiMed, % of BIOG NAV as of 30 June 2025. Some positions are double-counted because they use more than one technology.

FDA New Molecular Entity Approvals



Source: FDA CDER and CBER as of 12/31/24

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M&A Activity in Biotechnology



Source: Factset; M&A in BIOG reflects acquisitions from 2023-2025; Bloomberg Transcripts, quotes from 1Q24 Earnings Calls; biopharmadive.com

Big Pharma Patent Cliff Drives Biotech M&A



Note: Data from Wolfe Research, OrbiMed, Visible Alpha

(sales at risk represents consensus projected sales for year prior to year of expected loss of exclusivity)

Source: S&P Global report, Company Reports

Political Environment Overall Positive

What can we expect from President Trump's second term?

POSITIVES

A&M

INFLATION REDUCTION ACT

Trump issued an executive order to extend pricing exclusivity for small molecule drugs to 13 years (from 9).

Removal of Lina Khan as FTC Commissioner should pave the way for more biotech M&A.

INNOVATION administration is gen

Trump's administration is generally proinnovation, consistent with his broader domestic pro-business agenda.

POLITICAL SUPPORT

Trump administration wants domestic biotech industry to thrive. Senate report recently recommended further government support of biotech as a strategically important industry.

TAX CUTS

The industry benefited from Trump's previous corporate tax policy initiatives.

REDUCED REGULATION

FDA wants to reduce regulatory hurdles to drug approvals and speed up the drug development process.

UNCERTAINTIES

RFK Jr.

Noted vaccine skeptic as head of HHS may make vaccine development more difficult, but drugs likely not affected; FDA Commissioner Martin Makary seems science-based and reasonable

BUDGET CUTS

Trump has cut some government spending in order to pay for his tax cuts; FDA staffing has been cut, but HHS has no intention of cutting drug reviewers

DRUG PRICE REFORM

Trump's "most favored nation" pricing plan for drugs unlikely to pass Congress; pharmaceutical tariffs to be announced soon; biotech should be able to manage through both

Positives outweigh the uncertainties for Trump's second term

2025 Strategy and Outlook

Strategy to emphasize emerging biotech over large cap biotech will continue in order to fully capture sector rebound

- Will capitalize on compelling investment opportunities in companies that need to finance at low share prices
- o Portfolio has shifted to a more balanced allocation between small caps and mid caps
- Gearing level will generally stay between 5-10%



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Unprecedented low valuations in emerging biotech suggest significant upside from current levels

- Performance drawdown of small/mid cap biotech versus the S&P 500 has been the longest and most severe we have ever seen
- Percent of biotech companies with market caps below the net cash on their balance sheets well above historical norms
- Valuation levels are lower than the Great Financial Crisis and the bursting of the Dot Com bubble



Valuation contraction is disconnected from the fundamental innovation in the sector, which remains robust



We expect M&A activity to accelerate given lower valuations of biotech targets and urgent need for new products by Big Pharma



Trump administration supportive of the biotech industry despite some uncertainties

OrbiMed believes this is an excellent time to invest in biotech innovation at unprecedented low valuations.

BIOG

Appendix

BIOG Holdings As of 30 June 2025

UNITED STATES Emerging Biotechnology	Market Price \$ Millions	Pct. Value
Agios Pharmaceuticals Inc	8.2	3.0
Akero Therapeutics Inc	5.4	2.0
Alnylam Pharmaceuticals Inc	12.9	4.7
Alto Neuroscience Inc	0.6	0.2
Amicus Therapeutics Inc	6.6	2.4
Amylyx Pharmaceuticals Inc	2.0	0.7
Avidity Biosciences Inc	11.6	4.2
Axsome Therapeutics Inc	11.0	4.0
C4 Therapeutics Inc	2.5	0.9
CG oncology Inc	13.0	4.7
Catalyst Pharmaceuticals Inc	9.0	3.3
Corbus Pharmaceuticals Holding	2.7	1.0
Cullinan Therapeutics Inc	3.1	1.1
Cytokinetics Inc	2.3	0.8
Dyne Therapeutics Inc	2.6	0.9
Edgewise Therapeutics Inc	3.5	1.3
Engene Holdings Inc	1.1	0.4
Enliven Therapeutics Inc	0.8	0.3
Forte Biosciences Inc	6.9	2.5
Insmed Inc	1.4	0.5
Janux Therapeutics Inc	5.3	1.9
Kezar Life Sciences Inc	0.9	0.3
Korro Bio Inc	0.4	0.1
Krystal Biotech Inc	1.3	0.5
Mineralys Therapeutics Inc	2.6	0.9
Natera Inc	7.8	2.8

UNITED STATES	Market Price \$ Millions
Emerging Biotechnology (cont.)	5.0
Neurocrine Biosciences Inc	5.0
Prelude Therapeutics Inc	0.3
Repare Therapeutics Inc	0.1
Rhythm Pharmaceuticals Inc	5.0
Roivant Sciences Ltd	7.9
Structure Therapeutics Inc	5.2
Tarsus Pharmaceuticals Inc	2.7
Trevi Therapeutics Inc	5.5
Tyra Biosciences Inc	2.5
UroGen Pharma Ltd	6.6
Vir Biotechnology Inc	4.8
Xenon Pharmaceuticals Inc	10.0
	181.1
UNITED STATES	
Major Biotechnology	
Vertex Pharmaceuticals Inc	29.7
	29.7
United States Total	210.8

Pct.		Market Price	Pct.
Value	INTERNATIONAL	\$ Millions	Value
	Europe		
1.8	Emerging Biotechnology		
0.1	ADC Therapeutics SA	5.7	2.0
0.0	Argenx SE	22.0	7.9
1.8	Ascendis Pharma A/S	5.9	2.1
2.8	Immatics NV	3.9	1.4
1.9		37.5	13.5
1.0			
2.0	Europe Subtotal	37.5	13.5
0.9			
2.4	Far East		
1.7	Emerging Biotechnology		
3.6	Abbisko Cayman Ltd	5.7	2.0
65.1	Akeso Inc	6.8	2.5
	Cutia Therapeutics	1.4	0.5
	HUTCHMED China Ltd	2.5	0.9
10.7	Gracell Biotechnologies Inc	0.5	0.2
10.7	ImmuneOnco Biopharmaceutica	1.0	0.4
	Laekna Inc	2.7	1.0
75.8	LakeShore Biopharma Co Ltd	0.1	0.0
	New Horizon Health Ltd	1.1	0.4
	OrbiMed Asia Partners*	1.2	0.4
	Shanghai Henlius Biotech Inc	6.3	2.3
	Suzhou Basecare Medical Corp L	0.7	0.3
	Zai Lab Ltd	12.9	4.6
		42.8	15.4
	Major Biotechnology		
	CSPC Pharmaceutical Group Ltd	5.2	1.9
	Jiangsu Hengrui Pharmaceutical	12.3	4.4
		17.5	6.3
	Far East Subtotal	60.3	21.7
	International Total	97.8	35.2
	Cash	-30.6	-11.0
	Total Portfolio	278.0	100.0

Note: securities with asterisks (*) are unquoted

10-Year U.S. Government Yield Still Elevated

(USGG10YR Index) Data as of 30 June 2025





China Rapidly Advancing in Biotechnology



- Ever since China announced in 2015 that developing a domestic biotechnology industry was a priority in its 10-year "Made in China 2025" plan, biotechnology innovation in China has grown dramatically, as shown by China's clinical trial market share above
- Chinese companies' R&D capabilities are now on par, and in some cases better, than their U.S. peers
- They can conduct clinical trials faster and cheaper than their Western counterparts, shortening development times
- Potential new Pharma business model emerging: in-license a Chinese asset, conduct Phase 1-2 studies in China to maximize speed at low cost; conduct global Phase 3 pivotal trials to garner U.S. and European approval

Source: Citeline Trialtrove, Jan 2025; IQVIA Institute, Jan 2025

Selected Chinese Biotech Licensing Deals in 2024-2025

Pfizer in-licensing a PD-1/VEGF drug candidate from 3SBio with an upfront payment of USD1.25 billion marks a new high for China business development deals

Time	Asset	Target	Licensor	Licensee	Upfront Milestones (\$million) (\$million)	;
14-Jun-24	FG-M701	TL1A	FutureGen	AbbVie	150	1,560
14-Jun-24	Olverembatinib	BCR-ABL1	Ascentage Pharma	Takeda	100	1,200
13-Aug-24	CN201	CD3/CD19 TCE	Curon Biopharmaceutical	Merck	700	600
30-Sep-24	RGT-419B	CDK2/4	Regor Therapeutics	Roche	850	-
7-Oct-24	YS2302018	Lp(a)	CSPC Pharmaceuticals	AstraZeneca	100	1,920
17-Oct-24	Botanical cancer SM	N/A	Chengdu Baiyu	Novartis	70	1,100
30-Oct-24	CBM1A46	CD3/CD19/CD20	Chimagen Biosciences	GlaxoSmithKline	300	550
12-Nov-24	Acqu	uisition	Biotheus	BioNTech	800	150
15-Nov-24	LM-299	PD-1/VEGF	LaNova Medicines	Merck	588	2,700
18-Dec-24	HS-10535	Oral GLP-1R agonist	Hansoh Pharma	Merck	112	1,900
2-Jan-25	IBI3009	DLL3 ADC	Innovent Biologics	Roche	80	1,000
13-Jan-25	SIM0500	GPRC5D/BCMA/CD3	Simcere Zaiming	Abbvie	-	1,055
21-Mar-25	Biospecifics stra	tegic collaboration	Harbour BioMed	AstraZeneca	175	4,400
24-Mar-25	UBT251	GLP-1/GIP/GCGR	United Laboratories	Novo Nordisk	200	1,800
25-Mar-25	HRS-5346	Lp(a)	Jiangsu Hengrui	Merck	200	1,770
7-Apr-25	SHR-7280	RnRH	Jiangsu Hengrui	Merck KGaA	17	-
20-May-25	SSGJ-707	PD-1/VEGF	3SBio	Pfizer	1,250	4,800
20-May-25	SRSD107	FXI siRNA	Sirius	CRISPR	95	8,000
2-Jun-25	HS-20094	GLP-1/GIP	Hansoh Pharma	Regeneron	80	1,930
13-Jun-25	AI-based drug c	liscovery platform	CSPC Pharmaceuticals	AstraZeneca	110	5220

Note: Public companies shown here are not necessarily representative of portfolio holdings. Source: OrbiMed and company reports

China Healthcare Recovering From All-Time Lows

Innovation remains strong; recovery starting to occur



Source: Bloomberg, OrbiMed Note: Chart updated from 30 June 2021 through 30 June 2025.

Risk Management

- Diversification across multiple parameters
 - Size and stage of company (large cap, mid cap, small cap); revenue versus pre-revenue
 - Geographical diversification (U.S., Europe, China)
 - Therapeutic area
 - Drug development technologies
- Prudent position sizing
 - Sizing depends on conviction level, liquidity of the stock, and risk/reward of the idea
 - We aim not to lose more than 100 basis points of performance on a single catalyst
 - Typical position sizes: 5-10% for large cap biotech, 1-3% for emerging biotech
 - Position sizes are actively managed to navigate the catalyst paths of emerging biotech stocks, so turnover is relatively high (~90%)
 - The fund regularly trades around positions (trimming high, adding low) to ensure position sizes appropriately reflect risk/reward, especially in advance of binary events
 - Volatility of biotech increases markedly as one descends in market cap
 - Average annual volatility of stock constituents of NASDAQ Biotech Index:

As of 30 June 2025	1 Std. Deviation	
Large Cap (>\$10bn)	44.2	
Mid Cap (\$2bn-10bn)	57.9	
Small Cap (<\$2bn)	84.6	

• Fund's emphasis on emerging biotech with stock-moving catalysts means turnover will likely remain high to manage risk appropriately

Source: Bloomberg

Endnotes

General Notes

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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 fund 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.

No guarantee or representation is made that a fund will achieve its investment objective or that the fund's risk management strategies will be successful. The fund's performance may be volatile, and investors in the fund could lose all or a substantial amount of their investment. In addition, the fees and expenses charged in connection with an investment in the fund may be higher than those charged in connection with other investments, and in some market conditions, may offset trading profits. Accordingly, shares or interests are suitable only for sophisticated investors for whom an investment in the fund does not constitute a complete investment program and who fully understand and are willing to assume the risks involved in the fund's investment program.

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Endnotes

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Where stated as "Net" or "TR" (or total return), the index return numbers include reinvestment of dividends net of withholding taxes, as calculated by the index provider.

