

MADRID KNOWLEDGE EXCHANGE 2023

September 2023

HARVESTING INNOVATION: INVESTMENT OPPORTUNITIES IN HEALTH CARE

Global Life Sciences Fund

Andy Acker, CFA
Portfolio Manager

SFDR CATEGORISATION

Classified as Article 8

Note: In accordance with the Sustainable Finance Disclosure Regulation, the Fund is classified as Article 8 and promotes, among other characteristics, environmental and/or social characteristics, and invests in companies with good governance practices.

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ESG-related disclosures

Our firm-wide approach in summary

Being a global asset management organisation comes with important responsibilities. As an active manager, this means that integrating Environmental, Social and Governance (ESG) factors into our investment decision-making and ownership practices is fundamental to delivering the results clients seek.

- At a corporate level, ESG principles influence our people, our culture and our choices, helping to make us a better company.
- At an investment level, we have integrated ESG factors into our analysis and processes as appropriate, helping us identify opportunities and risks, and influence positive change as we engage with companies.
- Recognising that there is a lack of consistency in ESG implementation and articulation across the industry, we seek to be clear in our communication as well as providing insight and education for our clients.

Visit janushenderson.com for more information.

This product

For all investors:

Environmental, Social and Governance (ESG) or sustainable investing considers factors beyond traditional financial analysis. This may limit available investments and cause performance and exposures to differ from, and potentially be more concentrated in certain areas than, the broader market.

Pour les investisseurs en France:

L'attention des investisseurs est attirée sur le fait que ce fonds présente, au regard des attentes de l'Autorité des marchés financiers, une communication disproportionnée sur la prise en compte des critères extra-financiers dans sa gestion.

D10580

About the speaker



Andy Acker, CFA
Portfolio Manager

Firm since 1999 | Industry since 1996

- Responsible for managing the Global Life Sciences and Biotechnology strategies since 2007 and 2018, respectively
- Leads the firm's Health Care Sector Research Team
- Bachelor's degree in biochemical sciences from Harvard University, graduating *magna cum laude* and Phi Beta Kappa
- MBA with honours from Harvard Business School

Health care suite

\$12bn in global assets under management across all strategies and geographies

Supported by a team of specialists with more than 100 years of combined health care investing experience

Three distinct strategy offerings

Global Life
Sciences

Long-only
Biotechnology

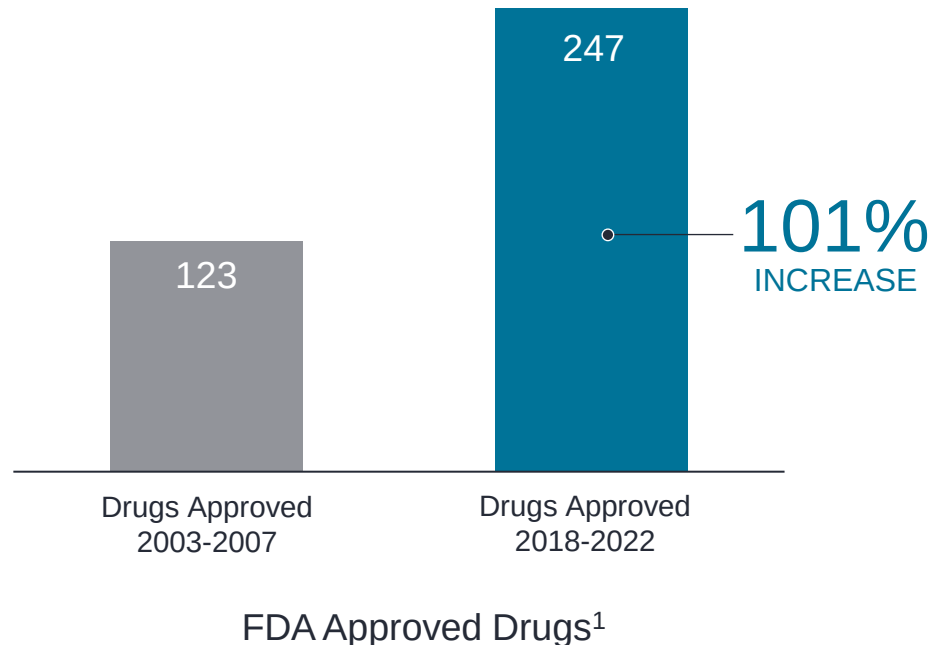
Long-short
Biotechnology

Source: Janus Henderson Investors, as at 30 June 2023.

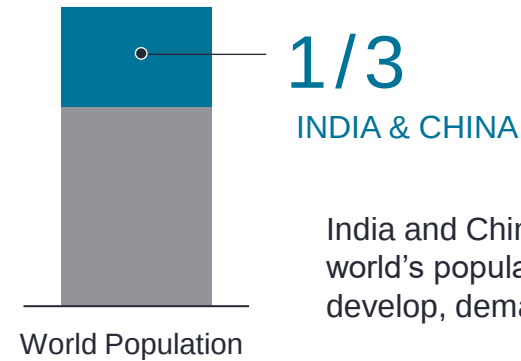
Why health care?

A powerful combination of factors are driving long-term sustainable growth

> ACCELERATING INNOVATION IN MEDICINE

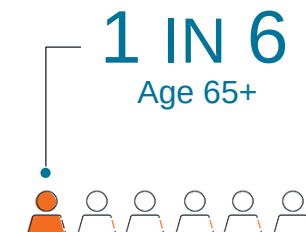


> RISING MIDDLE CLASS



India and China represent a third of the world's population. As their middle classes develop, demand for health care will increase.²

> AGING POPULATIONS



On average, those over the age of 65 spend 3x more on health care.

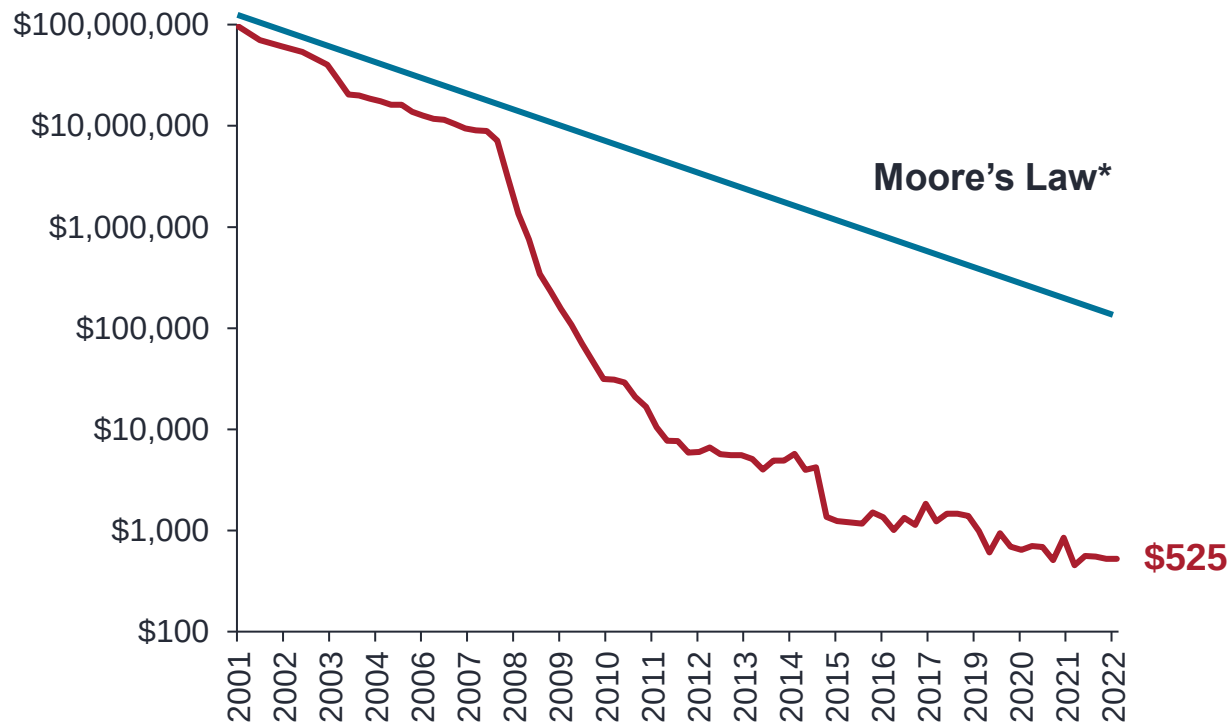
By 2050, one in six people in the world will be over age 65 (16%), leading to increased health care spending.³

Source: ¹FDA, as at 31 December 2022. ²World Population, as at 31 December 2022. ³United Nations, un.org as at 31 December 2019.

Accelerating innovation

Dramatic improvements in scientific productivity

Cost per genome



Gene sequencing advancing medicine

- The Human Genome Project was initially completed in 2000 after 13 years and \$3bn.
- Rapid innovation has driven cost below \$1000 per genome.
- Applications include targeted cancer therapy, prenatal screening, and treatments for rare genetic diseases.

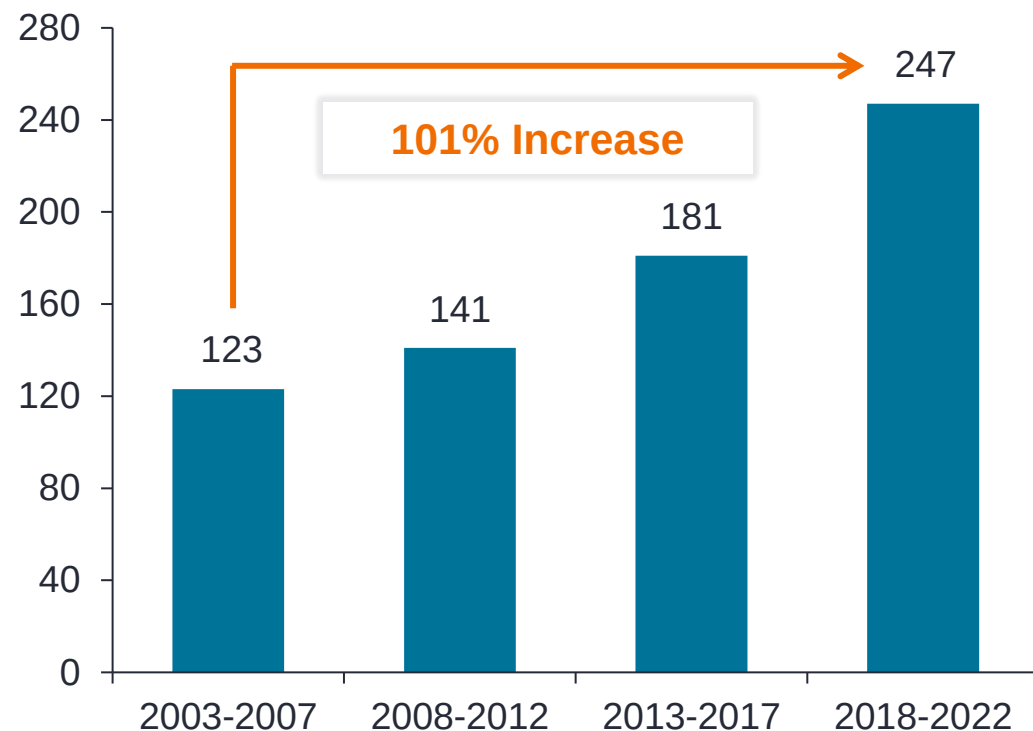
Source: National Human Genome Research Institute, as at May 2022.

Note: *Moore's law refers to a prediction by Intel co-founder Gordon Moore that the number of transistors on integrated circuits would double every two years. Moore's law has been credited with driving advances in the semiconductor industry that helped make computing widespread and affordable.

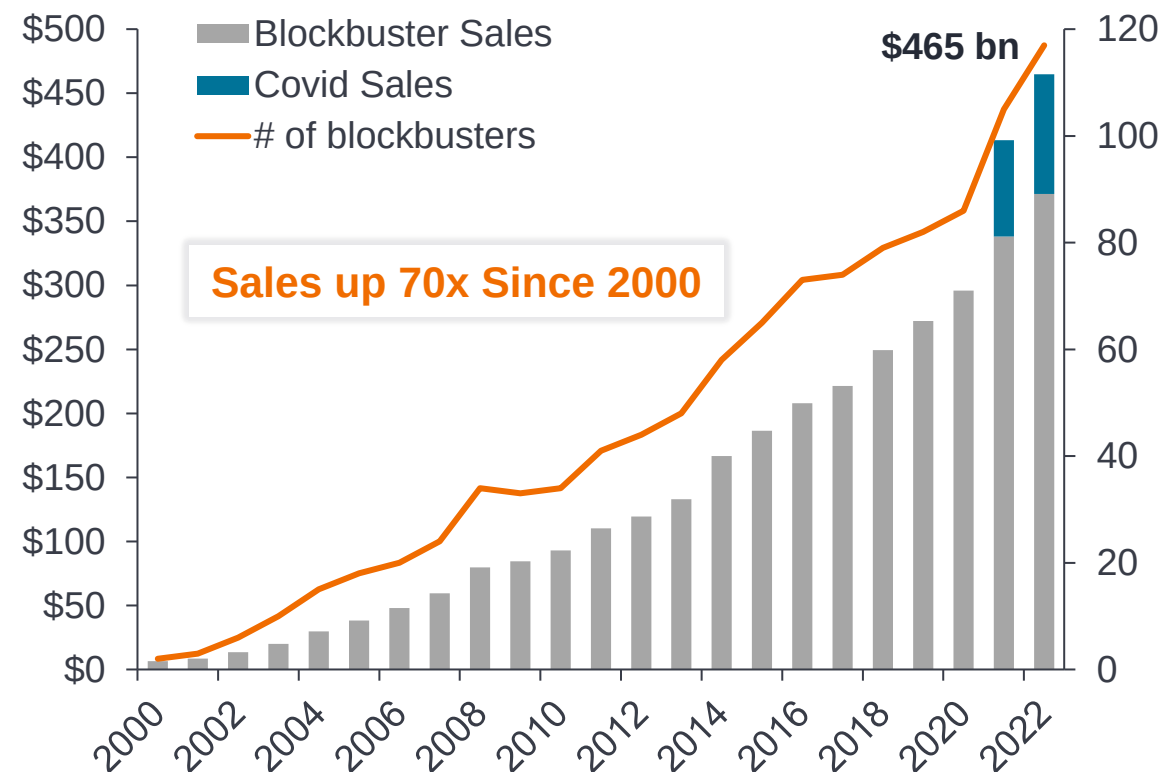
Accelerating innovation

Dramatic improvements in scientific productivity

Number of drugs receiving FDA approval



Blockbuster drug sales (\$Billions)



Source: Janus Henderson Investors, ISI Research, as at 31 December 2022.

Note: Blockbuster drug = > \$1 billion in annual sales. A blockbuster drug is a very popular drug that generates annual sales of \$1 billion or more for the company that sells it.

Source: FDA, as at 31 December 2022.

Opportunities for innovation

Accelerating innovation addressing unmet medical needs



GENETIC DISEASE

New advances in gene therapy offer the promise of treating 7,000+ genetic diseases, 95% of which currently have no available treatment.



CANCER

Moving from indiscriminate chemotherapies and radiation to more targeted immuno-oncology treatments that leverage, rather than attack, the body's immune system.



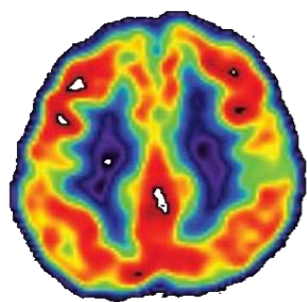
Source: Pharmaceutical Research and Manufacturers of America. American Cancer Society, 2018.

Opportunities for innovation

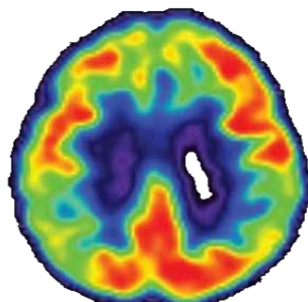
Accelerating innovation addressing unmet medical needs

ALZHEIMER'S

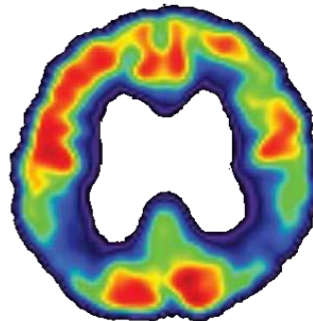
55 million people worldwide are living with Alzheimer's or dementia. The disease kills more people than breast cancer and prostate cancer combined. Death due to Alzheimer's has increased 145% from 2000 to 2019.



Normal



Mild Cognitive Impairment



Alzheimer's Disease

OBESITY

More than 40% of the global population is expected to be obese or overweight by 2030.¹ The cost of treating obesity-related health issues adds up to trillions of dollars (USD) annually.²



Source: ¹"Overcoming obesity: An initial economic analysis," Dobbs, Richard; Sawers, Corinne; Thompson, Fraser; et al., McKinsey Global Institute, November 2014.

²"The Heavy Burden of Obesity: The Economics of Prevention," OECD Health Policy Studies, 2019.

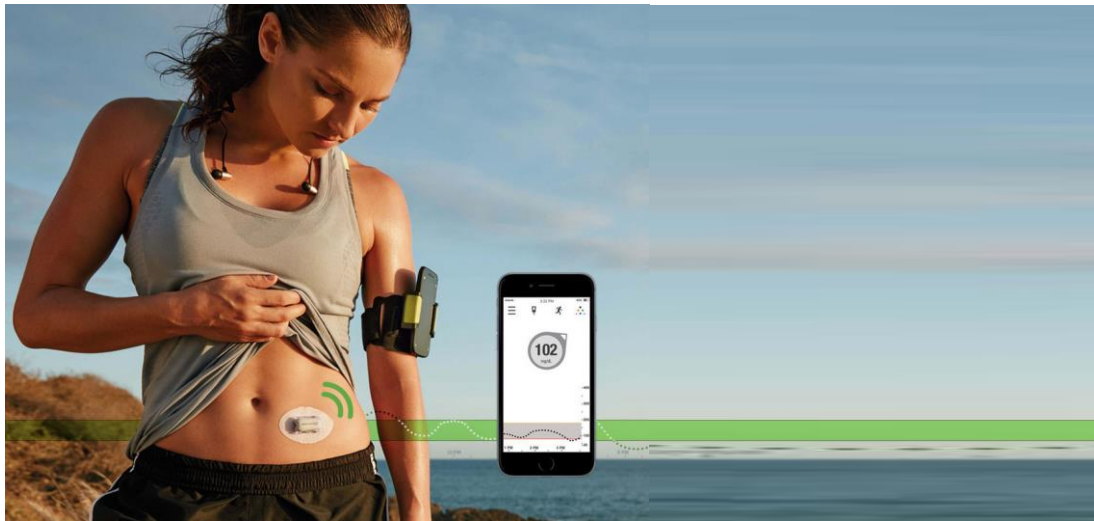
Source: <https://www.alz.org>.

Opportunities for innovation

Accelerating innovation addressing unmet medical needs

DIABETES

New medical devices harmonise the use of continuous glucose monitoring and insulin pumps to provide revolutionary ways for diabetics to manage their disease.



ROBOTIC SURGERY

Robotic surgery can improve accuracy beyond the human hand and offers patients the promise of better outcomes with less pain and faster recoveries.



Source: World Health Organization – Diabetes, as of June 2016.

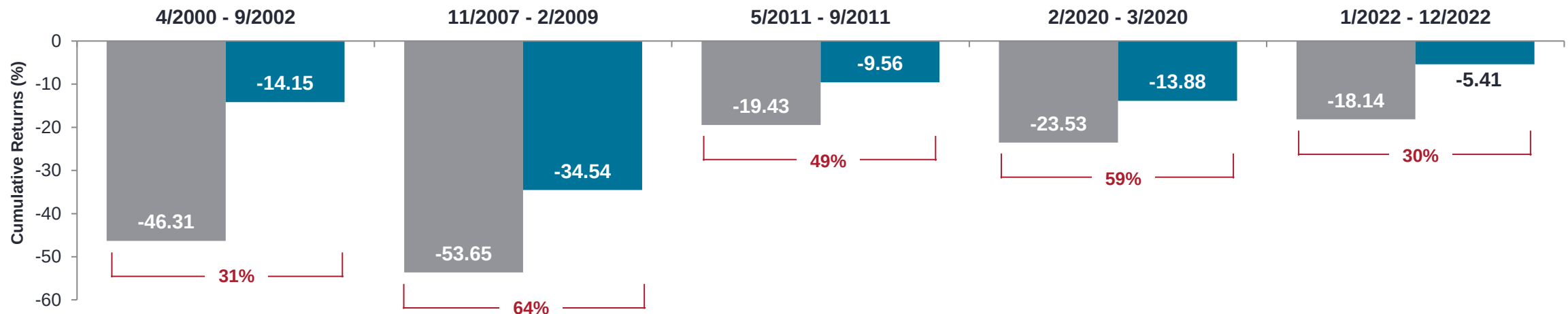
Defensive characteristics

Health care has typically held up better in down markets with lower volatility

Performance and downside capture

MSCI World Health Care IndexSM averaged 47% downside capture of MSCI World IndexSM returns

■ MSCI World IndexSM ■ MSCI World Health Care IndexSM ■ Downside Capture



Beta 10 Year Period (30 June 2013 – 30 June 2023)

MSCI World Health Care IndexSM vs. MSCI World IndexSM 0.74

Standard Deviation 10 Year Period (30 June 2013 – 30 June 2023)

MSCI World Health Care Index SM	13.32
MSCI World Index SM	14.62

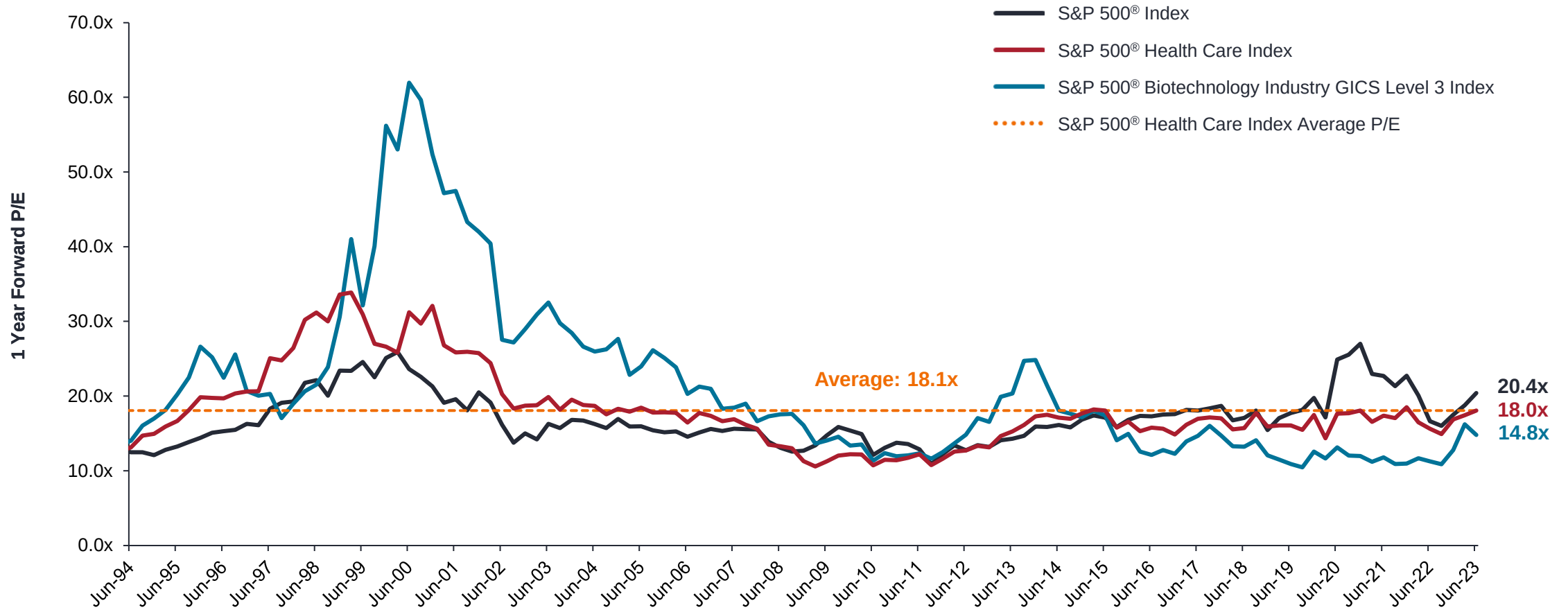
Source: Janus Henderson Investors, FactSet, as at 30 June 2023.

Note: Chart reflects market declines of 15% or greater in the MSCI World IndexSM since 2000.

Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

Historical valuations

Historical valuations of healthcare stocks (1-year forward P/E)
(30 June 1994 – 30 June 2023)



Source: Bloomberg, as at 30 June 2023.

Attractive entry point into biotech

Putting the recent biotech drawdown in historical context

Drawdown Analysis (%)					After-Drawdown Absolute Performance (%)					After-Drawdown Relative Performance vs. S&P 500 (%)				
Drawdown Period	# of Trading Days	Biotech	Biotech vs S&P 500	Biotech vs Health Care Sector	1 Wk	1 Mo	3 Mo	6 Mo	1 Yr	1 Wk	1 Mo	3 Mo	6 Mo	1 Yr
27 Feb 06 – 23 May 06	61	-21.8	-19.2	-13.3	4.4	1.2	5.2	21.0	27.2	4.2	2.3	1.4	8.3	3.8
10 Dec 07 – 10 Mar 08	62	-22.6	-7.0	-8.3	0.5	16.4	21.8	29.9	-4.4	0.2	11.9	14.4	29.3	37.7
14 Aug 08 – 5 Mar 09	138	-37.6	8.7	-3.9	8.9	9.4	12.7	23.2	36.8	-1.3	-13.0	-26.3	-25.5	-33.7
23 Mar 10 – 6 Jul 10	73	-19.5	-7.6	-6.5	7.5	13.5	21.2	27.0	49.2	0.9	4.2	7.7	2.2	16.2
22 Jul 11 – 8 Aug 11	12	-26.7	-10.0	-11.5	12.6	9.3	14.9	48.7	54.7	4.9	4.2	1.6	27.3	26.6
27 Feb 14 – 8 May 14	50	-29.1	-30.6	-26.2	4.0	23.2	21.3	41.0	87.4	4.1	20.0	19.0	31.6	72.3
17 Jul 15 – 8 Feb 16	140	-49.2	-37.4	-32.4	6.9	17.4	14.1	37.8	44.2	4.5	9.2	2.4	18.8	17.7
22 Sep 16 – 3 Nov 16	30	-22.6	-18.7	-12.6	23.4	14.7	22.2	32.7	61.0	19.6	9.5	12.4	17.2	34.6
20 Jun 18 – 24 Dec 18	129	-35.3	-21.1	-30.9	9.7	22.3	34.5	29.5	50.5	3.0	9.7	14.8	3.0	10.6
5 Apr 19 – 1 Oct 19	124	-20.4	-23.1	-18.3	0.5	12.1	26.8	3.7	51.3	2.1	7.6	15.5	15.0	34.1
19 Feb 20 – 16 Mar 20	19	-32.9	-3.5	-11.5	4.0	26.3	56.3	69.3	121.7	10.2	10.4	27.1	26.2	52.7
Current Drawdown (8 Feb 21 – 11 May 22)	316	-63.9	-66.2	-74.8	8.1	0.8	44.5	35.0	36.9	8.3	5.3	37.1	32.6	30.2
Average	96	-31.8	-19.6	-20.8	7.5	13.9	24.6	33.2	51.4	5.1	6.8	10.6	15.5	25.2
Median	68	-27.9	-19.0	-12.9	7.2	14.1	21.6	31.3	49.8	4.1	8.4	13.4	18.0	28.4

Source: Bloomberg, Janus Henderson Investors analysis, as at 11 May 2023.

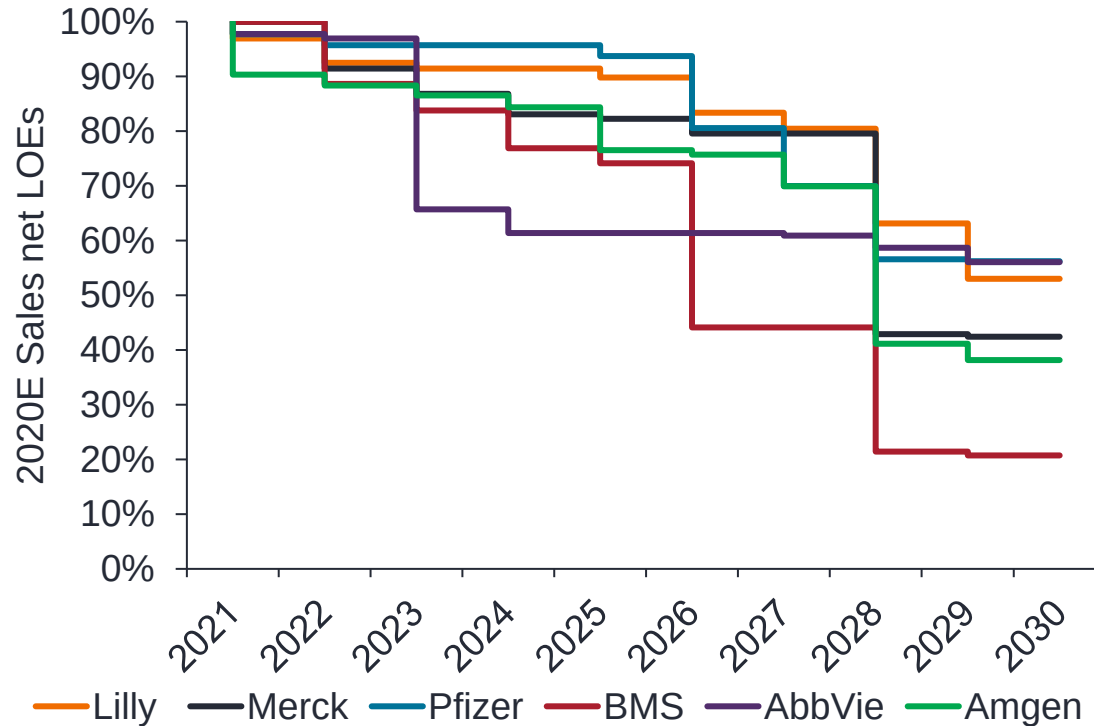
Note: Biotech performance is for SPDR® S&P® Biotech ETF (XBI). XBI is designed to correspond generally to the performance of the S&P Biotechnology Select Industry Index, which is equal weighted. XBI inception date was 31 January 2006. Short-term performance is not typical and may not be achieved in the future. Such results should not be the sole basis for evaluating material facts in making an investment decision. Index performance does not reflect the expenses of managing a portfolio as an index is unmanaged and not available for direct investment. References to exchange traded funds and indices are for informational purposes only and may include significant differences compared to the fund, including but not limited to; objectives, sales and management fees, liquidity, volatility, tax and other features, which may result in differences in performance. The fund may not invest in a significant portion of the securities represented by the benchmarks.

Past performance does not predict future returns.

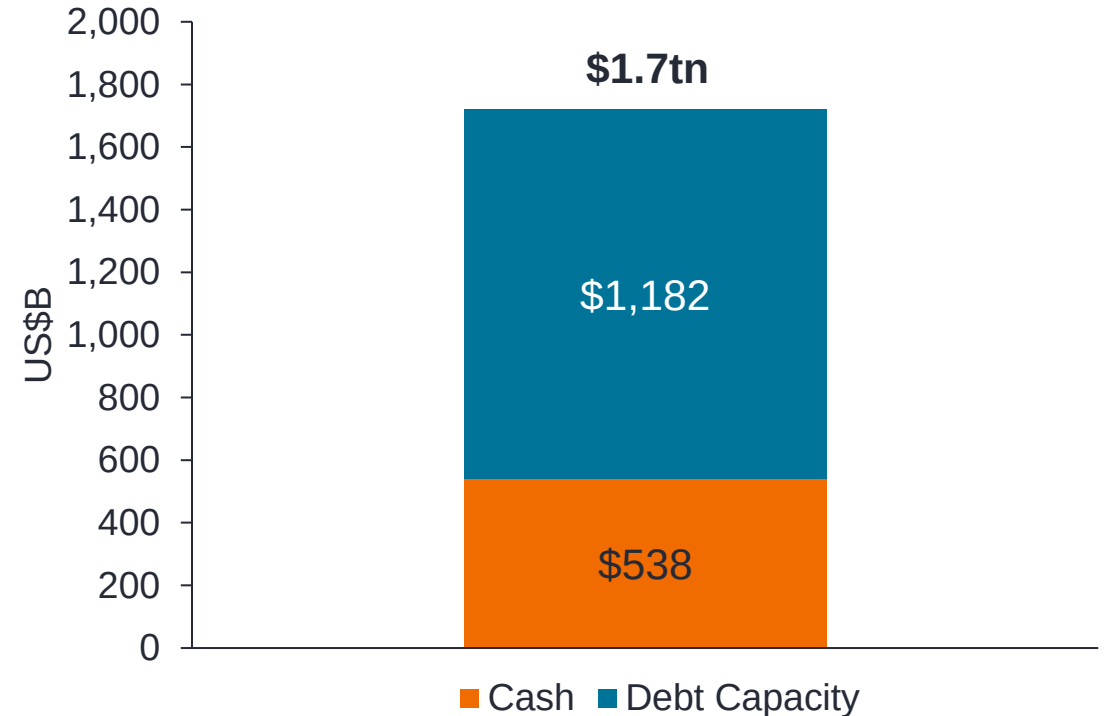
M&A potential is significant

Stage may be set for an uptick in activity

US biopharma faces significant loss of exclusivity beyond 2025...



...And maintains substantial deal-making capacity



Source: Bernstein Analysis. "Pharma is buying, but what's on the menu? Top takeout picks based on our M&A screen." 28 January 2022

Note: There is no guarantee that past trends will continue, or forecasts will be realized. References made to individual securities do not constitute a recommendation to buy, sell or hold any security, investment strategy or market sector, and should not be assumed to be profitable. Janus Henderson Investors, its affiliated advisor, or its employees, may have a position in the securities mentioned.

Source: SVB Leerink, as at 01 December 2021.

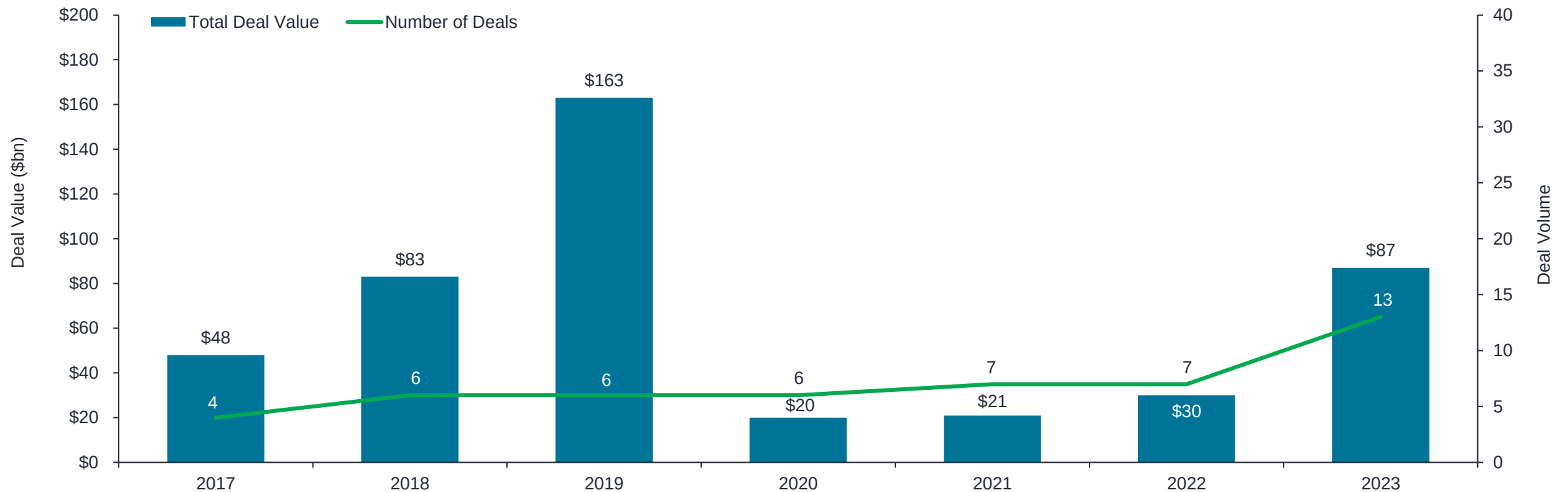
Note: Data aggregated for a cohort of 18 large-cap pharmaceutical companies.

M&A potential is significant

YTD M&A volumes and deal values have already doubled 2022

Major M&A deals in transaction value

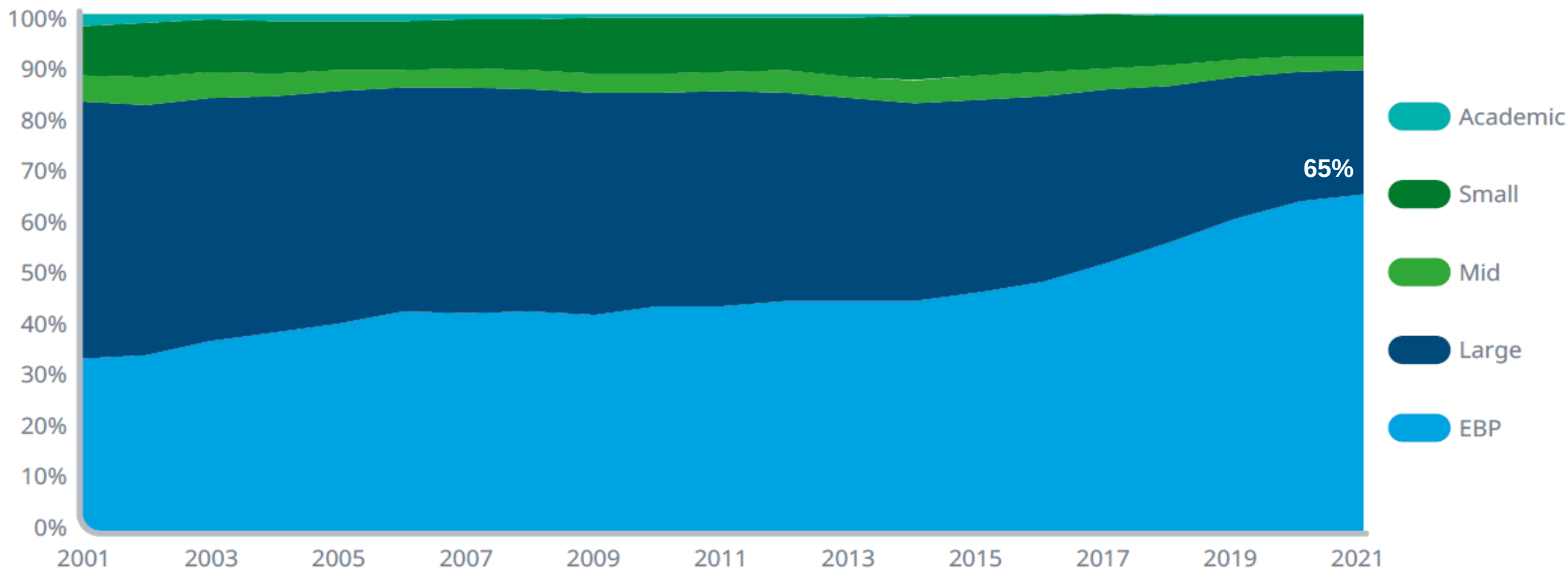
(01 January 2023 – 04 September 2023)



Source: Cowen, as of September 2023

Harvesting innovation in biotech

Emerging BioPharma (EBP) is responsible for 65% of new molecules in the R&D pipeline



Source: IQVIA Pipeline Intelligence, as at December 2021 and IQVIA Institute as at January 2022

Note: Emerging biopharma companies (EBPs) are defined as those with less than \$500 million in annual sales and R&D spending less than \$200 million per year.

Headwinds for healthcare

Political rhetoric and regulatory shifts often leading causes of volatility in the sector

The FDA's Dysfunction Is A Public Health Crisis Of Its Own

Pfizer Expects Drop in Revenue as Covid Vaccine Demand Wanes

Drugmaker says in earnings report it anticipates return in growth for Covid-products in 2024

Big Pharma dealmaking recovers with \$85bn M&A splurge

Drugmakers' appetite for takeovers surges but FTC action against Amgen raises risks for megadeals

HEALTH

Biotech Stocks, Once Booming, Enter Bear Territory

The biotech stock index, which rose early in the pandemic on hopes for Covid-19 treatments, has fallen amid research failures, slowing M&A and rising interest rates

Democrats Choosing Less Risky Path on Drug Prices

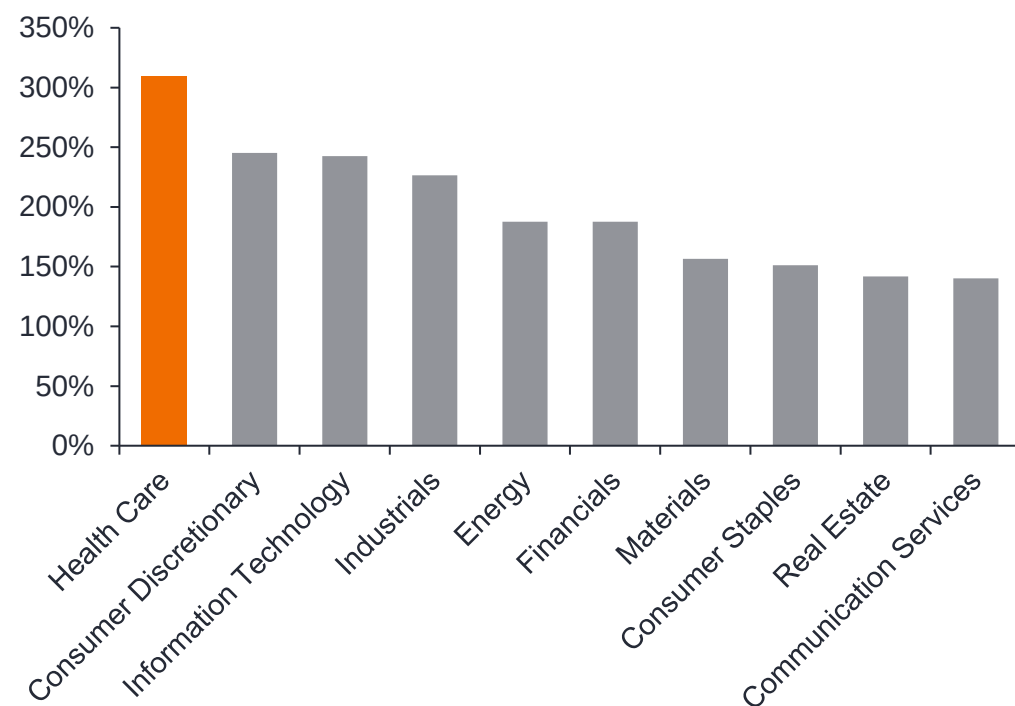
Experts say recent compromises could create less harm in balancing innovation with profits.

Source: The New York Times, The Wall Street Journal, Financial Times and Forbes.

The case for active management

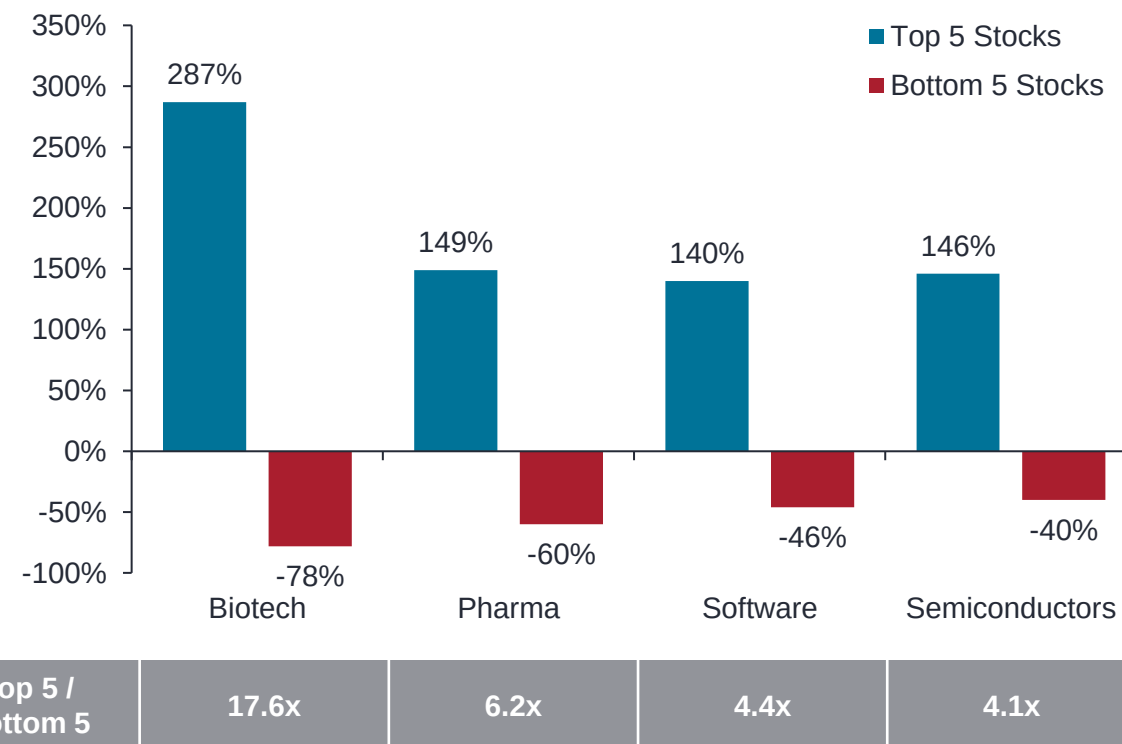
Health care has the biggest disparity between winners and losers

Disparity of returns by sector (over 10 years)



Source: Wilshire 5000 Index as at 31 December 2022.
Note: Includes average performance of stocks over \$500 mm in market cap
Past performance does not predict future returns.

10-year average return

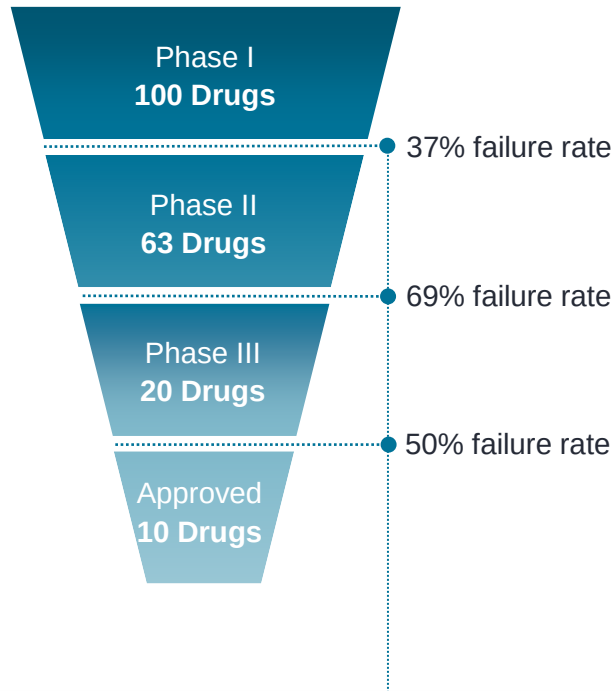


Source: Wilshire 5000 Index, 2013-2022.
Note: Based on analysis of ten-year period.
Past performance does not predict future returns.

The case for active management

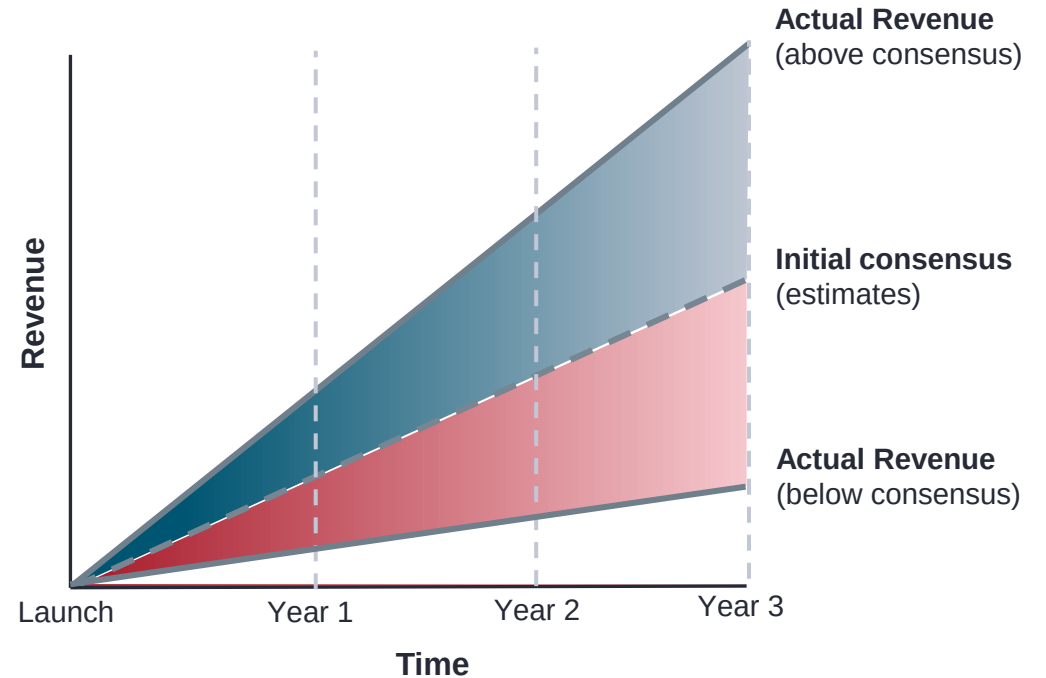
The 90/90 rule

Clinical Risk



90% of drugs that enter human clinical trials **NEVER** make it to the market.

Commercial Risk



90% of the time consensus estimates for new drug launches are **WRONG**, in our experience.

Source: BIO Industry Analysis, 2016.

Note: The 90/90 Rule is a concept created by Janus Henderson Investors. The above graphics are used for illustrative purposes only.

Investing with an experienced team

Global Life Sciences

- Specialised team with more than 100 years of combined health care investment experience
- Dedicated health care experts with three PhDs and one MD

PORTFOLIO MANAGEMENT



Andy Acker, CFA

Portfolio Manager

Financial Industry Experience: 27 years



Daniel Lyons, PhD, CFA

Portfolio Manager, Research Analyst

Financial Industry Experience: 23 years

- Joined Janus in 1999 as a research analyst focused on biotechnology and pharmaceuticals
- B.S. degree in biochemical sciences from Harvard University, graduating *magna cum laude* and Phi Beta Kappa
- MBA with honors from Harvard Business School

- Joined Janus Henderson in 2000
- PhD in immunology from Stanford and postdoctoral research with a Nobel laureate
- Bachelor's degree in biochemistry and chemistry from Rice University, graduating *magna cum laude*

BROADER HEALTH CARE TEAM

Agustin Mohedas, PhD

Biotechnology | 9 years of experience

Vish Sridharan, MD

Biotechnology | 3 years of experience

Luyi Guo, PhD, CFA

Pharmaceuticals | 10 years of experience

Tim McCarty, CFA

Medical Technology | 11 years of experience

Adam Poussard, CFA

Medical Technology | 17 years of experience

Lauren Petite

Medical Technology | 5 years of experience

Vicki Cohen

Associate Analyst | 3 years of experience

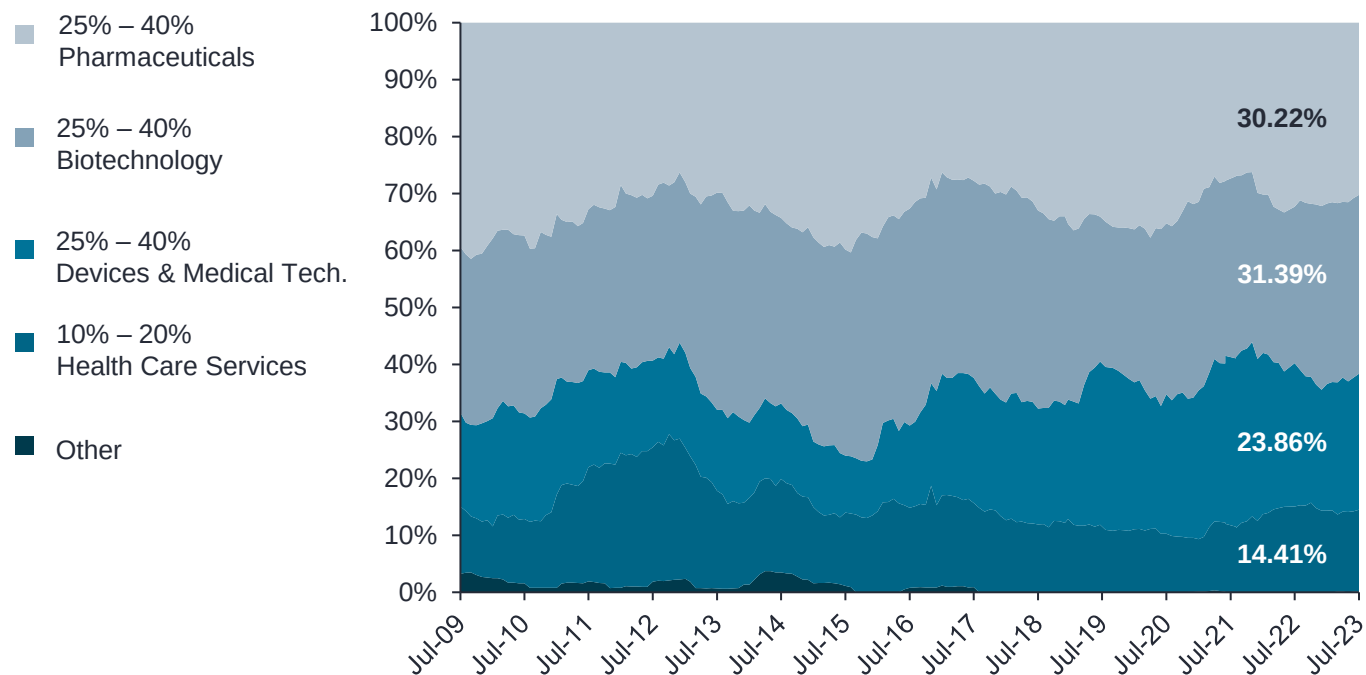
Source: Janus Henderson Investors, as at 31 July 2023.

Note: Years of experience refer to financial industry experience.

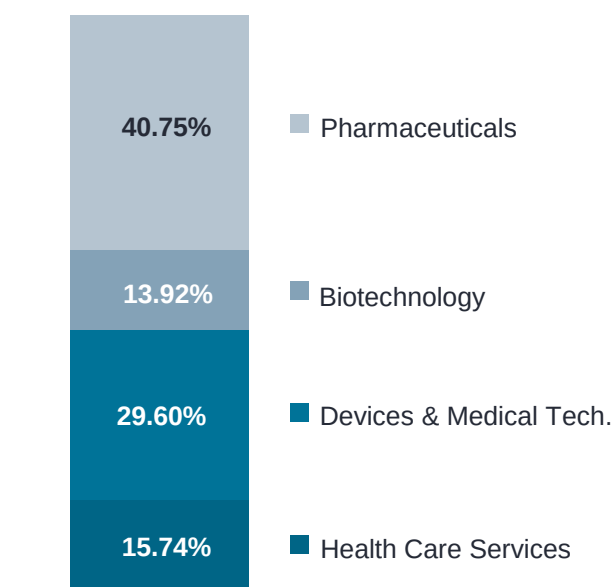
Portfolio composition

Diversified across the sub sectors within health care

Janus Henderson Global Life Sciences Fund



MSCI World Health Care IndexSM



Source: Janus Henderson Investors, as at 31 July 2023.

Note: Fund: Janus Henderson Global Life Sciences, in USD. Benchmark: MSCI World Health Care IndexSM

Allocations are subject to change without notice.

Performance

Overall Morningstar Rating™

Based on risk-adjusted returns as at 31 July 2023



12 USD Shares among 511 EAA Sector Equity Healthcare Funds

Note: © 2023 Morningstar, Inc. All Rights Reserved. A fee was paid for the use of this data. See appendix for important information.

Global Life Sciences Fund

Characteristics

Inception date:	31 March 2000	Benchmark	MSCI World Health Care Index SM
Managed since:	30 April 2007	Peer group	Sector Equity Health Care
Holdings range:	80 – 120 (public)	Min/max position size	Max 5.0% or 1.25x index weight (whichever is greater)
Non-US exposure	20% – 30%	Assets under management	Fund: \$3.8bn
Market cap range	All Cap		Strategy: \$11.4bn (as at 30 June 2023)

Investment Objective

The Fund aims to provide capital growth over the long term.

Performance target: To outperform the MSCI World Health Care Index by at least 2% per annum, before the deduction of charges, over any 5 year period.

The Fund invests at least 80% in shares (also known as equities) of companies, of any size, with a life sciences orientation, in any country.

The Fund may also invest in other assets including cash and money market instruments.

The investment manager may use derivatives (complex financial instruments) to reduce risk, to manage the Fund more efficiently, or to generate additional capital or income for the Fund.

The Fund is actively managed with reference to the MSCI World Health Care Index, which is broadly representative of the companies in which it may invest, as this forms the basis of the Fund's performance target. The investment manager has discretion to choose individual investments for the Fund with weightings different to the index or not in the index, but at times the Fund may hold investments similar to the index.

The investment manager looks to identify innovative healthcare companies addressing high unmet medical needs across fields including biotechnology, pharmaceuticals, healthcare services and medical technology. The team understands that success of drug development is binary in nature, creating wide disparities between winners and losers. The investment process leverages proprietary statistical models to analyse the probability of a company's success, focusing on products they believe can overcome the rigours of clinical development. Additional tools, such as physician surveys and prescription models, attempt to more accurately predict commercial viability.

Source: Janus Henderson Investors, as at 31 July 2023, unless otherwise noted.

Fund specific risks

Global Life Sciences Fund

Currency Hedging	When the Fund, or a hedged share/unit class, seeks to mitigate exchange rate movements of a currency relative to the base currency, the hedging strategy itself may create a positive or negative impact to the value of the Fund due to differences in short-term interest rates between the currencies.
Counterparty and Operational Risk	The Fund could lose money if a counterparty with which the Fund trades becomes unwilling or unable to meet its obligations, or as a result of failure or delay in operational processes or the failure of a third party provider.
Equities	Shares/Units can lose value rapidly, and typically involve higher risks than bonds or money market instruments. The value of your investment may fall as a result.
Smaller Companies	Shares of small and mid-size companies can be more volatile than shares of larger companies, and at times it may be difficult to value or to sell shares at desired times and prices, increasing the risk of losses.
Country or Region	If a Fund has a high exposure to a particular country or geographical region it carries a higher level of risk than a Fund which is more broadly diversified.
Investment Focus	The Fund is focused towards particular industries or investment themes and may be heavily impacted by factors such as changes in government regulation, increased price competition, technological advancements and other adverse events.
Derivatives and Leverage	The Fund may use derivatives towards the aim of achieving its investment objective. This can result in 'leverage', which can magnify an investment outcome and gains or losses to the Fund may be greater than the cost of the derivative. Derivatives also introduce other risks, in particular, that a derivative counterparty may not meet its contractual obligations.

(continued on next page)

Source: Janus Henderson Investors, as at 31 July 2023.

Fund specific risks

Global Life Sciences Fund (continued)

Exchange Rates	If the Fund holds assets in currencies other than the base currency of the Fund or you invest in a share/unit class of a different currency to the Fund (unless 'hedged'), the value of your investment may be impacted by changes in exchange rates.
Liquidity	Securities within the Fund could become hard to value or to sell at a desired time and price, especially in extreme market conditions when asset prices may be falling, increasing the risk of investment losses.

Note that any differences among portfolio securities currencies, share class currencies and costs to be paid or represented in currencies other than your home currency will expose you to currency risk. Costs and returns may increase or decrease as a result of currency and exchange rate fluctuations.

Fund charges will impact the value of your investment. In particular, the ongoing charges applicable to each fund will dilute investment performance, particularly over time. For further explanation of charges please visit our Fund Charges page at www.janushenderson.com.

The ongoing charge is calculated using the PRIIP methodology. The PRIIP methodology differs to the UCITS ongoing charge methodology, as the PRIIP methodology captures additional recurring charges, including but not limited to: Interest paid on borrowing (e.g. bank interest); Any fees incurred in relation to stock-lending activity (i.e. the fee paid to the lending agent); Any costs associated with holding closed-ended vehicles.

For EU investors

Lower Risk

1	2	3	4	5	6	7
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 Higher Risk

For Dutch investors only



Source: Janus Henderson Investors, as at 30 June 2023. Risk indicator refers to I2 USD share class. Rating is latest available as at 01 January 2023.

Note: UK investors should refer to the Key Investor Information Document as the risk rating may differ.

With effect from 1 January 2023, the Key Investor Information document (KIID) changed to the Key Information Document (KID), except in the UK where investors should continue to refer to the KIID. Availability of share classes shown may be limited by law in certain jurisdictions. Performance records/scenarios are detailed within the Fund's specific KIID/KID; fees and charges, and the respective risk rating may vary. Further information can be found in the Fund's prospectus and KIID/KID, which must be reviewed before investing. Please consult your local sales representative if you have any queries.

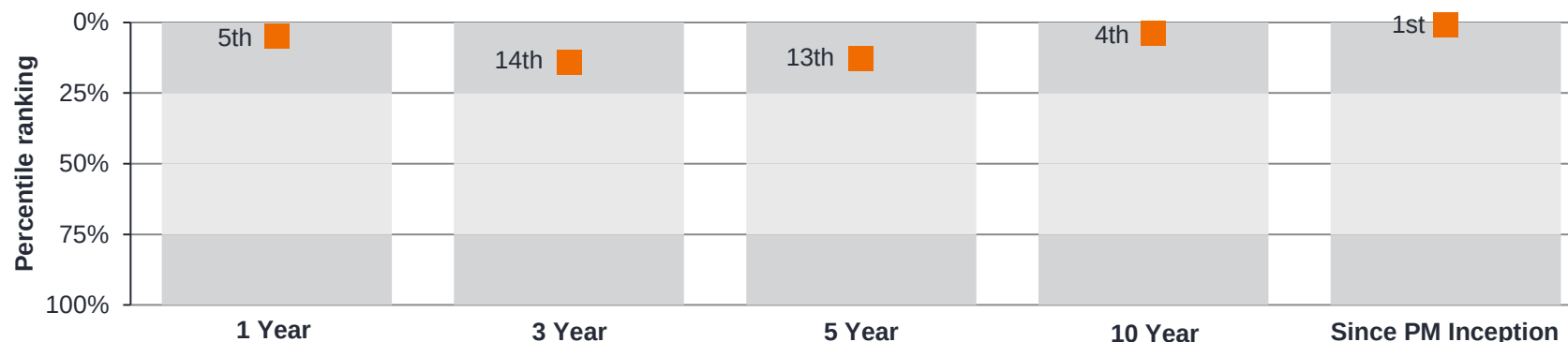
Performance

Strong long-term performance versus health care sector

Performance – USD (%)	YTD	1 year	3 year	5 year	10 year	Since PM inception (30/04/07)
Global Life Sciences Fund (Class I2 USD Shares)	3.95	11.68	7.35	8.91	11.61	10.99
MSCI World Health Care Index SM	2.08	4.27	7.55	9.02	9.87	8.58
Difference (Fund vs Index)	+1.87	+7.41	-0.20	-0.11	+1.74	+2.41

Performance Rankings

■ Janus Henderson Global Life Sciences Fund (I2 USD)



Source: Janus Henderson Investors, Morningstar, as at 31 July 2023. © 2023 Morningstar, Inc. All Rights Reserved.

Note: Fund: Janus Henderson Global Life Sciences Fund, I2 Shares, net of fees in USD. Benchmark: MSCI World Health Care IndexSM. Returns include reinvestment of dividends and capital gains. Returns greater than one year are annualised.

Morningstar ranking are shown using the Sector Equity Healthcare Category, net of fees in USD.

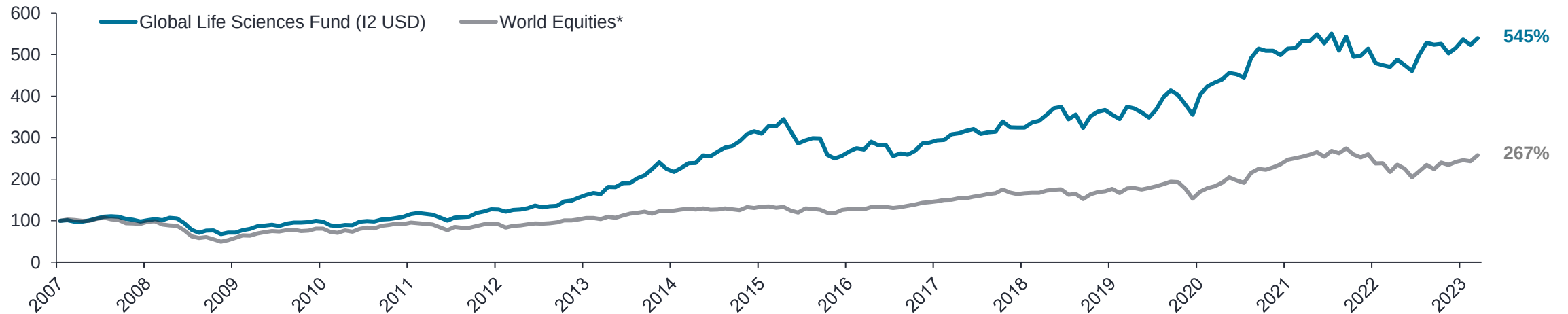
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Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

Performance

Strong long-term performance vs the broad world equities market

Performance – USD (%)	1 year	3 year	5 year	10 year	Since PM inception (30/04/07)
Global Life Sciences Fund (Class I2 USD Shares)	11.68	7.35	8.91	11.61	10.99
World Equities*	13.48	11.67	9.12	9.30	6.23
Difference (Fund vs Index)	-1.80	-4.32	-0.21	+2.31	+4.76



***World Equities is represented by the MSCI World IndexSM and is provided for informational purposes only. It is not a performance comparator/target or official benchmark for the fund, nor is it applicable to the investment process. For performance comparison against the fund's official benchmark please see slide 27.**

Source: Janus Henderson Investors, Morningstar, as at 31 July 2023. © 2023 Morningstar, Inc. All Rights Reserved.

Note: Fund: Janus Henderson Global Life Sciences Fund, I2 Shares, net of fees in USD

Returns include reinvestment of dividends and capital gains. Returns greater than one year are annualised.

Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

Calendar year returns

Performance (%)	YTD at 2Q23	2022	2021	2020	2019	2018	2017	2016	2015
Global Life Sciences Fund (Class I2 USD Shares)	2.96	-3.59	5.60	24.37	28.04	2.76	21.48	-13.17	6.55
MSCI World Health Care Index SM	0.78	-5.41	19.80	13.52	23.24	2.51	19.80	-6.81	6.60
Peer Group Average	1.64	-13.91	7.12	20.53	21.80	-3.40	21.68	-9.37	5.90
Quartile	2nd	1st	3rd	2nd	1st	1st	2nd	4th	2nd
Percentile	38th	9th	66th	30th	17th	21st	29th	81st	36th

Performance (%)	2014	2013	2012	2011	2010	2009	2008	2007*
Global Life Sciences Fund (Class I2 USD Shares)	33.68	53.88	23.73	7.03	7.16	25.68	-30.45	9.66
MSCI World Health Care Index SM	18.10	36.27	17.54	9.46	2.41	18.89	-21.50	-2.45
Peer Group Average	18.33	35.09	17.95	0.43	4.77	24.38	-24.71	-1.76
Quartile	1st	1st	1st	2nd	2nd	2nd	4th	1st
Percentile	2nd	5th	12th	28th	30th	30th	82nd	5th

Source: Morningstar, Janus Henderson Investors analysis, as at 30 June 2023.

Note: *Partial year performance from 01 May 2007, as at PM inception.

Fund: Janus Henderson Global Life Sciences Fund, I2 share class, net of fees in USD. Benchmark: MSCI World Health Care IndexSM

Peer Group Category: Morningstar USD Sector Equity Healthcare. Peer group average performance is based on valuation point of underlying funds in universe.

Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

Fund performance disclosure

Annualised performance (%)	I2 USD (Net)	Benchmark	I2 USD (Gross)	Target (Gross)*
5 Years	9.62	10.04	11.38	12.24
10 Years	12.62	10.33	14.48	12.54

Source: Janus Henderson Investors, as at 30 June 2023.

Note: Fund: Janus Henderson Global Life Sciences Fund, I2 share class, gross and net of ongoing charges, in USD.

Benchmark: MSCI World Health Care IndexSM

***Performance target: To outperform the MSCI World Health Care IndexSM by at least 2% per annum, before the deduction of charges, over any 5 year period.**

Performance/performance target related data will display only when relevant to the fund/share class inception date and/or the annualised target time period.

Please note the performance target is to be achieved over a specific annualised time period. Refer to the performance target wording within the objective.

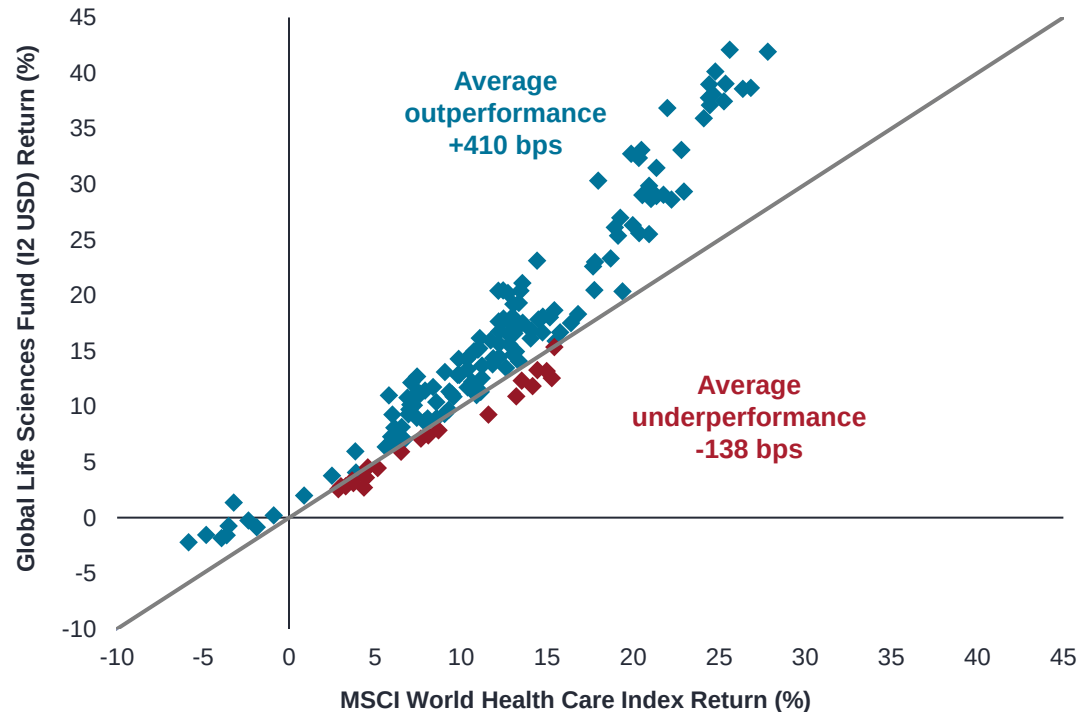
Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

Consistent outperformance

Global Life Sciences Fund

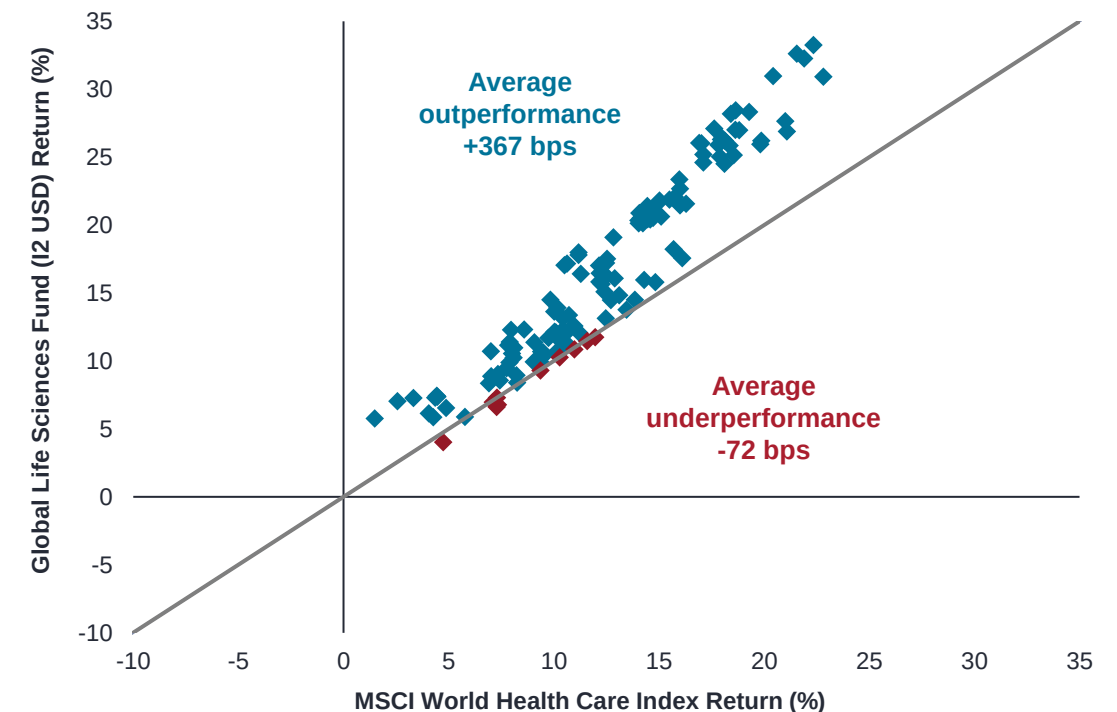
Rolling three-year excess returns

Global Life Sciences Fund (I2 USD) **outperformed** the MSCI World Health Care Index 120 of 164 periods, or **73% of the time**



Rolling five-year excess returns

Global Life Sciences Fund (I2 USD) **outperformed** the MSCI World Health Care Index 114 of 140 periods, or **81% of the time**



Source: Janus Henderson Investors, as at 31 July 2023.

Note: Fund: Janus Henderson Global Life Sciences Fund (I2 USD), net of fees, in USD. Benchmark: MSCI World Health Care IndexSM.

Data shown is for the "Value at Risk" approach period from 01 January 2007 – 31 July 2023. See page 27 for complete performance and important disclosures.

Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

Horizon Biotechnology Fund

Characteristics

Inception date:	10 December 2018	Benchmark	NASDAQ Biotechnology TR Index SM
Holdings range:	50 – 80	Peer group	Sector Equity Biotechnology
Non-US exposure	10% – 20%	Min/max position size	Typically 0.5% to 8%
Market cap range	Typically > \$300 mm	Assets under management	\$161 mm

Investment Objective

The Fund aims to provide capital growth over the long term.

Performance target: To outperform the NASDAQ Biotechnology Total Return Index by 2% per annum, before the deduction of charges, over any 5 year period.

The Fund invests at least 80% of its assets in shares (equities) and equity-related securities of companies, which are biotechnology or biotechnology-related, in any country.

The Fund may also invest in other assets including bonds (including convertible bonds), preference shares, cash and money market instruments.

The investment manager may use derivatives (complex financial instruments) with the aim of making investment gains in line with the Fund's objective, to reduce risk or to manage the Fund more efficiently.

The Fund is actively managed with reference to the NASDAQ Biotechnology Total Return Index, which is broadly representative of the companies in which it may invest, as this forms the basis of the Fund's performance target and the level above which performance fees may be charged (if applicable). The investment manager has discretion to choose investments for the Fund with weightings different to the index or not in the index, but at times the Fund may hold investments similar to the index.

The investment manager looks to identify innovative biotechnology companies addressing high unmet medical needs and trading at a significant discount to their intrinsic value. The team understands that success of drug development is binary in nature, creating wide disparities between winners and losers. The investment process leverages proprietary statistical models to analyse the probability of a company's success, focusing on products they believe can overcome the rigours of clinical development. Additional tools such as physician surveys, prescription models and scenario simulations attempt to more accurately predict commercial viability.

Source: Janus Henderson Investors, as at 31 July 2023.

Fund specific risks

Horizon Biotechnology Fund

Currency Hedging	When the Fund, or a hedged share/unit class, seeks to mitigate exchange rate movements of a currency relative to the base currency, the hedging strategy itself may create a positive or negative impact to the value of the Fund due to differences in short-term interest rates between the currencies.
Counterparty and Operational Risk	The Fund could lose money if a counterparty with which the Fund trades becomes unwilling or unable to meet its obligations, or as a result of failure or delay in operational processes or the failure of a third party provider.
Equities	Shares/Units can lose value rapidly, and typically involve higher risks than bonds or money market instruments. The value of your investment may fall as a result.
Smaller Companies	Shares of small and mid-size companies can be more volatile than shares of larger companies, and at times it may be difficult to value or to sell shares at desired times and prices, increasing the risk of losses.
Country or Region	If a Fund has a high exposure to a particular country or geographical region it carries a higher level of risk than a Fund which is more broadly diversified.
Investment Focus	The Fund is focused towards particular industries or investment themes and may be heavily impacted by factors such as changes in government regulation, increased price competition, technological advancements and other adverse events.
Derivatives and Leverage	The Fund may use derivatives towards the aim of achieving its investment objective. This can result in 'leverage', which can magnify an investment outcome and gains or losses to the Fund may be greater than the cost of the derivative. Derivatives also introduce other risks, in particular, that a derivative counterparty may not meet its contractual obligations.

(continued on next page)

Source: Janus Henderson Investors, as at 31 December 2022.

Fund specific risks

Horizon Biotechnology Fund (continued)

Exchange Rates	If the Fund holds assets in currencies other than the base currency of the Fund or you invest in a share/unit class of a different currency to the Fund (unless 'hedged'), the value of your investment may be impacted by changes in exchange rates.
Liquidity	Securities within the Fund could become hard to value or to sell at a desired time and price, especially in extreme market conditions when asset prices may be falling, increasing the risk of investment losses.
Transaction Costs (higher charges)	The Fund may incur a higher level of transaction costs as a result of investing in less actively traded or less developed markets compared to a fund that invests in more active/developed markets. These transaction costs are in addition to the Fund's Ongoing Charges.

Note that any differences among portfolio securities currencies, share class currencies and costs to be paid or represented in currencies other than your home currency will expose you to currency risk. Costs and returns may increase or decrease as a result of currency and exchange rate fluctuations.

Fund charges will impact the value of your investment. In particular, the ongoing charges applicable to each fund will dilute investment performance, particularly over time. For further explanation of charges please visit our Fund Charges page at www.janushenderson.com.

For EU investors

Lower Risk 1 2 3 4 5 6 7 Higher Risk

Source: Janus Henderson Investors, as at 31 July 2023. Risk indicator refers to I2 USD share class. Rating is latest available as at 01 January 2023.

Note: UK investors should refer to the Key Investor Information Document as the risk rating may differ.

With effect from 1 January 2023, the Key Investor Information document (KIID) changed to the Key Information Document (KID), except in the UK where investors should continue to refer to the KIID. Availability of share classes shown may be limited by law in certain jurisdictions. Performance records/scenarios are detailed within the Fund's specific KIID/KID; fees and charges, and the respective risk rating may vary. Further information can be found in the Fund's prospectus and KIID/KID, which must be reviewed before investing. Please consult your local sales representative if you have any queries.

Biotechnology carve-out

Biotech carve-out outperformed NASDAQ Biotechnology Index in 14 of 16 years since PM inception*

Calendar Year Returns (%)	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022
Global Life Sciences Strategy - Biotechnology Carve-Out (Gross)	17.89	-18.57	19.37	16.24	18.07	46.98	79.19	51.05	16.02	-15.91	25.86	-4.66	35.59	49.95	-15.24	11.63
Global Life Sciences Strategy - Biotechnology Carve-Out (Net)	17.39	-19.11	18.60	15.49	17.32	46.05	78.08	50.10	15.28	-16.46	25.06	-5.29	34.73	49.01	-15.79	10.91
NASDAQ Biotechnology TR Index SM	-0.76	-12.56	15.84	15.25	12.08	32.29	65.97	34.40	11.77	-21.35	21.63	-8.86	25.11	26.42	0.02	-10.12
Difference (Gross vs. Index)	+18.65	-6.01	+3.53	+0.99	+5.99	+14.68	+13.22	+16.64	+4.25	+5.44	+4.23	+4.20	+10.48	+23.53	-15.26	+21.75

Performance – USD (%)	1 Year	3 Year	5 Year	10 Year	Since PM Inception*
Global Life Sciences Strategy – Biotechnology Carve-Out (Gross)	11.63	12.37	12.90	19.81	18.40
Global Life Sciences Strategy – Biotechnology Carve-Out (Net)	10.91	11.65	12.17	19.05	17.64
NASDAQ Biotechnology TR Index SM	-10.12	4.36	5.32	11.95	11.24
Difference (Gross vs. Index)	+21.75	+8.01	+7.58	+7.87	+7.15

Source: Janus Henderson Investors, as at 31 December 2022.

Note: Global Life – Biotech Sector – is an in-house carve-out from the representative account for the broader Janus Henderson Global Life Sciences Strategy. Above performance does not represent the investment performance or the actual accounts of any investors.

Benchmark: NASDAQ Biotechnology TR Index.

*PM Inception date of 01 May 2007.

Returns greater than one year are annualised.

Global Life – Biotech Sector – is an in-house carve-out from the representative account for the broader Janus Henderson Global Life Sciences Strategy. This performance does not represent the investment performance or the actual accounts of any investors.

The rates of return are hypothetical and do not represent the returns of an actual investment.

Past performance does not predict future returns. Investing involves risk, including the possible loss of principal and fluctuation of value.

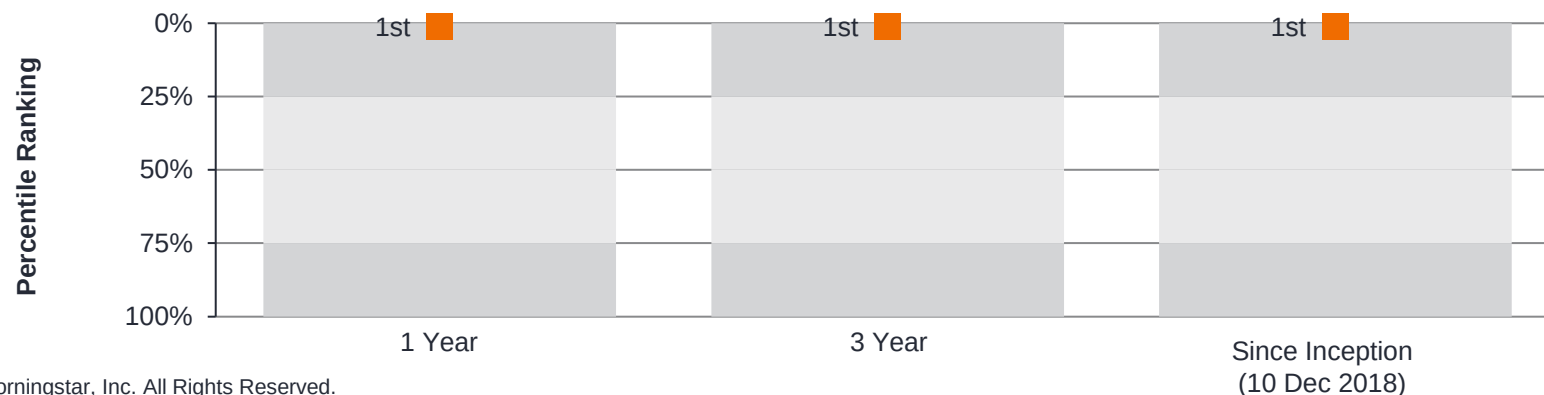
Performance

Horizon Biotechnology Fund

Performance – USD (%)	1 month	3 month	YTD	1 year	3 year	Since inception (10 Dec 18)
Horizon Biotechnology Fund – I2 USD Shares	1.61	6.42	12.37	46.39	11.81	17.21
NASDAQ Biotechnology TR Index SM	1.30	-1.15	-1.55	6.65	-0.01	5.89
SPDR® S&P® Biotech ETF	1.24	5.03	1.48	3.94	-8.18	1.39
Difference (Fund vs NASDAQ Biotech TR IndexSM)	+0.31	+7.57	+13.92	+39.74	+11.82	+11.32
Difference (Fund vs SPDR® S&P® Biotech ETF)	+0.37	+1.39	+10.89	+42.45	+19.99	+15.82

Performance Rankings

■ Horizon Biotechnology Fund – I2 USD Shares



Source: Janus Henderson Investors, Morningstar as at 31 July 2023. © 2023 Morningstar, Inc. All Rights Reserved.

Note: Fund: Janus Henderson Horizon Biotechnology, I2 Shares, net of fees in USD. Fund Inception: 10 December 2018.

Benchmark: NASDAQ Biotechnology TR IndexSM.

SPDR® S&P® Biotech ETF is designed to correspond to the performance of a modified equal weighting of the S&P® Biotechnology Select IndustryTM Index.

Returns include reinvestment of dividends and capital gains. Returns greater than one year are annualised. Janus Henderson Horizon Biotechnology Fund – A2 USD Shares Morningstar ranking compiled using Sector Equity Biotechnology category in USD net of fees.

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Why is health care attractive now?

Accelerating innovation, long-term growth potential, and attractive valuations

- **ACCELERATING INNOVATION:** Biotech companies are rapidly advancing our genetic understanding and ability to treat major diseases with new treatment modalities.
- **DEFENSIVE CHARACTERISTICS:** In the past five significant market downturns, healthcare fell roughly half as much as the overall market.
- **VALUATIONS:** After suffering its worst drawdown on record – biotech is trading at a substantial discount to the broader market, leading to “innovation on sale.”
- **M&A POTENTIAL:** Large-cap biopharma will hold in excess of \$500 bn in cash on their balance sheets – and have drug pipelines that need refilling.
- **LONG-TERM GROWTH POTENTIAL:** Innovation, aging populations, and globalisation should continue to support an attractive fundamental backdrop.



Note: The above are the Portfolio Manager's views and should not be construed as advice and may not reflect other opinions in the organisation. The views are subject to change without notice.

Appendix

Performance

Global Life Sciences Fund (Class I2 EUR Shares)

Performance – EUR (%)	YTD	1 year	3 year	5 year	Since inception (20/10/14)
Global Life Sciences Fund (Class I2 EUR Shares)	1.18	3.79	9.83	10.24	11.37
MSCI World Health Care Index SM	-1.18	-3.57	10.09	10.32	10.82
Difference (Fund vs Index)	+2.36	+7.36	-0.26	-0.08	+0.55

Source: Janus Henderson Investors, Morningstar, as at 31 July 2023. © 2023 Morningstar, Inc. All Rights Reserved.

Note: Fund: Janus Henderson Global Life Sciences Fund, I2 Shares, net of fees in EUR

Returns include reinvestment of dividends and capital gains. Returns greater than one year are annualised.

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SFDR Article 8 Application

FIRMWIDE EXCLUSIONS

Manufacturer or shareholding of ≥20% or greater in a manufacturer of controversial weapons

- Cluster munitions
- Chemical weapons
- Anti-personnel mines
- Biological weapons

FUND EXCLUSIONS

- Non-compliance of the UN Global Compact Principles and OECD
- 80% of the issuers covered by MSCI must be rated BB or above as per the ESG rating*

ENVIRONMENTAL AND/OR SOCIAL CHARACTERISTICS

- Support for the United Nations Global Compact principles
- Avoids issuers with the worst ESG risk ratings

Note: Please refer to the Prospectus for more detail. UN Global Compact cover matters including human rights, labour, corruption, and environmental pollution.
*By MSCI – <https://www.msci.com/>, or equivalent. OECD: Organisation for Economic Co-operation and Development.

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Important information

In accordance with the Sustainable Finance Disclosure Regulation, the Fund is classified as Article 8 and promotes, among other characteristics, environmental and/or social characteristics, and invests in companies with good governance practices.

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(continued on next page)

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The Morningstar Rating™ for funds, or "star rating", is calculated for funds with at least a three-year history. Exchange-traded funds and open-ended mutual funds are considered a single population for comparative purposes. It is calculated based on a Morningstar Risk-Adjusted Return measure that accounts for variation in a fund's monthly excess performance, placing more emphasis on downward variations and rewarding consistent performance. The Morningstar Rating does not include any adjustment for sales loads. The top 10% of funds in each category receive 5 stars, the next 22.5% receive 4 stars, the next 35% receive 3 stars, the next 22.5% receive 2 stars, and the bottom 10% receive 1 star. The Overall Morningstar Rating for a fund is derived from a weighted average of the performance figures associated with its three-, five-, and 10-year (if applicable) Morningstar Rating metrics, and may not indicate positive performance. Ratings may vary by share class.

The Morningstar percentile ranking is based on a fund's total return (including income and capital gains, if any, and excluding sales charges) relative to all funds in the same category for the period, and may not indicate positive performance. The highest (or most favorable) percentile rank is 1%, and the lowest (or least favorable) percentile rank is 100%. The top-performing funds in a category will always receive a rank of 1.

Ratings and/or rankings may be based, in part, on the performance of a predecessor fund or share class and are calculated by Morningstar using a methodology that differs from that used by Janus Henderson. Methodology differences may have a material effect on the return and therefore the rating/ranking.

Contact us

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Argentina Investors

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Brazilian Investors

The shares in the fund may not be offered or sold to the public in Brazil. Accordingly, the shares in the fund have not been and will not be registered with the Brazilian Securities and Exchange Commission (Comissão de Valores Mobiliários, the “CVM”), nor have been submitted to the foregoing agency for approval. Documents relating to the shares in the fund, as well as the information contained therein, may not be supplied to the public in Brazil, as the offering of the shares in the fund is not a public offering of shares in Brazil, nor used in connection with any offer for subscription or sale of securities to the public in Brazil. A seller of the shares in the fund may be asked by the purchaser to comply with procedural requirements to evidence previous title to the shares in the fund and may be subject to Brazilian tax on capital gains which may be withheld from the sale price. Persons wishing to offer or acquire the shares in the fund within Brazil should consult with their own counsel as to the applicability of these registration requirements or any exemption therefrom.

Chilean Investors

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An English translation:

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