

Major evolutions & trends in the fight against cancer

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Head of Healthcare

OCT. 2024



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Fondo Classe Retail (DXL)

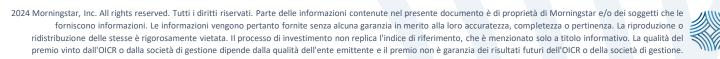
LUX IM ESG CANDRIAM Oncology Science LU2344409185

BG Stile Libero

Fondo	Rating Morningstar	Classe Istituzionale
CANDRIAM Equities L Oncology Impact (Euro hdg)	***	LU2015349330
CANDRIAM Equities L Biotechnology	****	LU1269736838

Coming Soon – Q1 2025 – in distribuzione diretta / Assicurativo

Fondo	Rating Morningstar	Classe Retail
CANDRIAM Equities L Biotechnology	****	LU1120766388
CANDRIAM Equities L Life Care	***	LU2223680658





Speaker

Servaas Michielssens

Head of Healthcare



Are we winning the fight against cancer?

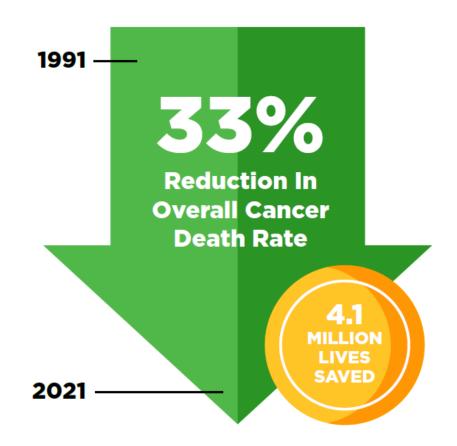


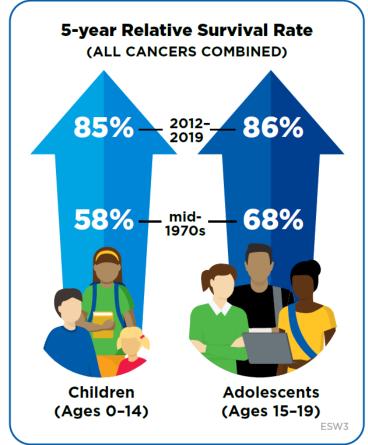
Survival rates are improving

Notable improvements in cancer survival

thanks to earlier detection and/or advances in treatment

US trends in survival rates

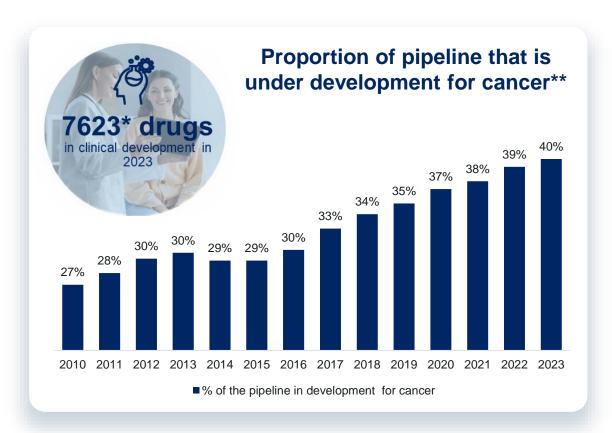


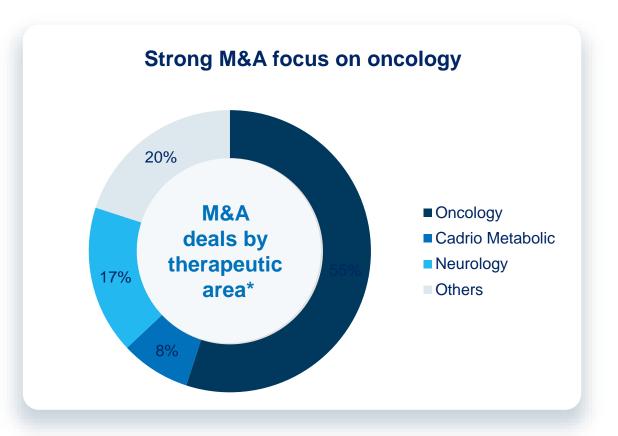




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Oncology continues to be the majority focus for sector R&D spend and M&A





Source: *Charles River Laboratories: presentation of the 39th Annual J.P. Morgan Healthcare Conference in 2021; McKinsey Oncology Market Outlook 2020 – Compounds in Clinical Development (Phase I-III) excluding Reformulations and Biosimilars, Pharmaprojects, 2019, pharmaintelligence.informa.com, representing approximately one-third of the global clinical pipeline across all therapeutic areas. **Global R&D Trends 2023: Activity, Productivity and Enablers

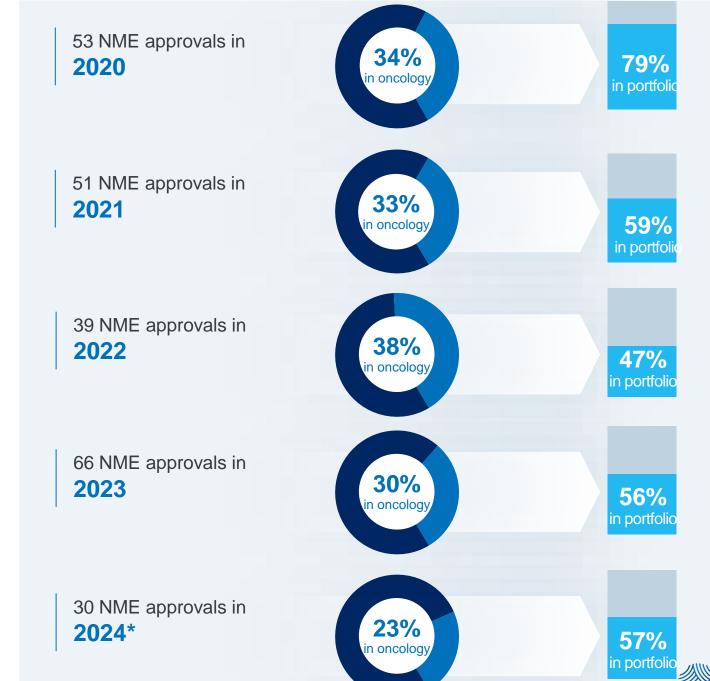


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A tangible impact: the Candriam strategy is involved in the majority of novel oncology drug approvals

With approximately one third of total new drug approvals (NME: new molecular entities), the oncology space takes a prominent place;

The Candriam strategy was involved in the vast majority of those oncology approvals.



Despite recent advances, the patient outlook is often still poor

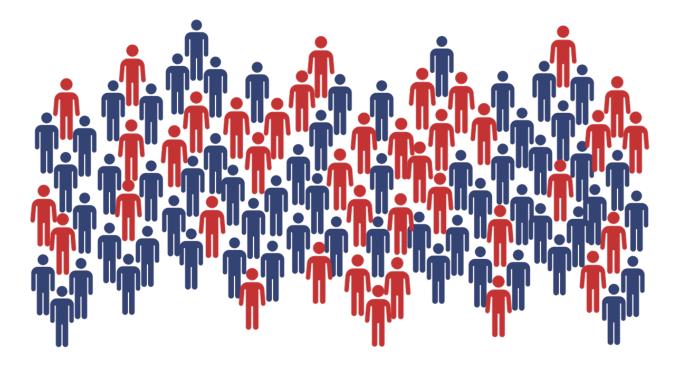


~1,600
DEATHS/DAY

due to cancer in the US

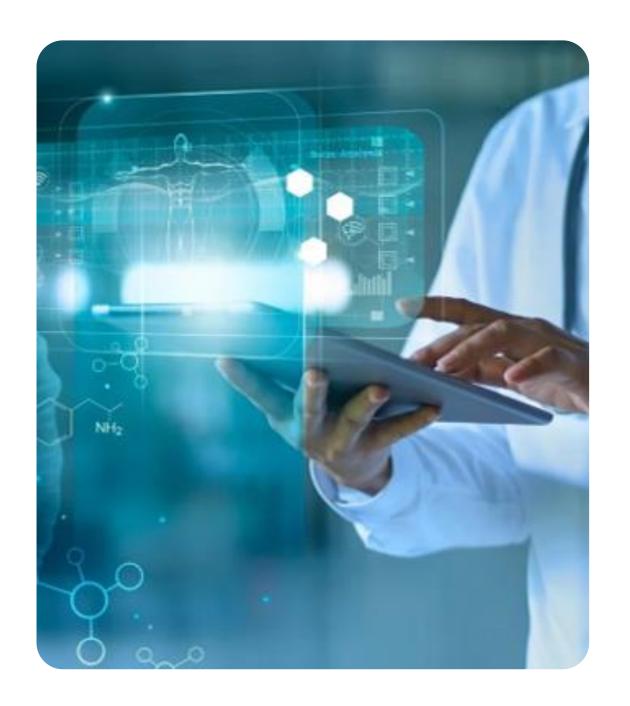
Cancer is the 2nd-leading cause of death globally ~40% LIFETIME RISK

38.4% of men & women will be diagnosed with cancer at some point during their lifetimes



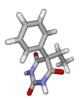


Cutting-edge evolutions



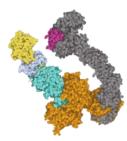
What is next? Innovation based on precision, selectivity and technology

Precision



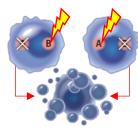
Small Molecules

Although small molecules have been used for decades, there is still room for innovation. Several generations of kinase inhibitors with increasing efficacy and safety.



Protein Degraders

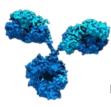
New class of therapeutic solutions based on small molecules that not only inhibit but entirely eliminate proteins responsible for various diseases, previously considered undruggable.



Synthetic Lethality

Exploiting the interactions between pairs of genes in cancer cells for therapeutic purposes: perturbation of either gene alone is viable but inhibiting the product of one gene while the other one is mutated leads to cell death.

Selectivity



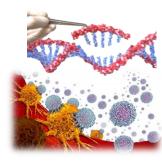
Antibody – Drug Conjugates

Designed to act more precisely on the cells that cause disease, without targeting healthy cells.



Bispecific antibodies

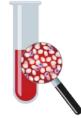
Antibodies with two targets, e.g., T-cell activators, to stimulate the natural immune system to fight the disease.



Gene and Cell Therapies

Modify the patient's genes to treat the disease, increase the level of a missing enzyme, or repair a genetic defect.

Technology



Liquid Biopsy

Non invasive diagnostic technique that allows for detecting cancerous cells or their DNA circulating in the blood sample.



Artificial intelligence assisted imaging - impressive accuracy and sensitivity of diagnosis



RNA Therapeutics

Allowing for modulation of protein expression, e.g., to reduce the levels of proteins responsible for a disease.



Exosomes

Extracellular vesicles generated by the cells, which can be used as targeted drug delivery vehicles, potentially overcoming the challenges of systemic toxicity

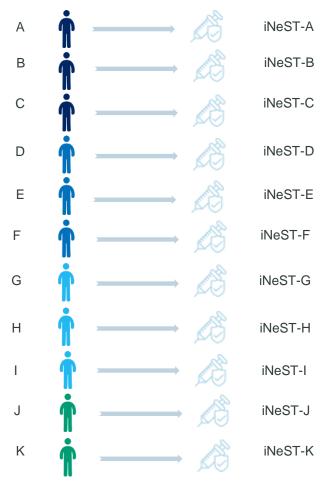


Cancer vaccines: a new era in individualized cancer treatment

- •Many tumor mutations are unique for each patient;
- Each vaccine can be uniquely tailored to each patient.

Personalized cancer therapy Patients with a type of cancer

Individualized cancer vaccine





Cancer vaccines: a new era in individualized cancer treatment

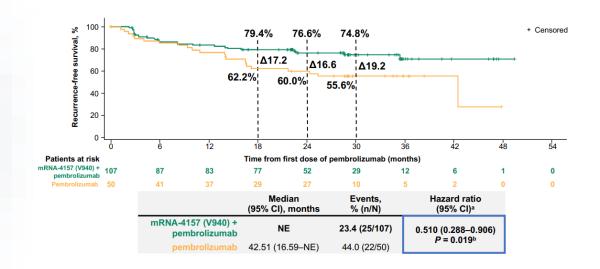
Impressive data from Moderna and Merck

~50%

less risk of recurrence or death.

74.8%

of the treated melanoma patients **remain cancer free** post surgery at 2.5 years.



Legend: CAR-T against BCMA target

CAR-T against BCMA target with the **best clinical outcome for multiple myeloma patients** who failed front- line therapy (Darzalex based regimens).

- > BCMA CAR-T, Carvykti, was approved in 2022 and offers the highest selectivity against tumor cells. Increased tumor target selectivity remains key to deliver more precise and chronic treatments to patients, while avoiding potential off target toxicities.
- Although many medicines are available for multiple myeloma patients, BCMA CAR-T is the only one to offer <u>one and done</u> modality.
- > BCMA CAR-T is a live therapy with higher probability of success.

CAR T-cell Therapy Remove blood from Make CAR T cells in the lab patient to get T cells Insert gene for CAR Chimeric antigen receptor (CAR) **CAR T cell** CAR T cells bind to cancer cells and kill them Cancer cell Grow millions of **CAR T cells** CAR T cell Infuse CAR T cells Cancer cell into patient (ciltacabtagene autoleucel) Suspension for IV Infusion Dead cancer BCMA-directed CAR-T therapy; delivered as a single infusion*3 *Although delivered as a single infusion, therapy with CARVYKTI® is part of a multi-step process.3



A.I. in health care



Impactful ways artificial intelligence improves healthcare



Facilities



Target discovery & drug design



Clinical development



Medical imaging



Diagnostics



Procedures & solutions

- Reduced administrative burden
- Towards reliable care
- Example: HCA Healthcare
- Faster development Safer, better drugs
- Example: Amgen
- The right drug for > the right patient
- Example: IQVIA
- Faster , assessment and , more precise medical imaging
- Example: Hologic
- Early diagnosis
 More convenient
- Example: Natera
- Safer procedures
- Less staff needed
- Example: Intuitive surgical



Revolutionizing drug development

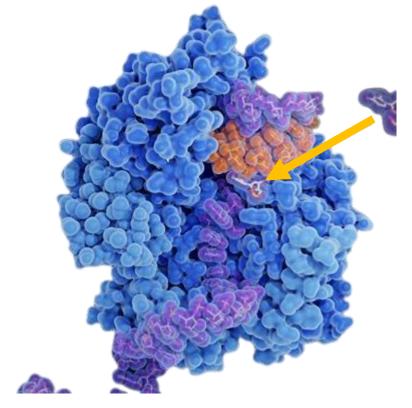
Background: proteins are key targets for drug discovery. The availability of correct protein structures is greatly enhancing the rational development of safe and efficacious drugs.



> The problem

For only ~0.1% of all proteins, an experimental structure is known.

Predicting that structure via simulations is an enormous challenge, as for a typical proteins there are more possibilities than atoms in the universe.



Picture source: Pharmacokentics



> The solution

A.I. made it possible to predict the structures of virtually all proteins, dramatically expanding the available targets for rational drug design.



Leveraging on DeepMind's ML computational approach to predict the structures of proteins

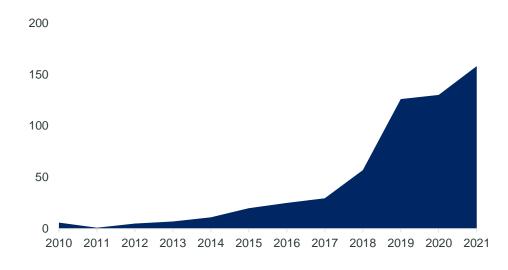




Drug development: A.I has the potential to fast-track drug discovery



A.I. co-discovered drugs - N° of assets in discovery and clinical trials*



^{*} Source: Candriam, Citigroup, Nature Reviews Drug Discovery via Recursion



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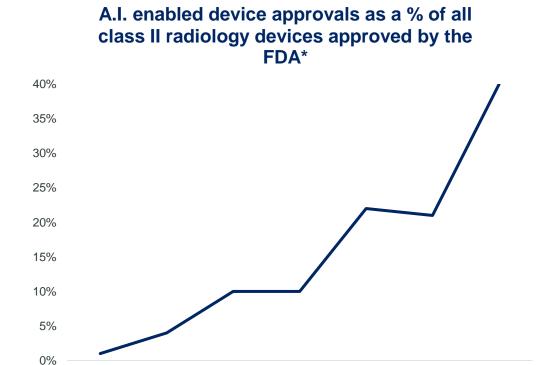
Hologic: Genius A.I.® Detection Solution

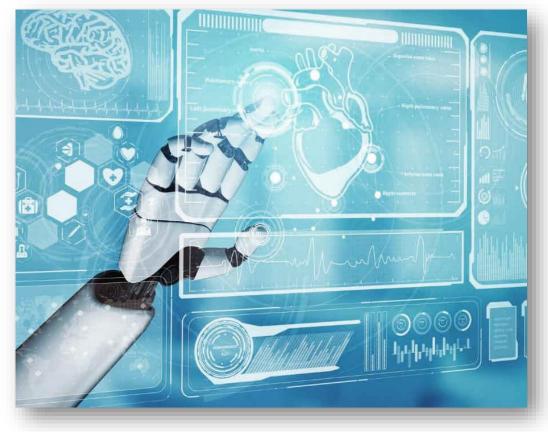


Genius Cervical A.I.®*:

- First and only FDA-cleared digital cytology system, launched this year.
- Combines deep-learning-based artificial intelligence with advanced volumetric imaging technology.
- 28% reduction in false negatives of high-grade squamous intraepithelial and more severe lesions compared to microscopic review.

Growing number of A.I. enabled device approvals







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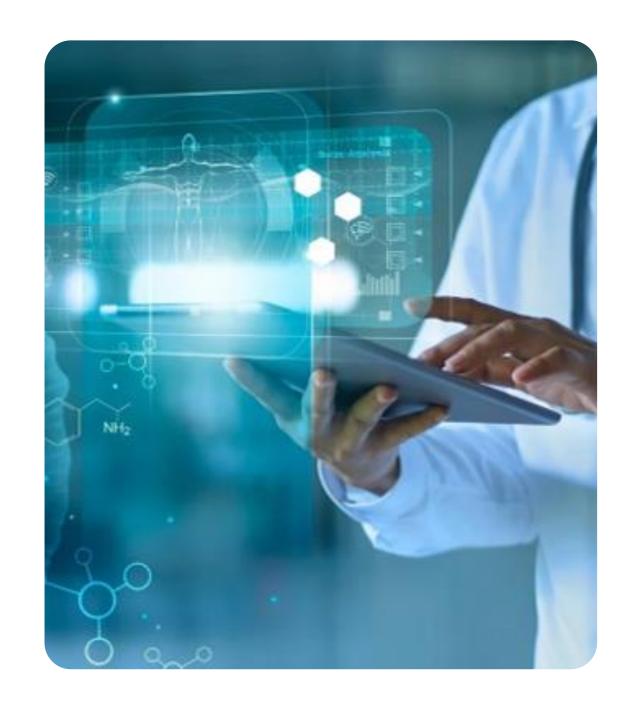
Intuitive Surgical: artificial intelligence improves robotic surgery

- Assessing the execution of a surgical procedure is not straightforward.
- Robotic-assisted surgery creates a wealth of (digital) data.
- Surgeons can compare this with their own modus operandi and learn/improve their skills very effectively via modules that Intuitive is making available.
- 3D modeling with A.I. helps surgeons pre-operatively understand a patient's specific anatomy for better surgical planning on the Da Vinci robot.



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Outlook and conclusions



Candriam supports cancer research





10%

Candriam pledges maximally
10% of the net management
fees of the Candriam
Equities L Oncology Impact
sub-fund to leading cancer
research institutes

BIG against breast cancer





€3.3m

donated
since 2018











Damit Heilung zur Regel wird.





Track Record Oncology Impact – Since Inception



Fund: CANDRIAM Equities L Oncology Impact - retail Class- Perf in Euro - LU1864481467

Data as at 30/09/2024. The fund is actively managed and the investment process implies referring to a benchmark index (MSCI World NR© (since 24/11/2020). Past performance is no quarantee of future results and is not constant over time. © 2024 MSCI Inc. Áll rights reserved. For further details on risks associated with investing in these strategies, a general description and explanation of the various risk factors is available in the section Risk Factors of the relevant Prospectuses. The prospectus is available on

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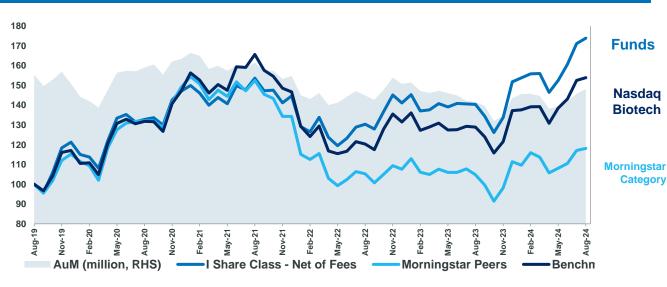


Biotechnology: Performance Strategy

Performance From 2000 –Retail Class- LU0108459040 Grafico crescita (EUR) 04-10-2024 Candriam Eqs L Biotech C USD Cap: 79.966,21 Morningstar Gbl Biotechnology N...: 61.651,57 × Azionari Settore Biotecnologia: 32.080,13 × 80.0K 70.0K 60.0K 50.0K 40.0K 30.0K 20.0K

Funds – CANDRIAM Equities L Biotechnology - LU0108459040 Retail Class
Morningstar Category: EAA Fund Sector Equity Biotechnology - Equity Market: Morningstar Global Biotechnology NR USD - Total Return
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factors is available in the section Risk Factors of the relevant Prospectuses. The prospectus is available on Candriam.com: Fund Detail.

Cumulative Performance & AuM over 5 years



Past performance is no guarantee of future results and is not constant over time.

Monthly data as at 31/08/2024. I Share Class USD, Net of fees. The fund is actively managed and the investment process implies referring to a benchmark index the NASDAQ Biotechnology Index NR ©. Past performance is no guarantee of future results and is not constant over time. Performances expressed in a currency other than that of the investor's country of residence are subject to exchange rate fluctuations, with a negative or positive impact on gains. If the present document refers to a specific tax treatment, such information depends on the individual situation of each investor and may change. Morningstar Category: EAA Fund Sector Equity Biotechnology ©2024Morningstar. All Rights Reserved. The information contained here: (1) is proprietary to Morningstar and/or its content providers; (2) may not be copied or distributed; and (3) is not warranted to be accurate, complete or timely. The quality of the ranking, award or label obtained by the fund or the management company depends on the quality of the issuing institution and the ranking, award or label does not guarantee the future results of the fund or the management company. For further details on risks associated with investing in these strategies, a general description and explanation of the various risk factors is available in the section Risk Factors of the relevant Prospectuses. The prospectus is available on Candriam.com; Fund Detail.

retail Class – LU0108459040 Update to 30 sept 2024	YTD	2023	2022	3 Y	5 Y	10 Y
CANDRIAM Equities L Biotechnology	+10.4%	+3.5%	+5.0%	+24.0%	+71.6%	+138.9%
Biotech Fund Category	+2.8%	+1.7%	-12.8%	-9.9%	+28.5%	+61.0%
Delta	+7.6%	+1.8%	+17.8%	+33.9%	+43.1%	+77.9%

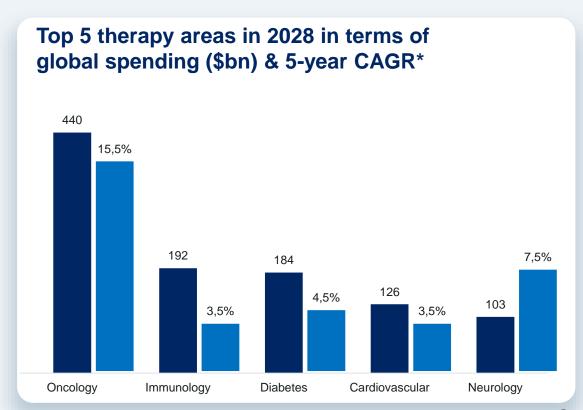
10.0K

2024

Morningstar Rating				
Overall	00000			
3 Years	00000			
5 Years	00000			



Continued R&D investments lead to success in the clinic and growing sales



Growth drivers in addition to increased early diagnoses and wider treatment availability



+100

drug approvals
including cell therapy,
RNA therapy and
immuno-oncology

treatments.



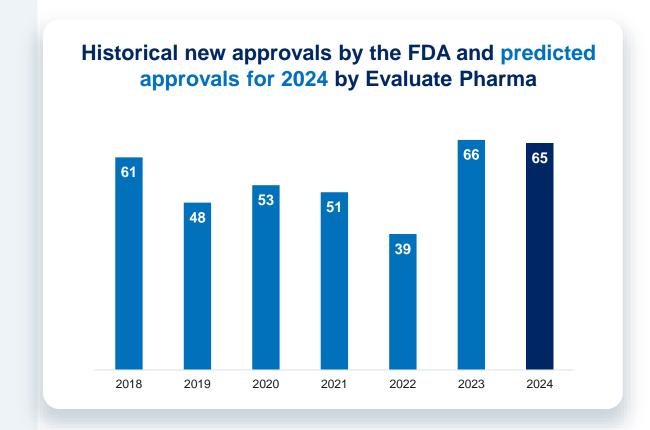
Precision medicine

including personalized treatment determined by biomarker testing & nextgen sequencing or Car Tcell therapies.

Source: Source: IQVIA Forecast Link, IQVIA Institute, December 2023 * median growth forecast; Covid vaccine and therapeutics are not included. Oncology includes therapeutic oncology only and not supportive care. Immunology includes small molecule and biologic treatments for a range of diseases. Neurology includes central nervous system disorder treatments and mental health treatments but excludes pain management and anesthesia. Pain includes narcotic and non-narcotic analgesics, muscle relaxants and migraine treatments. Cardiovascular includes hypertension and other cardiovascular treatments with the exception of lipid regulators.

2023 was a record year for new drug approvals





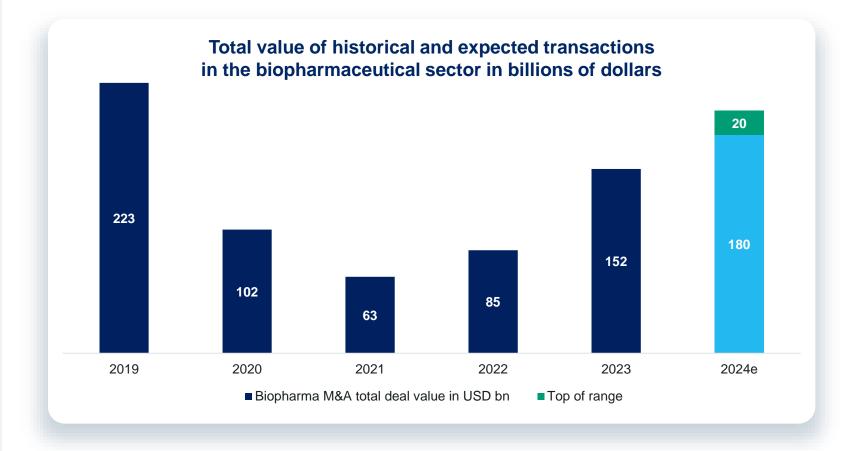
M&A activity should remain high







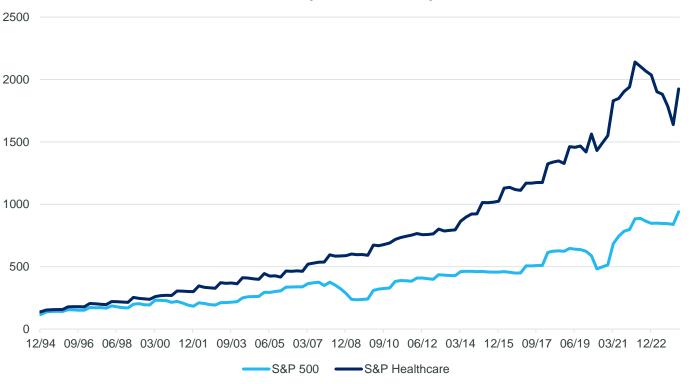




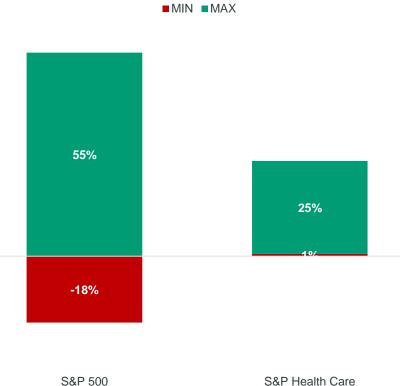


A sector that is less dependent on the economic environment





Forward EPS Y/Y (1995 - 2022*)



Valuations are reasonable today



12-month trailing earnings per share

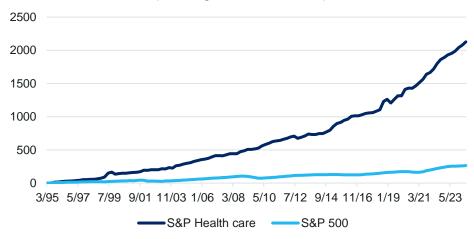
S&P Health care S&P 500

(% change since rebased 31/03/1995)



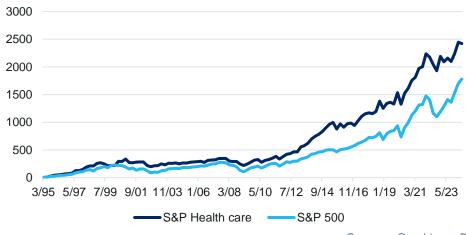
Sales/share

(% change since 31/03/1995)



Return comparison

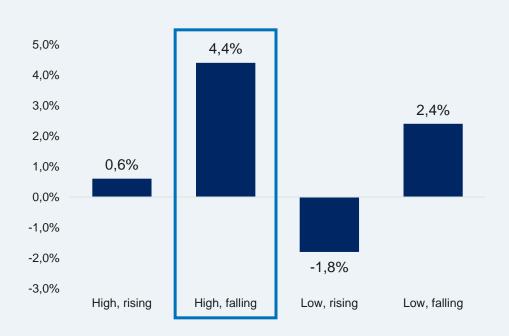
(% since 31/03/1995 – end of June 2024)





Environment supportive of biotechnology sector performance

Biotechnology relative performance during different interest rate regimes (avg quarterly)*



Nasdaq Biotechnology Index vs. 10-year US yield from 2006 to present.



Past performance is no guarantee of future results and is not constant over time



Valuation clinical-stage companies: reasonable from a historical view

Small/mid-cap biotechnology





Conclusion

- Innovation and investments in the oncology sector are thriving, leading to better outcomes for patients combined and double-digit sales growth rates
- > Drug approvals are on track in 2024, oncology continues to take an important share
- > A.I. is impacting the healthcare sector and supports innovation and improved outcomes
- > Macro-economic environment and valuations are supportive for the healthcare sector





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Swiss paying agent: CACEIS Bank, Montrouge, succursale de Nyon/Suisse, Route de Signy, 35, CH-1260 Nyon. Place of performance: Route de Signy 35, CH-1260 Nyon. Place of jurisdiction: Route de Signy 35, CH-1260 Nyon.

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Client Relation Offices

Management Centers

