

# Artificial Intelligence and Biosensors in Healthcare and its Clinical Relevance: A Review

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**Abstract**—Data generated from sources such as wearable sensors, medical imaging, personal health records, pathology records, and public health organizations have resulted in a massive information increase in the medical sciences over the last decade. Advances in computational hardware, such as cloud computing, Graphical Processing Units (GPUs), and Tensor Processing Units (TPUs), provide the means to utilize these data. Consequently, many Artificial Intelligence (AI)-based methods have been developed to infer from large healthcare data. Here, we present an overview of recent progress in artificial intelligence and biosensors in medical and life sciences. We discuss the role of machine learning in medical imaging, precision medicine, and biosensors for the Internet of Things (IoT). We review the most recent advancements in wearable biosensing technologies that use AI to assist in monitoring bodily electro-physiological and electro-chemical signals and disease diagnosis, demonstrating the trend towards personalized medicine with highly effective, inexpensive, and precise point-of-care treatment. Furthermore, an overview of the advances in computing technologies, such as accelerated artificial intelligence, edge computing, and federated learning for medical data, are also documented. Finally, we investigate challenges in data-driven AI approaches, the potential issues that biosensors and IoT-based healthcare generate, and the distribution shifts that occur among different data modalities, concluding with an overview of future prospects.

**Index Terms**—Artificial Intelligence, Explainable AI, Medical Imaging, Domain Adaptation, Biosensors, Federated Learning

## I. INTRODUCTION

About 10% of global gross domestic product (GDP) (10 trillion USD) is spent on healthcare annually [1]. The recent advancements in technology, especially data-driven methods and computational processing power can benefit, both the patients and the medical industry, as well as reduce the huge expenditures. Moreover, massive healthcare data is available from sources such as; electronic health records

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(EHRs), genomics profiles, medical imaging, chemical, and drug databases. Analytical methods, especially deep learning-based Artificial Intelligence (AI) methods, can provide the tools to design useful clinical and medical applications to process these large datasets. Data-driven methods could offer benefits in medical record digitization, clinical trials, diagnosis assistance, prognosis evaluation, and the design of optimal prevention and treatment strategies, as well as precision medicine, drug discovery, and health policy.

Advances in computational infrastructure have provided the capacity to generate, store, analyze and visualize large, complex, and dynamic datasets typical of modern biomedical studies [2]. New treatment options are being developed and tested in clinical trials [3]. In the last decade, artificial intelligence has moved from theoretical studies to real-time applications thanks to the rise in the computational capacity of GPUs and TPUs. Methods like AutoML [4] and explainable artificial intelligence (XAI) [5] are advancing, which have the potential to transform the current medical practice. However, there are still many bottlenecks to realizing the full potential of analytical methods in the healthcare industry. Important challenges for data science in medicine include data collection, standardization of data formats, missing data values, developing large and efficient computational infrastructure, data privacy and security, and others.

For example, to deal with the small sample size issue in medical images, generative models can be used to generate synthetic medical images of high quality. Generative Adversarial Network (GAN), a type of neural network that can generate synthetic data, can be used to generate synthetic magnetic resonance imaging (MRI) scans or positron emission tomography (PET)-scan images using computed tomography (CT) scans. A subset of images, regardless of size, is a subset of the universal set. Using that small subset, generative models learn the probability distribution of the universal training set. After extracting the representative features, the model can generate high-quality synthetic images by sampling from the probability distribution. These synthetic images can be used to build generalized medical image analysis models for various clinical applications.

The interrelated nature of biomedical data is one of its most important properties. Such data can be represented in the form of graphs. Graph machine learning allows for the modeling of unstructured multimodal datasets. Graph machine learning can model more complex relationships between disease and patients, understand tumor micro-environment, predict

drug response, and re-purposing. Additionally, graph machine learning coupled with attention mechanism may provide more interpretable machine learning models than typical traditional black-box models.

The recent breakthrough of the artificial intelligence (AI) system Alphafold2 [6] in predicting the three-dimensional structure of proteins solely from the amino acid sequence is a huge success. AlphaFold2 won the Critical Assessment of Structure Prediction (CASP) [7], the worldwide event for protein structure prediction, since 1994. Meta AI also joined the race and developed an AI system to predict structures of about 600 million proteins [8]. However, how to translate this into the in vivo situation is still an open question. AlphaFold2 can predict unbound protein structures; however, most practical applications require protein-drug complex predictions.

There have also been significant advancements in processing power and biosensor technologies. For example, with the help of parallel processing methods and powerful GPU clusters, such as NVIDIA-DGX, we can now process massive complex multi-dimensional biomedical datasets [9]. Moreover, wearable electronics, such as electronic tattoos (E-tattoos), epidermal electronics systems (EES), and flexible electrochemical bioelectronics, coupled with machine learning algorithms can be used to monitor various biomarkers in real time [10].

As the use of AI in healthcare has been a very active research area, several surveys were found covering this topic [11]–[13]. In [11], a discussion about the use of medical sensors with artificial intelligence is presented. In this respect, various sensing systems and the use of AI in medical decision-making are studied. The study in [12] provides coverage of the different wearable sensors for healthcare delivery, primarily from a hardware perspective, and briefly highlights the benefits and challenges of AI. More recent work [13] covers the use of AI in the internet of medical things and its different applications concerning various algorithms. AI methods for combating various medical diseases were also discussed. A survey about AutoML was presented in [14].

Given the enormous progress in recent years for AI in healthcare, an updated review will benefit the community. In this article, we present an updated survey of the recent progress in data-driven methods for healthcare. We specifically discuss practical applications of artificial intelligence, biosensors, and computational infrastructure, concerning clinical relevance. The recent methods which have the potential to become a part of the healthcare industry, such as AutoML [15], explainableAI [16], and Federated learning [17] are evaluated. Moreover, existing clinical tools and emerging AI-based start-up companies are presented. We also highlight the existing challenges for AI in healthcare and present some potential solutions. The use of AI for drug discovery, nano-medicine, and medical robotics is out of the scope of this review. The survey is organized as follows; Section II highlights applications of machine learning in various healthcare sectors. AI-based clinical tools and start-up companies are presented in Section III. Sections IV and V discuss applications of big data analytics and biosensors, respectively. Computational advances, federated learning, and edge computing are discussed in Section VI. The recent challenges in AI for healthcare with

potential solutions are explored in Section VII, and Section VIII concludes this review.

## II. MACHINE LEARNING IN HEALTHCARE

Data science and machine learning have been successful in many areas related to computer vision, such as self-driving cars, recognizing actions, image classification, and intelligent robots. These are well-posed tasks where the problem is known, and the solution is verifiable. However, healthcare-related tasks involve safety and security risks, leading to privacy concerns. These problems are neither well-posed nor well-defined, and their solutions can be hard to verify. Assessing the risk of life-threatening disease in people infected with the SARS-CoV-2 virus is a recent broad, complex, and urgent problem where data science has been used to suggest prognostic indicators from a wide variety of genetic and physiological markers and the presentation of symptoms [18]. Figure 1 shows an ecosystem for machine learning in healthcare tasks. Machine learning can produce actionable insights for clinical practice, provide recommendations to governments for optimal health policy, and help accelerate and optimize drug discovery and design processes. More established use cases of different machine learning applications in healthcare are presented in Table I.

### A. Explainable Artificial Intelligence

While machine learning models applied to biomedical data, have the potential to produce clinically useful judgments, the models, particularly deep learning, are frequently regarded as black boxes that are difficult for humans to understand [5]. This lack of transparency leads to a bottleneck in the clinical implementation of machine learning-based findings, as any decision will directly affect a patient’s health. One way to increase the transparency in machine learning predictions is to highlight the feature importance or to visualize features at different layers. This way, we can analyze each feature’s importance in the prediction model and better understand the predictions. One such method is known as Grad-CAM visualization [19], based on the target concept’s gradients, which flow into the final convolutional layer to build a coarse localization map highlighting significant locations or heat maps in the image for concept prediction. Explainable models, or explainable artificial intelligence, are needed to build the trust of healthcare professionals.

Explainable AI methods are classified based on the complexity and scope of their interpretability [20] and the level of dependencies in the AI model. Explainability has different levels of understanding, including interpretability, stability, robustness, and confidence. A user can not only see but also learn how inputs are mathematically transferred to outputs in an interpretable system, whereas a stable system is not misled by small perturbations or noise in the input data. The possibility of an event occurring is measured by confidence. The purpose is to quantify the level of confidence in the decision [21].

Complex deep learning models are generally less interpretable, and there can be a trade-off between accuracy and

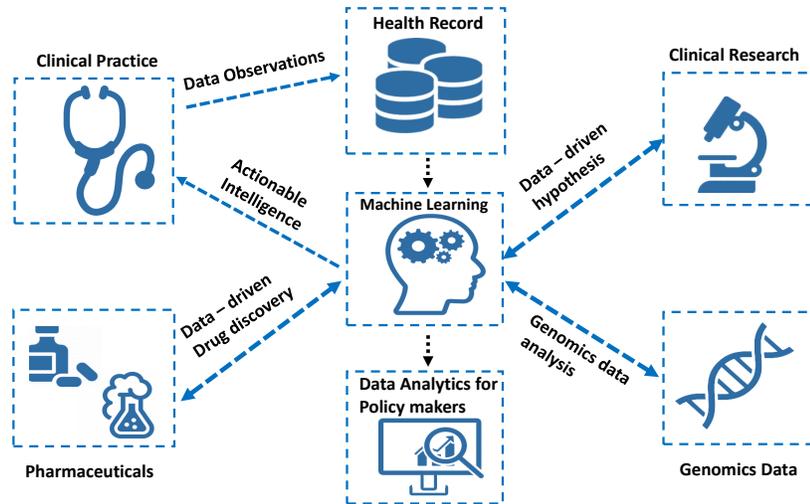


Fig. 1. An ecosystem for machine learning in the healthcare industry. Clinical decision support systems, policy-makers, and pharmaceutical companies can benefit from machine learning methods.

195 interpretability. Easy-to-interpret models could be designed, 231  
 196 but they may compromise accuracy. Highly complex, un- 232  
 197 interpretable models with high accuracy that require a separate 233  
 198 set of algorithms for interpretation are more commonly used 234  
 199 in XAI. Another way to explainability is to check whether 235  
 200 the model is agnostic or model-specific. Agnostic methods 236  
 201 are used for any machine learning algorithm, such as neural 237  
 202 networks and support vector machines, while model-specific 238  
 203 methods are limited to interpreting the specific model [22]. 239

204 It is also important to consider human factors when en- 240  
 205 hancing the model interpretability, such as a medical expert, 241  
 206 to guarantee the interpretability and explanations of the model. 242  
 207 It is expected that Explainable AI will further advance research 243  
 208 in machine learning for healthcare as it solves the critical 244  
 209 challenges of healthcare, such as fairness, transparency, safety, 245  
 210 security, privacy, and trust. 246

211 1) *Human and Machine Interpretable Visualizations:* One 247  
 212 important aspect of Explainable AI is the use of human 248  
 213 interpretable visualizations that allow humans to understand 249  
 214 the reasoning behind AI models easily. For example, deci- 250  
 215 sion trees, rule lists, and other interpretable models can be 251  
 216 visualized in a way that is easy for humans to understand. 252  
 217 In addition to human-interpretable visualization techniques, 253  
 218 machine-interpretable visualization techniques are also impor- 254  
 219 tant in Explainable AI. These techniques enable AI models 255  
 220 to explain their predictions or decisions in a way that is 256  
 221 easily understandable by other AI systems. For example, 257  
 222 SHAP (SHapley Additive exPlanations) [23] is a machine- 258  
 223 interpretable visualization technique that can be used to ex- 259  
 224 plain the output of complex machine learning models, such as 260  
 225 deep neural networks. 261

226 However, deep learning models work differently than hu- 262  
 227 mans, and it is difficult to interpret a model with billions 263  
 228 of parameters. For example, if we visualize the grad-cam 264  
 229 heatmap for a dog, we can see that most of the heat is 265  
 230 concentrated around the dog's ears. Humans recognize dogs

by the uniqueness of their shape.

2) *Causal Inference:* Health science-related tasks demand 232  
 more explanation than mere predictions. With the abundance 233  
 of data, many deep learning algorithms just only look for 234  
 correlations among variables and make predictions or clas- 235  
 sifications without explaining the actual cause. To be practical 236  
 and utilized in daily clinics, machine learning models must 237  
 have strong causal evidence. Several methods are developed 238  
 to convert the deep learning black box to a white box, for ex- 239  
 ample, feature visualization [24], gradcam visualization [25], 240  
 regularization via causal graph discovery [26], causal-aware 241  
 imputation via learning missing data mechanisms [27], domain 242  
 adaptation [28], tools such as Shared Interest [29] and learning 243  
 generalized policies [30]. 244

245 The causality can be defined in three stages. First is the 246  
 association, for example, between the training image and 247  
 its label. The second is intervention, which aims to predict 248  
 the outcome based on altering the system (treatment plan 249  
 or patients). The last one is counterfactual, which predicts 250  
 the output in a different condition and environment. Causal 251  
 machine learning models can guide us to make informative and 252  
 timely interventions and rethink different treatment regimens 253  
 and outcomes.

### B. Machine learning for precision medicine 254

255 Traditional medical models have treated an average patient 256  
 with a 'one size fits all approach'. Precision medicine, which 257  
 takes treatment approaches based on an individual patient's 258  
 unique clinical, genetic, epigenetic, and environmental infor- 259  
 mation, is a growing field of healthcare, and it is becoming a 260  
 viable alternative due to the increase in the amount of medical 261  
 data [31]. In Figure 2, we show a conceptual diagram for 262  
 precision medicine by utilizing different data modalities.

263 Data, such as a patient's age, weight, blood pressure, 264  
 medical history, and genomic sequences, can be used by 265  
 analysis algorithms to identify hidden patterns and identify

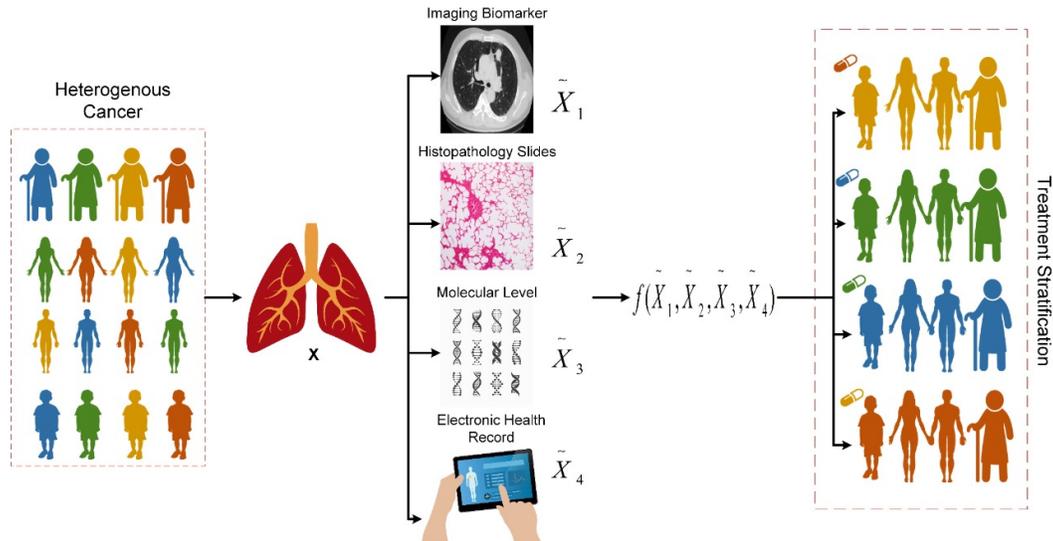


Fig. 2. A conceptual diagram for precision medicine, where different data modalities are used to find patient-specific features and treatment plans.

266 correlations between patient profiles and disease phenotypes.  
 267 A personalized drug response model developed for non-small  
 268 cell lung cancer patients [32] used the binding free energy of a  
 269 drug-mutant complex and personal features of the patient (age,  
 270 sex, smoking history, medical history) to build a personalized  
 271 drug prediction model. Extreme learning machines were used  
 272 to predict the drug response into two classes with an overall  
 273 accuracy of 95%, driven by the addition of personal features.  
 274 Personalized medicine is used for complex diseases such as  
 275 cancer, heart disease, and diabetes [33]. If it is used carefully,  
 276 this technology could improve performance in healthcare and  
 277 potentially reduce inequities (MIT-CSAIL).

### 278 C. AI in remote patient monitoring

279 The combination of edge artificial intelligence (machine  
 280 learning on edge devices) and the IoTs has facilitated the  
 281 deployment of remote healthcare systems. Such systems can  
 282 monitor a patient's vitals and other physiological parameters  
 283 in real-time while the patient remains at home and push it  
 284 to the cloud [34]. AI embedded in smart devices democratizes  
 285 healthcare by putting AI-enabled health services (for example,  
 286 AI-based clinical decision support) into patients' homes or  
 287 remote healthcare [35]. The centralized data gathered for the  
 288 patients can be used for knowledge discovery to improve  
 289 disease prognosis or by doctors to monitor the patient and  
 290 make/update prescriptions.

291 Several commercial wearable devices offer services mea-  
 292 suring physiological parameters such as heart rate, ECG, and  
 293 other variables through smartwatches and biosensors. There  
 294 have been considerable targeted systems proposed as well for  
 295 a variety of ailments, including but not limited to diabetes [36],  
 296 where devices can also be used for the management of insulin  
 297 as well [37], cardiac disease through ECG [38], sleep apnea  
 298 monitoring [39] or as generic monitoring platforms such as  
 299 smart-monitor [40] to provide 'a la carte' system based on the  
 300 patient health circumstances. Machine learning methods can

then be applied to these physiological signals for predictive  
 health management.

### III. CLINICAL AI TOOLS AND EMERGING AI HEALTHCARE COMPANIES

The primary question is when AI tools will be used in  
 ordinary clinical practice to support real-time health chal-  
 lenges, such as improved diagnostic and clinical decision  
 support systems [41]. Despite the promise of AI in solving  
 key healthcare challenges, several issues about the usage of  
 AI must be addressed. In this Section, we discuss some of the  
 practical AI tools in the clinics, as well as AI-based emerging  
 healthcare companies.

#### A. AutoML

Machine learning models have aided the healthcare industry  
 by lowering costs and improving outcomes, but only a small  
 number of hospitals are currently using them [4]. Healthcare  
 professionals likely lack the expertise to build, deploy and  
 integrate these models in clinical workflows. To assist the  
 deployment of machine learning models in daily work with  
 reduced input required from a data scientist or machine  
 learning engineer, AutoML [42], which automates machine  
 learning processes, has been developed. AutoML automates  
 fundamental steps like feature selection, model selection, and  
 hyper-parameter optimization, making it easier for health  
 professionals to develop machine learning models for clinical  
 data.

Generally speaking, about 80% of a data scientist's time  
 is spent on data preparation and feature engineering, which  
 also often requires domain knowledge experts [43]. The task  
 is to find the most discriminative features to provide insights  
 into the problem and to consider learning situations that  
 will be difficult for the classifiers. Several machine learning  
 frameworks have been developed to select, rank, and optimize  
 feature engineering processes [44].

A popular approach is expand-reduce, which applies transformation functions to obtain optimal features, and has been implemented in [45]. Genetic programming, based on the concept of natural evolution and a survival function, has been used for feature construction and selection.

Hyperparameters can also affect model performance, and optimizing them is an art that requires practical experience. Sometimes a brute force search is needed by a grid search with a manual specification of a subset of the hyper-parameter space. However, the dimensionality of the search space may make this impractical. Random searches, which sample hyperparameter configurations from a user-defined subset, can be limited to a specific computational budget. Another approach is a guided search that iteratively generates new configurations of the hyper-parameters based on the prior performance. AutoML automates this feature engineering and hyperparameter optimization and model selection process. Hence, non-technical professionals can use machine learning models to solve healthcare problems.

Auto-weka [45], another machine learning platform based on Bayesian optimization methods, can be used to optimize hyper-parameters and model selection [46]. Other practical products used are Google’s cloud AutoML system, Amazon’s Comprehend, and Microsoft’s Azure AutoML. The performance of AutoML models largely depends on the quality of the datasets. Adopting AutoML models in the healthcare environment will also require overcoming their operation as a black box.

### B. AI Tools and companies for clinics

The development and use of computer aided diagnosis or AI tools in clinical practice confront several hurdles despite the huge advancement in this new age of machine learning. For example, medical imaging is an essential diagnostic tool for various disorders. A variety of imaging modalities have been developed, with X-ray imaging, whole slide imaging, computed tomography (CT), ultrasound, magnetic resonance imaging (MRI), and positron emission tomography (PET) being some of the most widely utilized techniques. Moreover, several publicly available imaging and biological databases also offer excellent opportunities to build AI systems.

For example, PathAI [47] uses AI methods to assist pathologists in clinical diagnostics, clinical trials, and clinical translational research. Similarly, Viz.ai [48] is an AI-powered computer application to accelerate care coordination by reducing the time delays in clinical workflows. It uses AI to generate alerts and send them to clinicians for timely intervention. Similarly, Freenome [49] uses AI for cancer screening, diagnostics, prevention, and better management of cancer. Table II lists the companies that are completely based on AI tools to equip medical professionals to save lives.

1) *SaMD: Software as a Medical device*: SaMD [50] is meant to be used for one or more medical purposes and is not part of physical medical equipment. Since 1995, more than 500 software packages/applications have been approved by the FDA to assist doctors in various healthcare problems [51]. Most of these software packages are related to analyzing

TABLE I  
BROAD CATEGORIES AND APPLICATIONS OF AI IN HEALTHCARE INDUSTRY

Category	Specific Applications
Patient care	Diagnosis and Prognosis Real-time case prioritization Personalized medication Electronic health records, Smart health
Medical Imaging	Tumor segmentation and Detection Early diagnosis and Imaging Biomarkers Treatment effect monitoring
Management	Public Health Policy Market research Forecasting (Pandemics)
Biosensors	Remote health care Real-time health monitoring Soft computing
Computational Biology	Drug Discovery and efficacy analysis Single-cell analysis Multi-omics data analysis

radiology images. In many medical imaging tasks, AI algorithms have outperformed humans, and innovative companies have built AI-based systems to analyze radiology images and digital pathology slides. For example, Chan et al. [52] created a computer-aided diagnosis system to identify microcalcification on mammograms and carried out the first observer performance research that showed how well the developed tool improved breast radiologists’ ability to detect microcalcifications. Also see Table I.

AI researchers and developers must comprehend how clinicians desire to be assisted with different clinical works, construct efficient AI solutions, and produce interpretable results by considering the practical concerns in clinical settings. If properly created, verified, and applied, effective data analytics from AI technologies complement or support doctors’ intelligence to increase accuracy, workflow, and, ultimately, patient care.

## IV. APPLICATIONS OF BIG DATA ANALYTICS IN HEALTHCARE

The healthcare system consists of multiple stakeholders; patients, doctors, hospitals, industry, and policymakers, which are regulated by strict compliance. Healthcare systems generate a huge amount of data at a very high speed, which makes it a perfect avenue for big data analytics. Using big data analytics in healthcare may enable personalized medicine, timely interventions, better health policy management, and planning [64].

Big data analysis systems aim to collect, clean, extract, visualize, and analyze very large datasets and are associated with three key concepts. These are volume (large datasets), variety (highly dimensional/many attributes), and velocity (the speed at which the data is generated, made accessible, and analyzed). Healthcare datasets, usually large, complex, and arising from various sources, offer valuable opportunities for big data platforms [65]. For example, on average, a cancer patient generates 2GB of data annually in the form of images and medical records. New experimental techniques, such as immunotherapy, targeted therapy, omics research, high

TABLE II  
AI-BASED TOOLS AND COMPANIES IN THE FIELD OF MEDICAL SCIENCES.

Tool/Company	Services
Viz.ai [48]	It aims to reduce delays and make the healthcare team react faster with AI solutions regarding decision-making, treatment plans, and prescription providers.
PathAI [47]	It develops machine learning for pathologists to assist in diagnostics by reducing errors, specifically for cancer patients and personal treatment.
Buoy Health [53]	A chatbot attends to a patient and records the history, symptoms, and other health concerns; then guide the patient to the appropriate health facility. It is developed by a team at Harvard Medical School to speed up and optimize the treatment cycle.
Enlitic [54]	Enlitic creates deep learning radiology technologies. The company's deep learning engine analyses unstructured medical data to provide clinicians with improved insight into a patient's real-time demands.
Freenome [49]	It employs AI algorithms for cancer screenings, diagnostics, and blood work to identify cancer early and suggest innovative treatments.
Beth Israel Deaconess Medical Center [55]	It employs AI to diagnose blood disorders early. The robots were taught to detect germs using 25,000 blood sample photos. Machines learned to predict hazardous blood bacteria with 95% accuracy.
Iterative Scopes [56]	It uses AI for gastrointestinal diagnosis and therapy. They have submitted the first clinical study of their AI-powered SKOUT tool to the FDA for assessment.
VirtuSense [57]	It employs AI sensors to monitor patients' activities and alert them about accidents. VSTAlert can anticipate when a patient plans to get up and inform hospital services.
Caption Health [58]	It integrates AI and ultrasonography for illness detection. AI assists physicians through the scanning procedure in real time to collect early diagnosis results.
BioXcel Therapeutics [59]	It applies AI to develop immuno-oncology and neurological drugs. The company's medication initiative uses AI to uncover new uses for old pharmaceuticals.
BERG [60]	BERG is a clinical-stage, AI-powered biotechnology company taking a bold 'Back to Biology™' approach to healthcare.
Atomwise [61]	Atomwise utilizes AI to accelerate small molecule drug discovery and explores new undruggable targets to make them druggable.
XtalPi [62]	XtalPi's ID4 platform combines AI, the cloud, and quantum physics to anticipate small-molecule medicinal characteristics.
Deep Genomics [63]	Its AI platform finds neuromuscular and neurodegenerative medication possibilities. "Project Saturn" examines 69 billion cell molecules.

throughput screening, and parallel synthesis [66] may generate even larger amounts of data that require advanced data analytic methods.

In Figure 3, we show how complex high dimensional data from wearable sensors (ECG, Electromyograms (EMG), Electroencephalograms (EEG)), imaging data (X-rays, CT-Scans, MRI), electronic health records, and multi-omics (genome, proteome, and microbiome) data are generally collected and stored at a central repository, where pre-processing and data cleaning are performed. Missing values imputation methods may be used for further processing using statistical and machine learning methods. Centralized and mobile applications for patients, clinicians, hospitals, government agencies, and global health organizations can be developed. For example, the FDA has approved Ziopatch [67], which measures the heart rate and the ECG signal.

Multi-variate statistical methods, such as principal component analysis and other clustering methods, can be used to find patterns in a big dataset that may identify different disease states, mortality rates, susceptible age groups, forecast future

pandemics, and economic costs [68].

#### A. Multi-modal Data Fusion: A trash or a goldmine

Many quantities in the universe vary co-currently. Biological data is usually diverse, and a complete understanding of a complex biological system may require an ensemble of related data sets to extract hidden data dependencies [69]. However, combining these multi-modal data may result in a goldmine or trash. It requires domain knowledge and strong data engineering skills for efficient feature representation and any downstream analysis. For example, in [70] showed fusing histopathological, radiological, and clinicogenomics information improves risk stratification for cancer patients.

1) *Heterogenous Data*: The vast amounts of healthcare data generated daily, such as medical images, sensor data, medical histories, and genomic data, are heterogeneous. Machine learning is well suited to analyze multi-modal data and extract valuable insights.

Three major areas where multi-modal data fusion can be useful:

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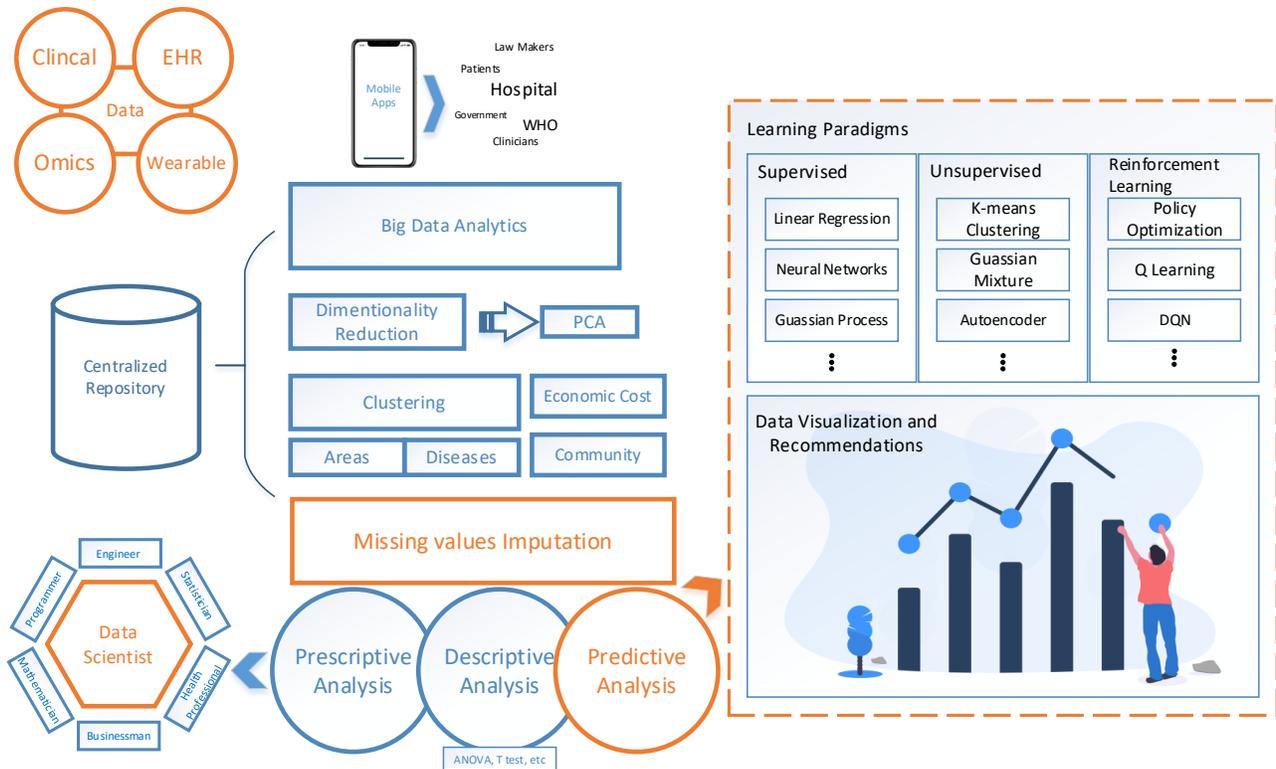


Fig. 3. Big data analytics in healthcare. Learning from various data modalities in the big data environment may aid patients, clinicians, hospitals, governments, and global health organizations. Different machine learning paradigms can be applied to analyze and visualize biomedical data.

- **Diagnosis:** Machine learning applied to health records and medical images can assist in the diagnosis of disease states.
- **Prognosis:** Applying machine learning algorithms to the heterogenous data available on a patient can predict the expected development of a disease from its early stages.
- **Treatment:** Optimal treatment plans can be generated by machine learning algorithms, especially reinforcement learning strategies, given the medical histories of patients and the number of treatment options available.

Medical data often consists of different data modalities such as images, signals, text, and molecular structures that are likely to be related. New machine learning or deep learning models enable us to integrate these diverse data sources, in a data-harmonization attitude [71] to produce multi-modal insights [72]. The extracted multi-modal features can also be used to form a knowledge graph to provide support for clinical decisions or understanding the mechanism of a specific disease [73] or visualisation for orthopaedic surgery [74]. In Figure 3, we show how multi-modal data can be used for different healthcare applications for patients, clinics, government and global healthcare organisation.

The integration of multiple data types may also increase the trust of clinicians. Since different data-modalities provide complementary information in describing a treatment plan or a disease process. In Figure 2, we show how different data-

modalities can be used for precision medicine. The main goal of methods used to combine multimodal data is to combine the data with values from various scales and distributions into a global feature space, where the data may be represented more consistently [75].

It is also pertinent to mention that in many real-world cases, fusing data from different data modalities may decrease the performance. The healthcare data are produced by extremely complex systems and instruments, including biological, environmental, social, and psychological ones, among others [76]. These systems are driven by a variety of underlying processes that are dependent on a wide range of variables, that may be not accessible in many cases [77]. In addition, the diversity among different data types; a number of samples, scales, and research questions further complicate the learning process. In small clinical cohorts, it may also suffer from the curse of dimensionality [78].

### B. Genomics Data Analysis

Genomic datasets, facilitated by next-generation sequencing, often contain vast amounts of raw data [79] and require big data analysis and computational methods. Examples are the encyclopedia of DNA elements (ENCODE) [80] gene annotation and expression data, the Cancer Therapeutics Response Portal (CTRP) [81], which can provide insights into the action of small molecules leading to personalized drug

519 discovery based on predictive biomarkers. The Cancer Cell  
520 Line Encyclopedia (CCLE) [82], and the Genomics of Drug  
521 Sensitivity in Cancer (GDSC) [83] database of large scale  
522 molecular screens on panels of hundreds of characterized  
523 cancer cell lines demonstrates the potential of modern machine  
524 learning algorithms to develop drug response predictors from  
525 molecular profiles.

526 However, current data resources are inadequate for reliable  
527 prediction of drug resistance or response [84]. Analyses of  
528 independent cohorts may reach different conclusions, and  
529 inconsistency between datasets and missing clinical informa-  
530 tion can hinder predictions. Data imputation techniques may  
531 address missing values, and the high dimensionality of the data  
532 could be dealt with by feature filtering techniques or sparse  
533 principal component analysis [85].

### 534 C. Medical Imaging

535 Deep learning can rapidly construct magnetic resonance  
536 (MRI) images directly from sensor data of partially observed  
537 measurements. Task-oriented reconstruction allows the recon-  
538 struction of a specific part of the image with high quality  
539 and a confidence score. Super-resolution images (high-quality  
540 images or sequences built from low-resolution images) can  
541 be constructed by deep learning, such as single (no reference  
542 information) brain MR images built using convolutional neural  
543 networks (CNNs) or super-resolution using GANs [86]. In  
544 Figure 4, we show various applications of deep learning in  
545 medical imaging.

546 For MRI images, image synthesis is a method to generate  
547 new parametric images or tissue contrasts from a collection of  
548 images acquired in the same session. Generative adversarial  
549 networks [87] could serve as a data augmentation tool as  
550 medical datasets tend to have limited numbers of samples,  
551 and they have been used to generate synthetic abnormal MRI  
552 images for a brain tumor based on pix2pix [88], [89].

553 Image registration, transforming data from multiple pho-  
554 tographs, different sensors, views, or depths to a single coordi-  
555 nate system is used, through deep learning, for medical image  
556 registration to improve accuracy and speed. Examples are  
557 deformable image registration, model-to-image registration,  
558 and unsupervised end-to-end for deformable registration of 2D  
559 CT/MR images [90].

## 560 V. WEARABLE BIOSENSORS

561 Wearable biosensors measure electro-physiological and  
562 electro-chemical signals from the body. Electrical activities  
563 emanating from various biological processes in the body, such  
564 as human heart activity (ECG), muscle activity (EMG), and  
565 sweat gland activity (Electro-Dermal Activity (EDA)) can  
566 be extracted from diagnostic machines or wearable sensors  
567 and provide vital information about one's health conditions.  
568 Analysis methods for these data, such as principal component  
569 analysis, discrete cosine transforms, auto-regressive methods,  
570 and wavelet transforms, can extract time and frequency domain  
571 features from the physiological signals [91]. Examples are a  
572 bidirectional deep long short-term memory (LSTM) network  
573 based on wavelet transform to classify ECG signals [92],

574 which achieved 99.39% accuracy on the MIT-BIH arrhythmia  
575 database [93] and a Fourier Transform and Wavelet-based  
576 feature model to classify patients with Alzheimer's Disease,  
577 Mild Cognitive Impairment and Healthy subjects from EEG  
578 signals [94].

### 579 A. AI-assisted design of biosensors

580 In the real world, medical signal data can also be passively  
581 gathered utilizing wearable sensors, such as smartphones or  
582 smartwatches [99]. The traditional way of acquiring signals  
583 has been through gel-electrodes that are placed on the body.  
584 In addition to the use of traditional wearables such as smart-  
585 watches and fitness trackers, recent advances in fabrication and  
586 electronics have led to the integration of bio-sensing electrodes  
587 in other devices such as eye-glasses [100], VR head-mounted  
588 displays [101], and textiles [96].

589 1) *Epidermal devices*: A new stream of computing devices  
590 termed *epidermal devices* allow for non-invasive capture of  
591 physiological signals through soft interactive tattoos [102],  
592 [103] (Figure 5). These epidermal devices can measure electro-  
593 physiological signals [96], [103] and electro-chemical signals  
594 in the body [104]. Another factor that has contributed to the  
595 widespread development of physiological sensing devices is  
596 the availability of open-source prototyping kits. Prototyping  
597 kits and platforms such as EMBody [105], Seeed<sup>1</sup>, OpenBCI<sup>2</sup>,  
598 Olimex<sup>3</sup>, BITalino<sup>4</sup> allow for rapid prototyping of custom  
599 physiological sensing systems. In addition to all these de-  
600 velopments, computational tools and AI-assisted approaches  
601 are being actively explored to automate and customize the  
602 design of biosensing wearables. For instance, Nittala et al. [97]  
603 developed a computational design tool built with an inte-  
604 grated predictive model to optimize the design of multi-modal  
605 electro-physiological sensing devices.

606 Machine Learning and Optimization Techniques for pro-  
607 cessing Physiological Signals

608 2) *Machine learning techniques on physiological signals*:  
609 Employing machine learning and deep learning techniques on  
610 physiological sensing is a commonly used approach. In the  
611 field of human-computer interaction, machine learning tech-  
612 niques have been commonly used for sensing gestures from  
613 EMG signals [106], identifying mood from EDA, Electroocu-  
614 lograms (EOG), EMG and ECG signals [101]. Deep learning  
615 approaches are also commonly applied on ECG data for de-  
616 noising data [107], for simulating signals and detecting heart-  
617 related anomalies [108], [109], emotion recognition [110] or  
618 to assess mental health by analyzing the EEG signals or to  
619 detect psychiatric disorders [111]. Classen et al. [112] detected  
620 brain activity using machine learning on the EEG recordings of  
621 brain-injured individuals who were clinically non-responsive,  
622 which is a predictor of eventual recovery.

## 623 VI. COMPUTATIONAL ADVANCES

624 Advances in computer hardware, and architectures are re-  
625 quired to process highly complex scientific problems. The

<sup>1</sup><https://www.seeedstudio.com/grove-emg-detector-p-1737.html>

<sup>2</sup><https://openbci.com/>

<sup>3</sup><https://www.olimex.com/Products/EEG/>

<sup>4</sup><https://www.pluxbiosignals.com/>

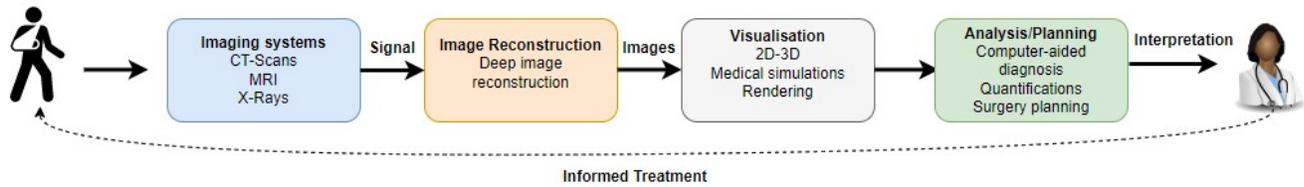


Fig. 4. Deep learning can be used to construct medical images at high speed, and facilitate the visualization and analysis of medical images.

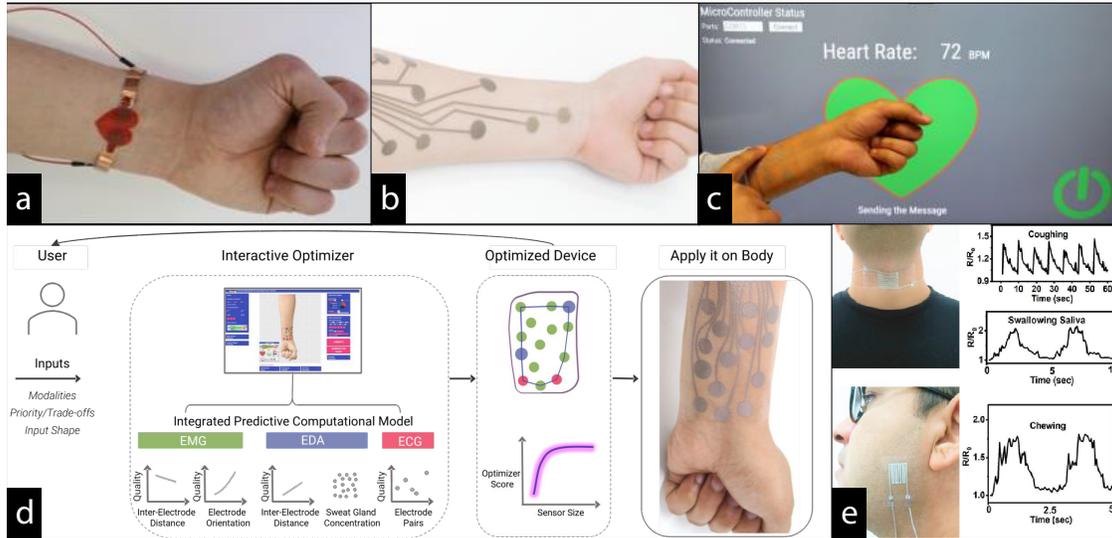


Fig. 5. Wearable Biosensors: (a) biosensors in a tattoo form factor that can sense electro-dermal activity (EDA) [95]. (b) multi-modal physiological sensing tattoo that can sense ECG, EDA, and EMG signals on the forearm [96]. (c) integration of user-interface controls e.g., touch buttons in bio-sensing tattoos [96]. (d) AI-assisted fabrication and optimization of multi-modal electro-physiological sensing devices [97]. (e) Ultra-thin and skin-conformable strain sensors on a decal transfer substrate, employed to detect subtle human body movements [98].

626 growth in fast processors, multicore-chips, accelerators, mem- 650  
 627 ory designs, interconnections, field programmable gate array 651  
 628 (FPGA) based processors, and GPUs with hundreds of cores 652  
 629 have made computationally intensive applications, such as 653  
 630 real-time image and video processing in healthcare, possible. 654

### 631 A. Accelerated Artificial Intelligence

632 Deep learning systems are often trained on multiple core 655  
 633 graphical processing units, which can optimize the highly 656  
 634 parallel matrix operations that are essential components of 657  
 635 deep neural networks. A recent example is the discovery 658  
 636 of faster matrix multiplication using reinforcement learn- 659  
 637 ing [113]. Google introduced a tensor processing unit (TPU) 660  
 638 as an accelerated artificial intelligence processor, especially for 661  
 639 its TensorFlow software [114].

640 Training of a deep neural network can be expedited by 663  
 641 either training more examples in parallel or training each 664  
 642 example faster. Operations that cannot be accelerated by 665  
 643 GPUs or TPUs, such as the earlier data processing stages or 666  
 644 input-output between devices or disks, need to be improved 667  
 645 in training. Data echoing [115], which reuses intermediate 668  
 646 outputs from earlier pipeline stages to reclaim idle capacity, 669  
 647 may be useful to ameliorate this. 670

648 As the quest to become a leader in AI continues, the model 671  
 649 sizes are increasing from millions of parameters to billions 672  
 673

of parameters (Openai GPT models). Google reported the 650  
 GLaM model with more than 1 trillion parameters (GPT-3 651  
 model had 175 billion parameters) [116]. The direct challenges 652  
 associated with these models are the training cost and the 653  
 porting out to small devices. One potential solution to enable 654  
 small models to learn the behavior of bigger models is to 655  
 use neural network compression techniques such as knowledge 656  
 distillation [117] or structural sparsity [118]. An analogy for 657  
 this is the teacher-student relationship, where the smaller 658  
 model (student) learns from the bigger model (teacher). A 659  
 survey in [119] presents efficient hardware architectures for 660  
 accelerating deep convolutional neural networks. 661

### 662 B. Edge Computing

663 Although most healthcare datasets are complex and large 663  
 664 and require massive computational resources (often in remote 664  
 665 computer clusters), processing data locally at the end nodes of 665  
 a cluster in a real-time application is appropriate for privacy 666  
 reasons or to reduce processing time and latency. The training 667  
 of the model locally on end nodes is known as edge computing. 668  
 In edge computing, edge (local) devices or servers can provide 669  
 data storage and processing, potentially giving fast, secure, 670  
 and real-time health analytics that may allow timely medical 671  
 interventions. Thus, an edge computing-based AI model could 672  
 provide better healthcare for patients far from major population 673

674 centers with limited connectivity and access. The localized  
675 processing power of edge computing may facilitate access to  
676 medical interventions by rapidly analyzing data from smart  
677 medical sensors.

678 To make AI models portable and compatible with prototyp-  
679 ing, the implementation of AI models on low-power devices  
680 is important. For example, Owais et al., [120] recently showed  
681 the implementation of the U-Net segmentation model on the  
682 Intel Neural Compute Stick. The work demonstrated that  
683 inference could be obtained on the NCS with proper tuning  
684 and suitable modifications of the U-Net model. However, the  
685 implementation was achieved with a trade-off for performance.  
686 Nevertheless, experimental results on brain MRI images and  
687 heart MRI images showed promising performance in terms  
688 of the dice scores for the segmentation tasks. Hence, such  
689 inference-enabled devices can aid in the clinical transforma-  
690 tions of AI methods in real-time healthcare settings.

### 691 C. Federated Learning

692 Data privacy and protection are general requirements for  
693 medical data, and new frameworks for training models are  
694 required that do not expose the underlying data. One such  
695 approach is Federated or Collaborative Learning [121], which  
696 is a machine learning technique that trains an algorithm across  
697 multiple edge devices or servers without exchanging local  
698 data samples. Multiple parties, for example, several hospi-  
699 tals/research centers, actively collaborate to train algorithms  
700 without centralizing their datasets. In developing AI models  
701 for medical data from multiple locations, federated learning  
702 has recently been shown to be effective. For example, with  
703 the rapid spread of COVID-19 globally, researchers needed  
704 to come up with quick responses and rapid developments  
705 of mechanisms for the assessment of COVID-19 patients.  
706 Multiple institutes around the globe collaborated to expedite  
707 AI model development for disease clinical support systems.  
708 However, sharing COVID-19 patient data from different lo-  
709 cations had ethical and legal bottlenecks that complicated the  
710 process. Hence, the research community resorted to federated  
711 learning to make use of data from diverse sites without the  
712 need for data sharing. In [122], a federated learning model was  
713 developed to predict future oxygen requirements for COVID-  
714 19 patients making use of clinical and radiology (chest X-rays)  
715 data. The model referred to as the EXAM model facilitated the  
716 use of data from 20 different institutes from various countries.

717 Federated Learning frameworks are implemented with dif-  
718 ferent topologies (also see Figure 6). To accomplish model  
719 training at multiple sites, the framework may execute model  
720 training at each site independently and then share the weights  
721 with other sites (a peer-peer topology), or the individual  
722 sites may share the weights with a centralized server node  
723 (client-server topology). According to the federated learning  
724 topology, the stochastic gradient descent (SGD) optimization  
725 of the model training is transformed into federated stochastic  
726 gradient descent (FedSGD) [123], [124].

## VII. THE RECENT CHALLENGES IN AI FOR HEALTHCARE WITH POTENTIAL SOLUTIONS

AI has shown great promise to improve the healthcare indus-  
try, and it is expanding as technology advances. However, there  
are some limitations in this field that prevent AI from being  
integrated into current healthcare systems. In this section, we  
discuss some of the key challenges and provide suggestions  
to overcome these to improve healthcare.

### A. Data issues

Data availability and access are two critical success factors  
for data science in healthcare. Moreover, the data quality,  
sample size, labels, disparity among labels, privacy, and ethical  
concerns, are the most prominent challenges that must be ad-  
dressed to fully exploit the potential of AI in healthcare [125].  
The first principle to build robust data-driven healthcare sys-  
tems is to capture clean, accurate, and properly formatted data  
for use in multiple healthcare applications. A perspective about  
sharing biomedical data for strengthening the role of AI is  
presented in [126].

Machine learning methods can also assist in automated la-  
beling, anomaly detection, missing value imputation, and other  
data cleaning processes [127]. For example, in [128], deep  
learning is used to identify bleeding events from electronic  
health records. Deep learning models are frequently used to  
improve the quality of radiology or pathology scans [129] or  
to identify anomalies in biosensors [130]. Some IT vendors  
also provide automated scrubbing tools that use logic rules to  
compare, contrast, and correct large datasets.

Another issue is the widespread perception in the com-  
munity that larger datasets are required to make accurate  
predictions. The data quality, proper annotations, and hypoth-  
esis in consultation with healthcare experts are necessary to  
build robust machine learning models. The data generated by  
the push of technology, without appropriate hypothesis and  
domain knowledge, will remain difficult to analyze.

Data security is another top priority for healthcare organi-  
zations. Risks include high-profile data breaches, hacking, and  
ransomware incidents [131]. Machine learning can be used to  
make data and systems more secure. It allows security systems  
to analyze and learn from patterns to help prevent similar  
attacks and respond to changing behavior.

To deal with imbalanced, complex, unlabeled, and poorly  
understood data, the type of learning paradigms and evaluation  
metrics used is also important. To address these challenges and  
generate hypotheses for understanding complex diseases and  
signaling pathway patterns, unsupervised or semi-supervised  
learning can be used [132].

1) *The challenges in distribution shifts and different data modalities:* Many real-world clinical AI systems suffer from the training and testing distribution shifts in the data. To deal with these distribution shifts, domain adaptation techniques are adopted in machine learning. In domain adaptation, we train a neural network on a source dataset  $X$  and achieve high accuracy on a target dataset  $Y$ , where  $X$  and  $Y$  have different data distributions.

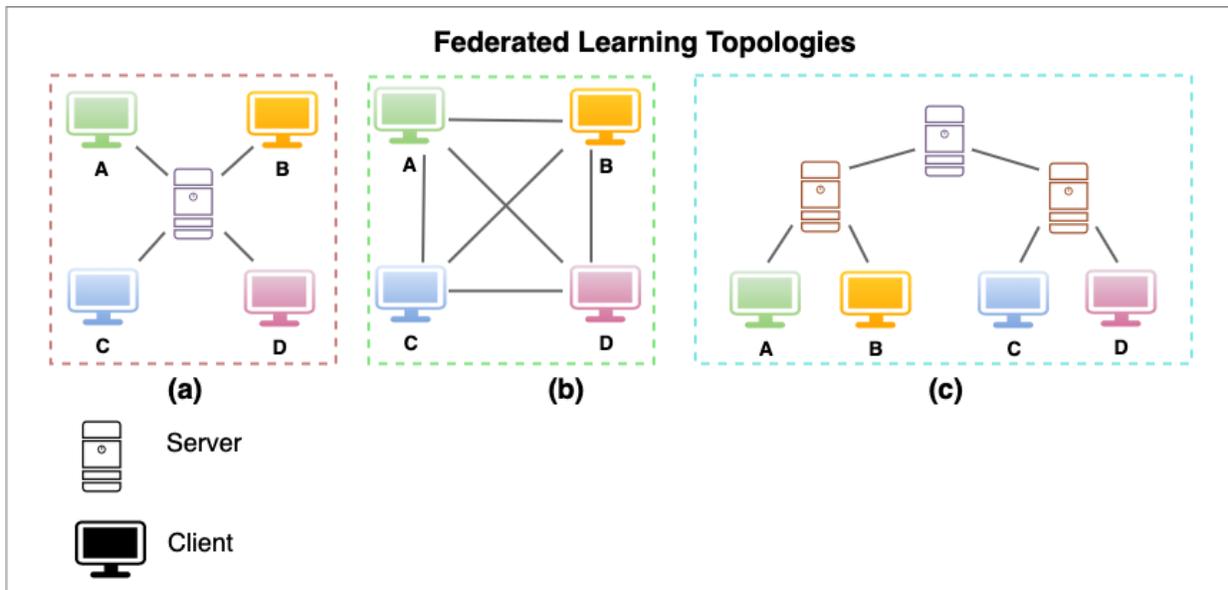


Fig. 6. Common topologies of federated learning. (a) Client-Server. (b) Client-Client. (c) Federation of sub-federation (mix topology).

782 Domain adaptation can be sliced down into three categories: 783 supervised, semi-supervised, and unsupervised learning, 784 depending on the type of data from the training dataset. In 785 supervised fast-expanding target dataset is substantially smaller 786 than the source dataset since the target domain's data has been 787 labeled. While unsupervised learning makes use of unlabelled 788 data from the target domain, semi-supervised learning uses 789 both labeled and unlabelled target domain data. As a result, 790 deep domain adaptation was suggested to improve the model's 791 performance and overcome the issue of insufficient labeled 792 data by utilizing deep network features. Discrepancy-based, 793 reconstruction-based, and adversarial-based adaptation are the 794 three main deep-domain adaptation strategies that have been 795 established.

796 In a discrepancy-based approach, the features that can be 797 transferred come up with drawbacks due to its delicate co- 798 adaptation and representation specificity. [133] has illustrated 799 that fine-tuning can improve generalization ability. When the 800 fine-tuning is conducted on the deep model, a base network 801 is trained using source data, and the first 'n' layers of the 802 target network are then used directly. The target network's 803 remaining layers are randomly initialized and trained using a 804 loss function based on the discrepancy. Finally, considering 805 the size of the target dataset and how closely it resembles the 806 source dataset, the initial layers can be fine-tuned or frozen 807 during the training procedure. Another deep domain adapta- 808 tion [134] technique, reconstruction-based domain adaptation, 809 uses an autoencoder to reduce reconstruction error and learn 810 transferable and domain-invariant representations to align the 811 discrepancy between domains.

812 Stacked Auto Encoders (SDAs) can be used to represent 813 source and target domain data in a high-level representation 814 manner [135]. However, because SDAs are computationally 815 expensive, the marginalized SDA (mSDA), which does not 816 require the use of stochastic gradient descent, was presented 817 in [136] to overcome the computational cost. Transfer learning

with deep autoencoders (TLDA) [137] used a softmax loss to 818 encode the source domain's label information. In contrast, the 819 embedding encoding layer uses the KL divergence to minimize 820 the distance in distributions between domains. 821

822 Generative Adversarial Networks (GANs) obtain transfer- 823 able and domain-invariant characteristics by minimizing the 824 distribution discrepancy between domains. GANs are also used 825 in the adversarial domain adaptation techniques [138]. CoGAN 826 was suggested in [139], which generated synthetic target data 827 and linked it with synthetic source data.

828 An approach for simulated-unsupervised learning was es- 829 tablished in [140], in which adversarial and self-regularisation 830 loss were minimized, using unlabelled real data to enhance 831 the realism of synthetic images.

832 2) *Challenges in Medical Imaging:* Perhaps, medical imag- 833 ing is the most disruptive area where AI has made tremendous 834 progress. However, there are various challenges in medical 835 imaging as well [141]. Medical images are often three- 836 dimensional, and the three-dimensional convolutional neural 837 networks to process these 3D volumes require more memory 838 and computational time. Generally, researchers treat 3D CNNs 839 as stacks of 2D CNNs. However, adding a newer dimension 840 adds additional constraints. Most deep learning models are 841 built on anonymized public data, making privacy-related issues 842 less relevant. However, this does not offer a permanent solution 843 to handle privacy-related problems in medical imaging. One 844 conclusion is that when these datasets are made public, there 845 are always associated risks of leaking patient privacy [142].

846 High diversity of clinical scenarios is another challenge in 847 medical imaging. This is because medical imaging can be used 848 in various clinical situations, such as disease detection, in- 849 cluding localization and classification and disease surveillance. 850 On the other hand, deep learning is also being used for data 851 quantification, such as pediatric bone age prediction [143]. As 852 a result, there are many different clinical activities from the 853 standpoint of medical imaging, and it is challenging for one

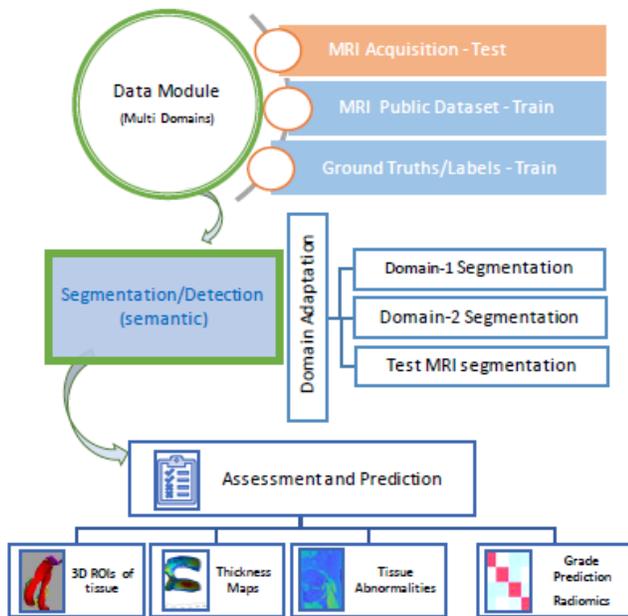


Fig. 7. Domain adaptation in medical imaging

854 individual or model to manage all of these operations using  
 855 present methodologies. Developing task-aware deep learning  
 856 solutions is the way forward.

857 Another significant challenge in medical imaging is the  
 858 lack of transparency in algorithms and issues with validation  
 859 and testing. AI-based applications differ in terms of data  
 860 ingestion to output, and there is currently no established  
 861 standard procedure. For example, algorithms with similar  
 862 performance may use different strategies to solve the same  
 863 problem, necessitating special pre-processing techniques be-  
 864 fore inference. As a result, scalability, which is critical in  
 865 commercial AI-based products, becomes difficult because each  
 866 application may require its own server or virtual environment.  
 867 The transferability of the algorithm presents another challenge  
 868 due to the stringent medical regulations in different nations.  
 869 However, there is no statistical method available to evaluate an  
 870 algorithm's transferability. One such initiative is the petabyte  
 871 'medical-imagenet' project of radiology and pathology images  
 872 by Stanford University with genomics and electronic health  
 873 record information for rapid creation of computer vision  
 874 systems(Stanford-AIMI).

875 The challenge of a lack of large datasets can be addressed by  
 876 image synthesis and data augmentation. Models may be hard  
 877 to generalize as the distribution of the training data, usually  
 878 high-quality images, may differ from real-world clinical data,  
 879 which may cause a deep learning model to produce unexpected  
 880 results. Transfer learning, fine-tuning, or pre-training can ad-  
 881 dress this [144]. Transfer learning leverages the weights of  
 882 a network already trained on a similar task. More emphasis  
 883 might be placed on unsupervised machine learning models  
 884 to overcome sample size issues. In Figure 7, we show the  
 885 applications of domain adaptation for image segmentation  
 886 tasks.

### 3) Biosensors and flexible bioelectronics: A way forward: 887

888 Despite increasing advancements in the last few years, there  
 889 are still numerous significant obstacles to overcome before  
 890 AI biosensors for Internet of Things-based applications are  
 891 commercially mature. For commercial applications, flexible  
 892 bioelectronic materials are a key component. The human body  
 893 and its internal organisms are naturally elastic and flexible.  
 894 In this instance, integrating electronics into platforms made  
 895 of flexible material is required. Current soft wearables on  
 896 the skin are dominantly reliant on capturing physiological  
 897 signals and transmitting those signals to an external computing  
 898 infrastructure (e.g. mobile, laptop, etc.). Flexible bioelectronics  
 899 is advantageous to match the human body and organs (such  
 900 as skin, eyes, and muscles) with low mechanical damage to  
 901 tissues and lessen adverse effects after long-term integration  
 902 because of its exceptionally flexible mechanical qualities.  
 903 Similarly, Medical AI biosensors will play a pivotal role in  
 904 developing key technologies in the future with the help of nan-  
 905 otechnology. They will continue to advance in miniaturization,  
 906 scalability, low power consumption, low cost, high sensitivity,  
 907 multifunction, safety, non-toxicity, and degradation [145].

908 4) *Adaptability*: Another issue is that the majority of ML-  
 909 enhanced biosensors currently lack adaptive learning capa-  
 910 bilities. Biosensors can learn from their surroundings with  
 911 adaptive learning rather than only depending on manually  
 912 input training sets. An adaptable model continually improves  
 913 and optimizes itself by learning from the environment, un-  
 914 like a non-adaptive system. This might lessen the chance  
 915 of disastrous mistakes and erroneous results, which a single  
 916 fixed model can cause. On the other hand, while non-adaptive  
 917 ML models' excellent local performance may be sacrificed in  
 918 the name of generalisability, particularly in clinical practice,  
 919 adaptive learning provides a solution to resolve this conflict.

920 5) *Bigdata in smart sensors*: Establishing a smart sensor  
 921 system that relies on enormous datasets and algorithms, is a  
 922 significant barrier regarding the platform for data processing  
 923 and storage. In recent years, cloud computing has been used to  
 924 process sensor signals since it offers superior computational  
 925 power and data storage. Cloud and biosensor integration is  
 926 nothing new, especially for monitoring applications where the  
 927 volume of data is continuously growing over time. The direct  
 928 connection of many sensors to the cloud is sometimes too  
 929 expensive and sluggish due to the exponential growth in the  
 930 number of sensors. Edge computing has so been introduced in  
 931 recent years. Instead of a single data centre, edge computing  
 932 enables data processing at scattered edge devices. It benefits  
 933 from great computational effectiveness, rapid network process-  
 934 ing, low cost, and more. Therefore, biosensors will likely use  
 935 this cutting-edge technology.

### B. Opening the black box of deep learning 936

937 A big hurdle in AI implementation is the black-box nature of  
 938 the deep learning models; in critical healthcare scenarios, we  
 939 can not fully rely on model predictions. We need interpretable  
 940 and transparent models to make critical healthcare decisions.  
 941 As the input data propagates through the layers of the neural  
 942 network, it gets compressed and generates some predictors for

943 the target label. Moreover, we do max-pooling at each layer  
 944 and drop out certain neurons in the final layers to avoid over-  
 945 fitting. Given these compressed representations, it is difficult  
 946 to explain the predictions at each level; however, we can have  
 947 a high-level idea about the inner-working of the model. Since  
 948 complex deep learning models consist of hundreds of millions  
 949 of parameters and, in our opinion, are nearly impossible to  
 950 interpret at every point.

951 In Figure 8, we show various methods used to explain the  
 952 working of the deep learning model. These methods can ex-  
 953 plain the predictions to a certain level without losing accuracy.  
 954 There is a trade-off between accuracy and explainable AI,  
 955 which depends on the problem at hand.

956 In a very intriguing study [146] proposed information  
 957 bottleneck [147] to explain the working of deep neural net-  
 958 works. The information bound is the theoretical limit proposed  
 959 by [147], at which the model can do the best given the set  
 960 of features; no further compression is possible. The paper  
 961 suggests that most of the training epochs are spent on learning  
 962 the efficient representations of the input; the representation  
 963 compression begins when training error starts to decrease. The  
 964 model starts to converge, layer by layer, and the last layer  
 965 keeps only the most relevant features to predict the output  
 966 label.

967 1) *Model fairness and accountability*: One of the chal-  
 968 lenges that the deployment of biosensors with AI will entail  
 969 is the need to ensure no biases in the outcomes determined.  
 970 Studies have shown [148], [149] that ML algorithms can  
 971 sometimes provide unequal outcomes for different population  
 972 groups, especially with populations already under-served in  
 973 society. In this regard, several steps need to be taken and  
 974 devised when working on ML applications using biosensors.  
 975 These can include actions such as a conscious inclusion  
 976 of diversity in the data collection process and developing  
 977 robust policies governing post-application performance audits  
 978 to quantify the impact on vulnerable communities. From a  
 979 technical perspective, aspects to look for would be logging  
 980 model performance to detect drift of performance in the  
 981 model. Such processes included in deploying and monitoring  
 982 biosensors utilizing AI applications would ensure healthcare  
 983 professional and patient confidence in the services offered.

### 984 C. Large Language Models for Healthcare

985 While the development of Large Language Models (LLMs)  
 986 has been the focus of researchers [150]–[152] for a while  
 987 relating to application towards machine translation, text sum-  
 988 marizing and paraphrasing and generation of text, the recent  
 989 release of ChatGPT [153] from OpenAI has brought the  
 990 potential use of chatbots into mainstream consumer use.  
 991 LLMs are deep learning models trained on a large amount  
 992 of textual data to cater to multiple tasks related to Natural  
 993 Language Processing. LLMs make use of complex transformer  
 994 architectures that enable it to capture longer dependencies  
 995 than is possible with typical sequential models such as RNNs.  
 996 LLMs also have the advantage of being able to be fine-tuned  
 997 for specific tasks, thereby performing well in some desired  
 998 niche or even work as the backbone for generic chatbots

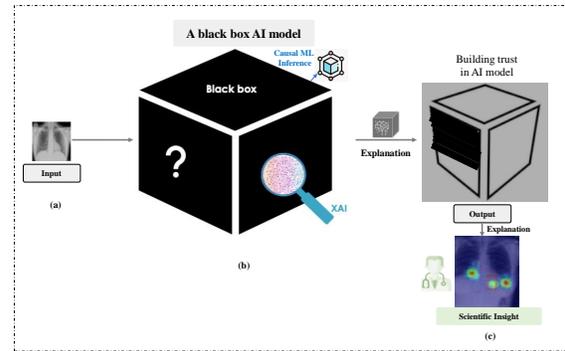


Fig. 8. AI- black-box model. Algorithms like Explainable AI, feature visualization or causal inference can be used to interpret the predictions. Gradcams visualization can highlight important regions that can build the trust of healthcare professionals.

999 too with a fine tuned performance. Infact, Open AI’s GPT-  
 1000 3 has been used as the back-end of several such offerings,  
 1001 including JasperChat (tailored for business use) and Poe by  
 1002 Quora, both of which are based on OpenAI’s base models.  
 1003 The multifaceted use of LLMs for special domains has also  
 1004 been true for the case of healthcare, medical data, as part of the  
 1005 used training data corpus enables chatbots powered by LLMs  
 1006 to be useful in assisting healthcare practitioners. One such way  
 1007 this was performed was suggested by Wang. et al. [154] who  
 1008 incorporate LLMs in to a CAD system for medical images  
 1009 called ChatCAD. They do this by generating prompts based  
 1010 on the output of different image based classifier/segmentor and  
 1011 report generator. These outputs are converted in to a prompt  
 1012 and are then passed on to the LLM so that its logical reasoning  
 1013 capabilities could be used to provide better and interactive care  
 1014 to patients. In order to provide a focused discussion on the  
 1015 potential use of LLM based chatbots for use in healthcare, we  
 1016 briefly discuss the current as well as potential uses of ChatGPT  
 1017 in this section.

1018 a) *ChatGPT for healthcare*: The OpenAI’s language  
 1019 chatbot ChatGPT [153] is an artificial intelligence language  
 1020 model that has been pre-trained on a large corpus of text data  
 1021 and is capable of generating human-like responses to natural  
 1022 language queries. Having passed successfully part of the US  
 1023 medical licensing exam, attesting to its capability to work with  
 1024 medical queries, ChatGPT has the potential to revolutionize  
 1025 clinical applications in many ways [155]. In Table III, we enlist  
 1026 several applications of ChatGPT.

## 1027 VIII. CONCLUSION AND FUTURE WORK

1028 The use of AI and biosensors has been gaining increasing  
 1029 traction in the healthcare industry for different purposes. AI-  
 1030 based methods are being embraced in the healthcare indus-  
 1031 try, where low-cost, intelligent, and adaptable methods are  
 1032 influencing fields such as clinical decision support, diagnos-  
 1033 tics, prevention, remote healthcare, public health policy, and  
 1034 clinical recommendation. More user-friendly machine learning  
 1035 technologies, such as AutoML, ClinicalAI, patient-centricAI,  
 1036 and explainable AI, are required to boost the confidence of  
 1037 healthcare stakeholders and to make machine learning an

TABLE III  
CHATGPT APPLICATIONS IN HEALTHCARE

Application	Description	Advantages	Disadvantages
Patient communication	ChatGPT can be used to communicate with patients and provide them with general medical advice. This can help reduce the workload on healthcare providers and improve patient satisfaction.	Provides immediate medical advice, available 24/7, can handle large volumes of inquiries simultaneously	May not be able to fully replace human interaction and empathy, may not be able to handle complex or critical cases, raises concerns about patient privacy and confidentiality.
Telemedicine	It can facilitate virtual consultations between patients and healthcare providers. By providing patients with access to medical advice and expertise, ChatGPT can help improve healthcare access and outcomes, particularly in rural or underserved areas.	Improves access to healthcare, reduces travel costs and wait times, increases patient engagement	May not be suitable for all types of medical consultations, may not be able to perform physical exams or provide hands-on care, raises concerns about patient privacy and security.
Medical education	Can be used as a tool for medical education, providing students and healthcare professionals with access to medical information and resources. By analyzing medical data and answering questions, It can improve medical knowledge and training.	Improves medical education accessibility, personalizes learning experience, can be used for quick reference and knowledge consolidation	May not be able to provide hands-on training, raises concerns about patient privacy and confidentiality, may perpetuate health disparities for students or institutions who do not have access to the technology or resources
Medical research	ChatGPT can be used in medical research to analyze large amounts of medical data and identify new patterns and trends.	Enables faster and more efficient analysis of large amounts of data, can identify previously unknown correlations and patterns	May require significant computing resources and expertise, may not be able to fully replace human researchers and medical experts.
Diagnosis support	It can assist healthcare providers in diagnosing diseases by analyzing patient symptoms, medical history, and other data.	Improves accuracy and consistency of diagnoses, saves time and reduces errors, can support rare and complex cases	May not be able to fully replace human diagnostic skills and expertise, and all clinical factors.

1038 integral part of daily clinical practice. Combining biosensors  
1039 and imaging data, or other data modalities, may increase the  
1040 model performance, as well as the confidence of clinicians.

1041 In this regard, this review provides researchers and health  
1042 practitioners with an overview of the state of technology  
1043 in this area, both from a technical and clinical perspective.  
1044 Various applications of AI towards diagnosis, prognosis, treat-  
1045 ment as well as monitoring have been discussed, along with  
1046 traits related to explainability and the tools useful in clinical  
1047 practice. Moreover, technologies that enable the usage and  
1048 development of biosensors for healthcare applications have  
1049 been presented. Lastly, open research issues and challenges  
1050 related to biosensor-based healthcare systems have been talked  
1051 about, which require further work.

1052 AI has great potential to transform the healthcare systems  
1053 and improve the lives of patients and health professionals.  
1054 However, clinical AI implementation is currently on a smaller  
1055 scale due to trustworthiness, lack of coordination, data col-  
1056 lection and privacy issues, and patient reluctance. We need to  
1057 develop patient-centric AI systems and build the trust of health  
1058 professionals in this exciting technology. AI can only assist  
1059 health professionals and improve lives, and in no way can it  
1060 replace them, of-course nobody would like to be treated with  
1061 a robot. AI, in any sense, can not replace the human touch,  
1062 which is the essence of every field. AI and clinicians should  
1063 work in synergy to maximize the benefits for patients. In this  
1064 regard, this article will guide further research and development  
1065 in AI for healthcare. Given the enormous amount of data and  
1066 processing power available today, we expect an increasing role  
1067 of AI and biosensors in the clinics that will augment or help  
1068 healthcare professionals and reduce their workload.

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