

REVIEW

Artificial intelligence for HIV care: a global systematic review of current studies and emerging trends

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Abstract

Introduction: Artificial intelligence (AI) and, in particular, machine learning (ML) have emerged as transformative tools in HIV care, driving advancements in diagnostics, treatment monitoring and patient management. The present review aimed to systematically identify, map and synthesize studies on the use of AI methods across the HIV care continuum, including applications in HIV testing and linkage to care, treatment monitoring, retention in care, and management of clinical and immunological outcomes.

Methods: A comprehensive literature search was conducted across databases, including PubMed and ProQuest Central, Scopus and Web of Science, covering studies published between 2014 and 2024. The review followed PRISMA guidelines, screening 3185 records, of which 47 studies were included in the final analysis.

Results: Forty-seven studies were grouped into four thematic areas: (1) HIV testing, AI models improved diagnostic accuracy, with ML achieving up to 100% sensitivity and 98.8% specificity in self-testing and outperforming human interpretation of rapid tests; (2) Retention in care and virological response, ML predicted clinic attendance, viral suppression and virological failure (72–97% accuracy; area under the curve up to 0.76), enabling early identification of high-risk patients; (3) Clinical and immunological outcomes, AI predicted disease progression, immune recovery, comorbidities and HIV complications, achieving up to 97% CD4 status accuracy and outperforming clinicians in tuberculosis diagnosis; (4) Testing and treatment support, AI chatbots improved self-testing uptake, linkage to care and adherence support. Methods included random forests, neural networks, support vector machines, deep learning and many others.

Discussion: AI has the potential to transform HIV care by improving early diagnosis, treatment adherence and retention in care. However, challenges such as data quality, infrastructure limitations and ethical considerations must be addressed to ensure successful implementation.

Conclusions: AI has demonstrated immense potential to address gaps in HIV care, improving diagnostic accuracy, enhancing retention strategies and supporting effective treatment monitoring. These advancements contribute towards achieving the UNAIDS 95-95-95 targets. However, challenges such as data quality and integration into healthcare systems remain. Future research should prioritize scalable AI solutions tailored to high-burden, resource-limited settings to maximize their impact on global HIV care.

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Keywords: artificial intelligence; chatbots; diagnostics; HIV care; machine learning; retention to care; treatment monitoring; viral suppression

Additional information may be found under the Supporting Information tab of this article.

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1 | INTRODUCTION

The HIV pandemic has had devastating consequences. Since its onset, an estimated 85.6 million people have acquired HIV,

and 40.4 million people have died from AIDS-related causes. In 2022, approximately 39 million people globally were living with HIV. In the same year, approximately 1.3 million individuals newly acquired HIV, while 630,000 lost their lives to AIDS-

related illnesses [1]. Notably, nearly 29.8 million people had access to antiretroviral therapy (ART) [1], and UNAIDS estimates revealed considerable progress towards the 95-95-95 targets. The achievement of 86%-89%-93% (86% of people living with HIV [PLWH] knowing their status, 89% of those with a known status on treatment and 93% of those on treatment achieving viral suppression) is a significant improvement from the 71%-67%-83% estimates of 2015 [1]. Much work, however, still needs to be done: at the end of 2023, \pm 5.5 million PLWH were not yet aware of their status; \pm 9 million were not on ART; and \pm 11.3 million were not virally suppressed. To attain the global HIV targets, innovative strategies will be needed.

Artificial intelligence (AI) is at the forefront of global innovation, with its potential to enhance health outcomes gaining increasing recognition. AI refers to the capacity of machines, such as computers, to replicate aspects of human intelligence [2]. Machine learning (ML), a subset of AI, entails the utilization of algorithms and statistical models programmed to learn from data, thereby discerning patterns and executing tasks without explicit human instructions [2]. Supervised learning, a type of ML, learns to map inputs to outputs through pre-matched (labelled) pairs, contrasting with unsupervised learning, which identifies patterns in data without prior labelling [2]. Neural networks, inspired by biological systems, employ interconnected neurons with adjustable weights to make predictions, while deep learning refers to neural networks with multiple hidden layers of neurons [2]. These concepts collectively underpin the potential of AI to revolutionize various fields, including healthcare, by enhancing decision-making processes and ultimately improving outcomes [2].

AI's integration into clinical HIV care offers promise for the improvement of disease diagnosis, treatment selection and risk assessment. This has the potential to enable healthcare systems to more accurately identify PLWH, initiate ART in a timely manner and implement targeted retention strategies, ultimately improving patient and public health outcomes. While systematic reviews have been conducted on the role of AI in managing conditions like hypertension and diabetes, there does not appear to have been an exploration of the potential applications of AI in various aspects of HIV care [3-6]. This study aimed to systematically identify, map and synthesize studies on the use of AI methods across the HIV care continuum, including applications in HIV testing and linkage to care, treatment monitoring, retention in care, and management of clinical and immunological outcomes.

2 | METHODS

This systematic review followed the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) guidelines [7]. The systematic review was registered with PROSPERO to ensure transparency and adherence to the predefined methodology and reporting standards (https://www.crd.york.ac.uk/PROSPERO/display_record.php?RecordID=517798).

2.1 | Search strategy

The search strategy was developed by a librarian, author LM, in consultation with SN. This search strategy was designed to gather relevant articles on HIV care and the application of AI in healthcare, spanning the period from 2014 to 2024. It included queries in four key databases: PubMed, Scopus, Web of Science and ProQuest Central. The search terms focused on HIV-related care and treatment, including HIV management, ART and HIV/AIDS interventions, paired with terms related to AI and healthcare technologies such as ML, automated systems and digital health. Filters for article type, language (English) and time span (2014-2024) were applied to ensure the results were up to date and relevant. The search results from these databases included 265 articles from PubMed, 355 from Scopus, 314 from Web of Science and 2799 from ProQuest Central. The detailed search strategy is provided in Supplementary Material 1.

2.2 | Screening

In accordance with PRISMA guidelines, a total of 3733 records were identified through database searching. After duplicates were removed, 3185 records remained for screening eligibility. The title and abstract screening phase was conducted by two authors, SN and TB, who reviewed the 3185 remaining records based on predefined inclusion and exclusion criteria. Of these, 3060 records were excluded based on predefined criteria. Subsequently, 125 full-text articles were assessed for eligibility, of which 74 were excluded for various reasons such as irrelevant study designs, populations that did not focus on PLWH or interventions that were not related to AI in HIV care. Ultimately, 47 studies were included in the narrative synthesis for analysis (Figure 1).

2.3 | Study selection criteria

This systematic review included observational studies, experimental trials and studies that investigated AI applications in HIV care. We defined HIV care to encompass clinical activities and outcomes following HIV diagnosis, including ART initiation, adherence and retention, viral suppression, immunologic monitoring (e.g. CD4/CD8 ratios), management of comorbidities, prediction of treatment-related outcomes (e.g. frailty, mortality, coinfections) and clinical decision support for ongoing care. While our search strategy primarily targeted HIV care and management terms, we also included studies on AI interventions that facilitated entry into care through mechanisms such as HIV self-testing promotion that was linked to clinical follow-up or ongoing care engagement (e.g. AI chatbots promoting self-testing with linkage to services). The inclusion criteria were established to capture a comprehensive range of AI applications in HIV diagnosis, treatment monitoring and patient management. Studies were excluded if they were published in languages other than English, were conducted before 2014 or focused on AI applications without a direct connection to HIV care. Additionally, studies that applied AI techniques but did not report outcomes related to HIV care were excluded to ensure relevance.

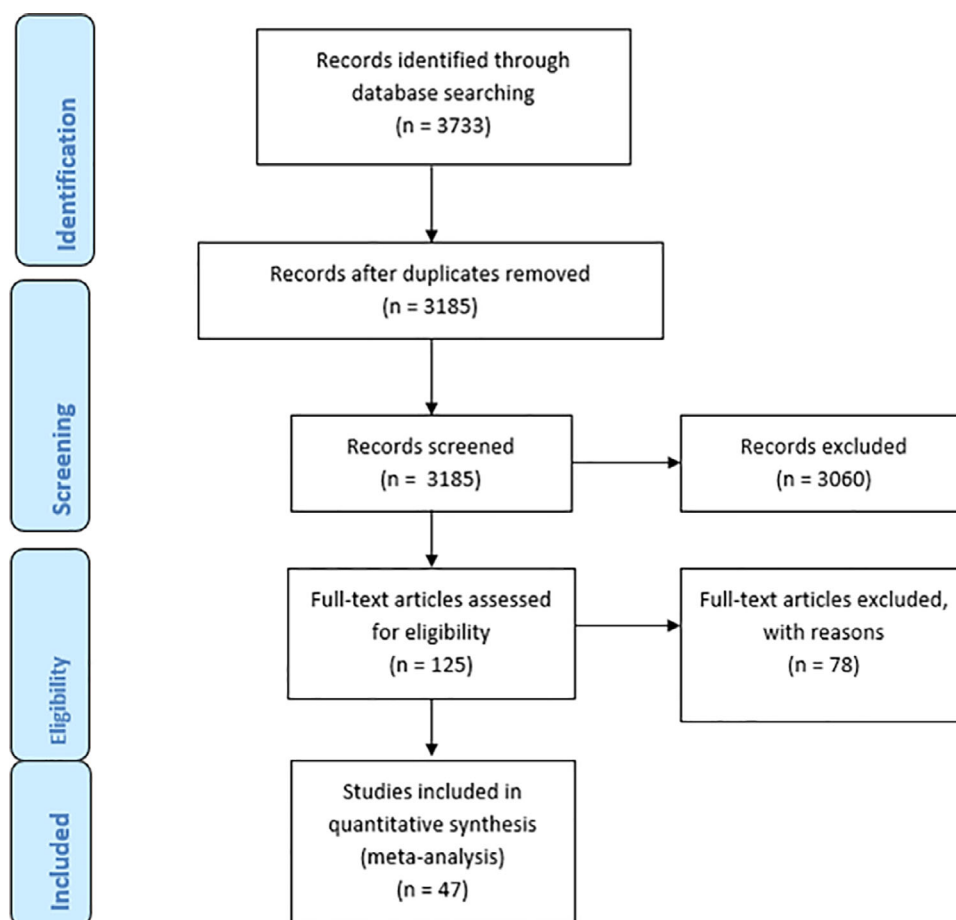


Figure 1. PRISMA 2020 flow diagram of study selection.

2.4 | Data collection process

The data extraction process followed a structured approach to maintain accuracy and consistency. SN and TS independently extracted data, and the compiled dataset was subsequently reviewed and refined by author ML, who acted as the third reviewer. Given the complexity of ML methodologies, JS, a mathematician specializing in AI, conducted an independent review of the extracted ML techniques to ensure precise classification and correct interpretation of algorithmic approaches. To enhance consistency, the research team conducted virtual consensus meetings, where reviewers discussed discrepancies and reached a final agreement on study inclusion and data categorization. Data extraction adhered to PRISMA guidelines and was systematically recorded in Microsoft Excel. The extracted variables included: Study ID (Article Title), File Name, ML Techniques Used, Study Design, Study Setting, Study Population/Participants, Sample Size, Intervention/Exposure, Comparator/Control, Outcomes of Interest, Effect Size/Measure of Association, Statistical Methods, Key Findings/Results and Conclusion/Summary. This structured approach facilitated a comprehensive synthesis of the data while ensuring consistency across studies. ML techniques were categorized using standard taxonomies found in authoritative sources such as Scikit-learn, The Elements of

Statistical Learning and Topol's framework for AI in health-care.

2.5 | Quality appraisal

The methodological rigour of the included studies was assessed using the Critical Appraisal Skills Programme (CASP) checklist (Supplementary Material 2) [8]. TS evaluated the studies, focusing on key aspects such as study design, risk of bias, validity and generalizability. Any discrepancies in quality assessment were discussed and resolved through consensus to maintain the integrity of the review.

2.6 | Synthesis strategy

The synthesis of findings was structured around four key thematic areas that reflected the different applications of AI in HIV care. Studies were categorized into: (1) HIV testing; (2) Retention in care and virological treatment response; (3) Clinical and immunological treatment outcomes; and (4) Testing and treatment support. In parallel, ML techniques were analysed based on the ML methods employed, such as random forests (RFs), logistic regression (LR), support vector machines (SVMs) and neural networks, to map their usage in various aspects of HIV care.

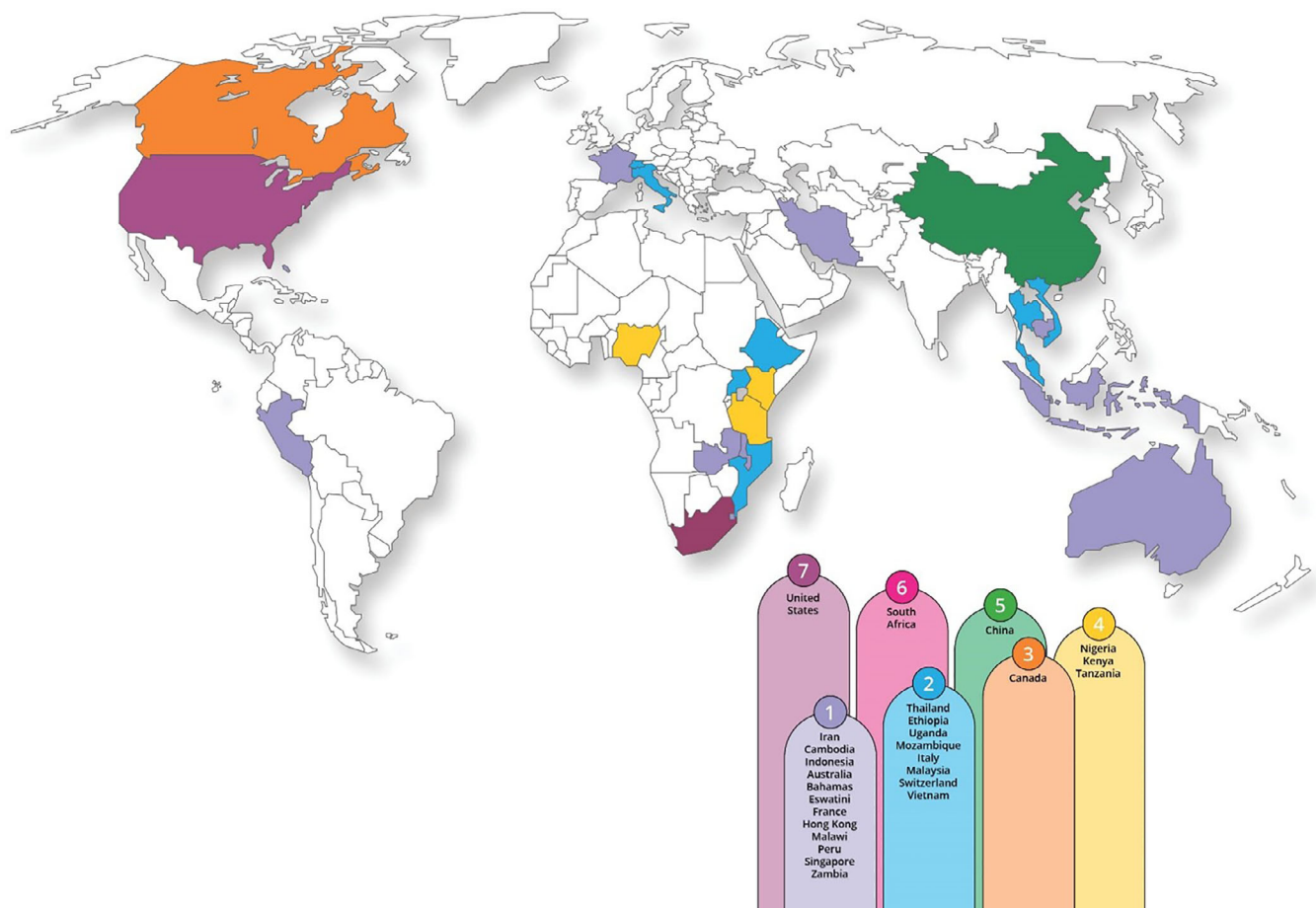


Figure 2. Geographic coverage of included studies on AI in HIV care.

3 | RESULTS

Table 1 summarizes the 47 included studies, detailing authors, study design, setting, population, sample size, outcomes and AI methods used. Figure 2 presents the country distribution of included studies, with the highest representation from the United States ($n = 7$), South Africa ($n = 6$), China ($n = 5$), Canada ($n = 3$), and Kenya, Nigeria and Tanzania ($n = 4$). Studies were conducted across both high- and low- to middle-income countries, reflecting the global interest in applying AI methods across the HIV care continuum. Study designs ranged from retrospective and prospective cohorts to randomized controlled trials and cross-sectional analyses.*

The results are structured into four thematic areas: (1) HIV testing; (2) Retention in care and virological treatment response; (3) Clinical and immunological treatment outcomes; and (4) Testing and treatment support.

3.1 | HIV testing

ML models have been employed to predict HIV and other sexually transmitted infection testing uptake [48], while predictive models have been applied to classify field-based rapid

diagnostic test images for HIV diagnosis [47]. The application of ML algorithms enhanced the specificity and positive predictive value in interpreting HIV rapid diagnostic tests, significantly reducing false positives compared to traditional visual methods [50]. AI-assisted HIV self-testing in Kenya demonstrated perfect sensitivity (100%) and high specificity of 98.8%, ensuring accurate interpretations and enhancing service delivery in pharmacy-based HIV testing [56]. In rural South Africa, ML algorithms trained on 11,374 HIV rapid test images achieved 97.8% sensitivity and 100% specificity, outperforming human interpretation and supporting REASSURED mobile diagnostics [47].

3.2 | Retention in care and virological treatment response

Predictive AI has been applied across multiple domains in HIV care, including retention in care [9, 15, 18, 22], viral load suppression [16, 18, 20, 57] and virological failure [14, 21]. ML models have been employed to forecast missed ART appointments [34, 58] and predict clinic attendance [22], supporting adherence and retention strategies. ML models for retention in HIV care showed strong predictive performance across diverse contexts. Models achieved an area under the curve (AUC) of 0.69 for predicting clinic visit attendance in South

* Numbers represent the count of included studies conducted in each country.

Table 1. Details of the studies included in the review

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[9]	Jeni et al. (2022)	Retrospective study	Nigeria and Mozambique	PLWH	360	Logistic regression, ANN, random forest, XGBoost, extra trees	Random forest, boosted trees predicted ART loss to follow-up with AUPRC = 0.65 in Mozambique and 0.52 in Nigeria, outperforming standard methods and supporting targeted interventions across sex and age groups.
[10]	Tu et al. (2021)	Longitudinal study	Canada	PLWH; three patient groups: peripheral neuropathy (n = 111)—included HIV-associated distal sensory polyneuropathy (n = 90); mononeuropathy (n = 21); and non-neuropathy (n = 408)	540	Random forest, logistic regression	Random forest models (AUC > 0.80) outperformed logistic regression in predicting peripheral neuropathy, with HIV-1 duration, peak viral load, age and low CD4+ T-cell count as top predictors.
[11]	Roy et al. (2022)	Cohort study	Canada	PLWH	186	Random forest, logistic regression	Random forest models (AUC = 0.76–0.79) outperformed logistic regression (p = 0.0001) in predicting carotid plaques using age, smoking, axial strain, pulse pressure and HIV status. Random forest achieved 82.8% accuracy, 78% sensitivity and AUC = 0.73 in predicting 1-year mortality or AIDS progression, outperforming six other models including logistic regression.
[12]	Dominguez-Rodriguez et al. (2022)	Prospective study	South Africa and Mozambique	Children with perinatally acquired HIV who began ART <3 months of age	100	Random forest, SVM, k-nearest neighbours, naïve Bayes, ANN, elastic net, logistic regression	Gradient boosting, random forest and support vector regression predicted >5% weight gain with >90% accuracy when body composition data (DEXA) were included; simplified models were sufficient to rule out gainers.
[13]	Federico et al. (2023)	Observational study	Italy	PLWH	3321	Gradient boosting, SHAP (explainability)	

(Continued)

Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[14]	Esber et al. (2023)	Cohort study	Sub-Saharan Africa: Uganda, Kenya, Tanzania and Nigeria	PLWH	2941	LASSO regression, random forest	LASSO regression (AUC = 0.82) and random forest (AUC = 0.75) predicted viral failure using 94 variables; key predictors included CD4 count, ART regimen, adherence, age and duration on ART. An ensemble decision tree model using routine EMR data predicted 6-month disengagement from care (≥28-day no-show) with 75.2% accuracy and 54.7% sensitivity among the top 30% highest risk group.
[15]	Fahey et al. (2022)	Cohort study	Tanzania	PLWH	178	Ensemble decision trees	Using 4 million VL records from 4265 Kenyan health facilities, a random forest model predicted viral load hotspots (≥20% unsuppressed) with 78% accuracy, F1 score = 69% and Brier score = 0.139, correctly identifying 434 of 446 hotspots.
[16]	Kagendi and Mwau (2023)	Cross-sectional study	Kenya	Pseudonymized participants	4 million tests and 4265 facilities	Random forest	A random forest model predicted virological failure with AUC = 0.9989, sensitivity = 1.00 and F1-score = 0.993; top predictors included low CD4 count, ART duration, TDF-3TC-EFV regimen and lack of CPT/TPT use.
[17]	Mamo et al. (2023)	Cross-sectional study	Ethiopia	PLWH	5264	Random forest, association rule mining	

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[18]	Maskew et al. (2022)	Longitudinal study	South Africa	PLWH	445,636 patients (retention model) 363,977 (viral load model)	Logistic regression, random forest, AdaBoost	Random forest, logistic regression and AdaBoost predicted next-visit attendance (AUC = 0.69) and viral load suppression (AUC = 0.76), with key predictors including prior late visits, treatment duration and number of prior VL tests. Using pseudo amino acid composition features, MLP and logistic model tree algorithms predicted anti-HIV-1 peptides with 96.15% and 83.71% accuracy, respectively, making MLP the most accurate classifier for sequence-based peptide prediction.
[19]	Poorinmohammad and Mohabatkar (2015)	Descriptive study	Iran	Three datasets: positive dataset, experimentally tested peptides and independent test sets	230, 212 and 47	ANN, logistic model tree, random forest, K star, J48 decision tree	XGB predicted viral load >1000 copies/ml with 96% accuracy, 97% sensitivity and AUC = 0.99, while GB achieved the highest accuracy for CD4 <200 classification; RF ranked second in viral load prediction.
[20]	Seboka et al. (2023)	Retrospective study	Ethiopia	PLWH	2907	K-nearest neighbours (KNN) Support vector machine (SVM) Logistic regression (LR) Decision tree (DT) Gaussian naive Bayes (GNB) Random forest (RF) Gradient boosting (GB) eXtreme gradient boosting (XGB)	

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[21]	Steiner et al. (2020)	Retrospective study	USA	HIV-1 sequence data and drug resistance assay	No study population. 18 ART drugs	Multilayer perceptron (MLP), bidirectional recurrent neural network (BRNN), convolutional neural network (CNN)	CNN achieved the highest drug resistance classification accuracy (mean AUC = 0.95), outperforming MLP (AUC = 0.91) and BRNNs (AUC = 0.93), with model interpretability confirming importance of key drug resistance mutations.
[22]	Xianglong et al. (2022)	Retrospective study	Australia	Men who have sex with men	1627	Logistic regression, LASSO, ridge regression, elastic net, SVM, k-nearest neighbour, naïve Bayes, random forest, XGBoost, MLP	XGBoost predicted timely post-reminder clinic attendance (AUC = 62.8%, F1 = 70.8%), while elastic net regression best predicted HIV/STI testing uptake within 30 days (AUC = 82.7%, F1 = 85.3%).
[23]	Peng et al. (2022)	Qualitative study	Malaysia	Men who have sex with men in Malaysia	Five web-based focus group interviews; 31 MSM	NLP	Participants expressed interest in AI chatbots for HIV prevention if anonymity, accurate information, linkage to services and stigma-sensitive design were ensured; concerns included privacy, usability and legal risks.
[24]	Murnane et al. (2021)	Retrospective study	Sub-Saharan Africa	Pregnant women (2011–2014)	1321	LASSO, logistic regression, SuperLearner, random forests, gradient boosting	SuperLearner and LASSO models using routine clinical data predicted viraemia (>50–1000 copies/ml) with moderate accuracy (AUC = 0.74–0.78), but flagged 64% of women as high risk to achieve 90% sensitivity—limiting utility for targeted interventions.

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[25]	MacPherson et al. (2021)	Randomized controlled trial	Malawi	Adults (≥18 years) with cough attending acute primary services in Malawi	473, 492 and 497	ANN	HIV-TB screening with computer-aided chest X-ray (CAD4TBv5) plus universal HIV testing led to faster TB treatment initiation (median = 1 vs. 11 days in SOC) and 40% same-day TB treatment versus 0% in SOC; undiagnosed HIV dropped from 2.7% to 0.2% (RR = 0.09). ChatGPT provided accurate and comprehensive responses to ART-related queries, demonstrating utility as a supportive tool for HIV counselling, though its advice remained generic and required clinical contextualization.
[26]	Koh et al. (2024)	Qualitative evaluation of chatbot responses	Singapore	Simulated patient queries	23 prompts	NLP	ML models (SVM, RF, MLP) predicted CD4/CD8 ratio from clinical data; SVM performed best for CD4<200 ($R^2 = 0.365$), RF best for CD4≥200 ($R^2 = 0.341$). RF and LR models were applied to immunologic and reservoir data from 115 ART-treated individuals; prediction of high versus low total or intact HIV DNA yielded ~70% balanced accuracy across models. Notably, CD127 expression on CD4+ T cells and years on ART were strong features in classifying reservoir level.
[27]	Olatosi et al. (2021)	Retrospective study	USA	PLWH	8888	Bayesian network, random forest, decision trees, ANN	
[28]	Semenova et al. (2023)	Cross-sectional study	USA	PLWH	115	Linear regression, RF, PaCMAP (pairwise controlled manifold approximation projection) GOSDT (global optimal sparse decision trees), LOCO (leave-one-covariate-out) Inference	

(Continued)

Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[29]	Aybar-Flores et al. (2023)	Retrospective study	Peru	Adolescents and young people	10,565	Random forest, quasi-binomial logistic regression, k-nearest neighbours, decision tree, artificial neural network	Random forest performed best (accuracy = 64.3%) to predict HIV/AIDS knowledge, identifying 14 significant predictors: gender, area of residence, wealth index, region of residence, age, highest educational level, ethnic self-perception, having heard about HIV/AIDS, HIV/AIDS screening test history, mass media access, marital status, working status, health insurance coverage and internet use.
[30]	Bala et al. (2023)	Retrospective study	Nigeria	PLWH	2500	ANN, ANFIS, SVM, multiple linear regression (MLR)	ANFIS outperformed all models in predicting ART outcomes, with $R^2 = 0.903$ (training), 0.904 (testing) and MSE = 7.961 (training), 3.751 (testing); CA showed the strongest association between ART drugs and marital status (explaining 93.7% of variation).
[31]	Bisaso et al. (2018)	Observational study	Uganda	PLWH	EFV cohort = 233; IDI cohort = 484	Multitask temporal logistic regression (MTLR), patient-specific survival prediction (PSSP), simple logistic regression (SLR)	MTLR outperformed PSSP and SLR in predicting early virological suppression (AUROC = 0.92 vs. 0.75 vs. 0.53; Brier = 0.08 vs. 0.19 vs. 0.11); external validation showed 92.9% accuracy, AUROC = 0.878 and false positive rate = 6.9%.

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[32]	Chikusi et al. (2022)	Observational study	Tanzania	HIV index clients and their notified contacts	6346 HIV index clients and 7226 contacts	Random forest, XGBoost, artificial neural network (ANN)	RF performed best (MAE = 1.1261) in predicting HIV index testing outcomes compared to XGBoost (MAE = 1.2340) and ANN (MAE = 1.1268); model visualizations identified Kilimanjaro as having the highest partner-notified cases and 82.6% of notified contacts were female.
[33]	Cotiugno et al. (2020)	Cohort study	Italy	PLWH	23	Elastic net (glmnet function in R) [20], support vector machines (svm.fs function in R) [21] and random forests	An AdaBoost model using expression profiles from 46 genes across five lymphocyte subsets and conditions predicted vaccine immunogenicity with 95.6% accuracy and only one misclassified case, demonstrating high potential to identify non-responders to influenza vaccination among children living with HIV.
[34]	Rachel et al. (2023)	Cross-sectional study	South Africa	Patients from public health clinics	202,817	Categorical boosting and ADABOOST	The categorical boosting model achieved 62% sensitivity, 67% specificity and 20% PPV, predicting 22,119 of 35,985 treatment interruptions; performance was highest during the first 6 months of ART.

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[35]	Li et al. (2022)	Retrospective study	China	PLWH	96	SVM, random forest, ANN	SVM, RF and MLP models using clinical indicators effectively predicted immune function (CD4/CD8 ratio) in long-term ART patients; SVM outperformed in low-CD4 (<200) cases, RF performed best in higher CD4 (≥200) cases.
[36]	Li and Li (2020)	Qualitative study	China	Morbidity models	NA	ANN, ARIMA (auto-regressive integrated moving average model)	The BP-ANN model achieved a mean absolute percentage error (MAPE) of 3.08%, outperforming the ARIMA model with an MAPE of 7.92%, indicating superior prediction accuracy for monthly AIDS incidence in Henan Province, China, from 2004 to 2016.
[37]	Luckett et al. (2021)	Cross-sectional study	USA	297 PLWH and 1509 HIV-negative controls	297	ANN	The saliency (SAL), parietal memory (PMN) and frontoparietal (FPN) networks had the highest predictive weights (up to 1.0) for classifying HIV status and cognitive impairment.
[38]	Luckett et al. (2019)	Cross-sectional study	USA	PLWH	125	ANN	Deep neural networks (DNNs) classified cognitive impairment among PLWH with 82–86% accuracy (AUC = 0.81–0.87) and frailty status with 75% accuracy, using cerebral blood flow features from key cortical and subcortical regions like the amygdala, hippocampus, thalamus, and temporal and parietal lobes.

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[39]	Marathe et al. (2022)	Mixed-methods study	Canada	PLWH with hepatitis C virus	2018 HIV-HCV co-infection; 717 food security sub-study (1934 total visits)	Random forest, feature importance analysis	A random forest model predicted depressive symptoms (CES-D-10 ≥ 10) in individuals with HIV and HCV with AUC = 0.82 using 137 predictors; key predictors were employment, HIV stage, revenue source, BMI and education.
[40]	Cheah et al. (2024)	Mixed-methods study	Malaysia	Men who have sex with men	14	NLP; local language adaptation	An AI chatbot promoting HIV testing and PrEP among Malaysian MSM was found feasible and acceptable, with 93% finding it useful and 79% willing to reuse it; users valued its stigma-free, user-friendly design and suggested adding local language and more health content.
[41]	Mulyadi and Qomariyah (2023)	Retrospective secondary data analysis	Indonesia	National-level HIV surveillance data: ART coverage (%), HIV cases, HIV-related deaths, total population per country	75 countries	K-nearest neighbours, logistic regression, support vector machine, XGBoost, decision tree, random forest, K-means clustering	Random forest achieved the highest accuracy (90%) in predicting ART outcomes, while K-means identified two clusters (15 vs. 60 countries) based on ART coverage and HIV prevalence.
[42]	Mutai et al. (2021)	Retrospective study	Sub-Saharan Africa: Tanzania, Zambia, Malawi and Eswatini	Individuals tested for HIV	263,829	Agglomerative hierarchical clustering	XGBoost predicted HIV status with 96% accuracy and AUC = 0.99; top predictors included age, number of lifetime partners, HIV testing history and condom use.

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[43]	Mutai et al. (2023)	Longitudinal study	13 sub-Saharan African countries	Population-based HIV impact assessment data	146,733 male and 155,622 female	Agglomerative hierarchical clustering	Two country clusters (per sex) were identified using agglomerative hierarchical clustering and PCA on 302,355 participants from 13 sub-Saharan African countries; HIV positivity varied significantly by predictors like school enrollment, urban residence and known HIV status.
[44]	Paul et al. (2020)	Randomized controlled trial	Thailand Cambodia Vietnam Indonesia	Children with perinatally acquired HIV	285	Gradient-boosted multivariate regression	GBM models predicted neurocognitive trajectories with 79% AUC at baseline and up to 90% with longitudinal data; mental health, hematocrit and CD4 count were key predictors.
[45]	Paul et al. (2020)	Cohort study	USA	PLWH	105	Gradient-boosted multivariate regression	GBM identified frailty with 71% F1 score, 84% precision and 66% sensitivity; key predictors included low CD4, poor psychomotor function and visuomotor neuroimaging features, with added risk from female sex and depressive symptoms.
[46]	Pranav et al. (2020)	Cross-sectional study	South Africa	PLWH with suspected tuberculosis	677	CNN	Deep learning model achieved 79% accuracy versus 65% for clinicians assisted by the model ($p = 0.002$), showing it significantly improves TB diagnosis in PLWH patients using chest X-rays.

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Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[47]	Roche et al. (2024)	Observational study	Kenya	PLWH	1500	Object detection (YOLOX Nano), classification (MixNet), image quality assurance	AI algorithm interpreting HIV self-test images achieved 100% sensitivity, 98.8% specificity, 100% NPV and 81.5% PPV, outperforming both clients (93.2% sensitivity) and providers (97.7%) in detecting true positives.
[48]	Andresen et al. (2022)	Longitudinal study	Switzerland	Men who have sex with men	6354	Hierarchical clustering, regression analysis	Unsupervised ML clustering of MSM living with HIV identified behaviour-based subgroups that improved prediction of 10 sexual behaviour and STI outcomes (e.g. condomless sex, syphilis), with AUROC increases of 0.03–0.17 over models using individual predictors alone.
[49]	Shi et al. (2022)	Observational study	China	PLWH with talaromycosis	1927	Logistic regression, XGBoost, k-nearest neighbours (k-NN), SVM	XGBoost model predicted in-hospital mortality with 0.90 AUC, 0.69 sensitivity and 0.96 specificity; key predictors included septic shock, respiratory failure, UA, urea, PLT count and AST/ALT ratio.
[50]	Turbé et al. (2021)	Feasibility study	South Africa	HIV rapid test images	11,374	CNN	A deep learning model trained on 11,374 HIV rapid test images collected in rural South Africa achieved 97.8% sensitivity and 100% specificity in field deployment, outperforming human readers and supporting REASSURED diagnostic principles for scalable, connected HIV testing.

(Continued)

Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[51]	Wang et al. (2023)	Randomized controlled trial	The Bahamas	Grade 10–12 students	2564	Support vector machine (SVM), logistic regression, decision tree, random forest, Boruta feature selection	Random forest predicted intervention non-responsiveness with 84.3% sensitivity, 67.1% specificity, AUROC 0.85; key predictors included self-efficacy, parent monitoring and HIV knowledge. SVM radial model using 10 clinical factors (e.g. low Hb, CD4+, PLT; high LDH, BUN; no ART; imaging findings) predicted BSI in PLWH with high performance (AUC = 0.916, sensitivity = 82.4%, specificity = 85.5%).
[52]	Wu et al. (2023)	Retrospective study	China	PLWH	498	SVM with radial basis kernel	A LASSO regression model predicted increased comorbidity burden (Charlson Index) in 28.2% of PLWH using EHR data, identifying key predictors such as age at diagnosis, tobacco use, low CD4+ and retention duration, achieving high predictive performance.
[53]	Yang et al. (2021)	Cohort study	USA	PLWH	8253	LASSO regression	

(Continued)

Table 1. (Continued)

Ref	Author	Study design	Country/region	Population	Sample size ^a	AI methods used	Key findings
[54]	Yu et al. (2019)	Qualitative study	China	HIV and hepatitis C virus targets from therapeutic target and ChEMBL databases	HIV: 11 targets; Hepatitis C: 4 targets	Naive Bayes, support vector machine	A multiple QSAR approach using naive Bayes and SVM with MACCS and ECFP6 fingerprints predicted 20 multitarget HIV/HCV inhibitors with high accuracy (AUC 0.83–1.0); seven of nine novel compounds were experimentally validated to act on both HIV-1 and HCV targets.
[55]	Zhang et al. (2018)	Retrospective study	Yale University	PLWH	1137	Ensemble learning, elastic net, LASSO, random forest, SVM, XGBoost	Ensemble ML identified 698 smoking-associated CpGs predictive of HIV frailty (AUC = 0.73–0.78) and 5-year mortality (HR = 1.46, <i>p</i> = 0.02) in PLWH, highlighting integrin signalling involvement.

Abbreviations: ADABOOST, adaptive boosting; AI, artificial intelligence; ANFIS, adaptive neuro-fuzzy inference system; ANN, artificial neural network; ARIMA, auto-regressive integrated moving average model; ART, antiretroviral therapy; AUC, area under the curve; BRNN, bidirectional recurrent neural network; BSI, bloodstream infection; CASP, Critical Appraisal Skills Programme; CD4, cluster of differentiation 4; CHWs, community health workers; CNN, convolutional neural network; DT, decision tree; EFV, efavirenz; FBP, filtered back projection; GB, gradient boosting; GNB, Gaussian naive Bayes; GOSDT, global optimal sparse decision trees; HCV, hepatitis C virus; HIV, human immunodeficiency virus; IDI, Infectious Disease Institute; KNN, k-nearest neighbours; LASSO, least absolute shrinkage and selection operator; LMT, logistic model tree; LOCO, leave-one-covariate-out inference; LR, logistic regression; ML, machine learning; MLP, multilayer perceptron; MLR, multiple linear regression; MSM, men who have sex with men; NLP, natural language processing; PLWH, people living with HIV; PNP, peripheral neuropathy; PRISMA, Preferred Reporting Items for Systematic Reviews and Meta-Analyses; RDT, rapid diagnostic test; RF, random forest; RSN, resting state network; SHAP, Shapley additive explanations; SVM, support vector machine; TB, tuberculosis; TDM, therapeutic drug monitoring; TIV, trivalent inactivated influenza vaccine; VL, viral load; WTV, weighted total variation; XGB, extreme gradient boosting; YOLOX Nano, You Only Look Once Nano.

^aUnless otherwise stated, sample size values represent the number of participants; some entries specify number of tests, facilities or datasets.

Africa [59]; demonstrated the potential for identifying high-risk patients with AUC values of 0.65 and 0.52, respectively, in Mozambique and Nigeria [9], and 75% accuracy [15]. A categorical boosting ML model, applied in a South African cohort, correctly identified 22,119 of 35,985 missed clinic visits, achieving a sensitivity of 62%, specificity of 67% and a positive predictive value of 20% [34, 58].

AI methods applied to predict VL suppression and identify patients in HIV care at risk of virological failure utilized algorithms such as RFs, LR and SVMs [16, 18, 20, 57]. These models, trained on routine patient data, including clinic attendance, viral load results and treatment history, generally achieved varying levels of accuracy (approximately 80–96%) and sensitivity (72–97%), but successfully identifying patients at a high risk for disengagement and highlighting the potential for targeted interventions to improve outcomes and optimize resource allocation. In South Africa, these models predicted retention and viral suppression with an AUC of up to 0.76 [59]. Models achieved 78% accuracy in identifying viral load hotspots in Kenya [16], 97% accuracy in predicting virological failure [20].

3.3 | Clinical and immunological treatment outcomes

AI has shown promise in improving the monitoring of clinical outcomes in PLWH. Predictive models have been applied to predict clinical progression to AIDS [12], diagnose PLWH with laboratory-confirmed bacteraemia or fungemia (hereafter, bloodstream infection [BSI]) [52], predict immune function recovery [35] and identify comorbidity predictors [53]. Additionally, various ML models have been employed to identify individuals at high risk for HIV-associated morbidity—for example development of peripheral neuropathy [10] and mortality.

Models achieved 97% accuracy in predicting CD4 status in Ethiopia, and 90% accuracy in clustering regions by ART outcomes [20, 41]. In Yunnan, China, a study developed ML models to predict long-term immune function changes in PLWH on ART. The SVM model performed best for CD4+ T cell recovery in patients with counts <200 cells/ μ l, while the RF model excelled for those with counts \geq 200 cells/ μ l [35]. At Wenzhou Central Hospital in China, a study developed an AI diagnostic model to identify PLWH with BSI on the basis of eight clinical factors combined with age and gender. The model, built using the SVM with Radial Basis Function Kernel (svmRadial) algorithm, achieved an AUC curve of 0.916, sensitivity of 0.824 and specificity of 0.855, demonstrating excellent performance in diagnosing PLWH with BSI [52].

Furthermore, ML and AI have significantly enhanced diagnostic tools for HIV-related conditions. In Malawi, a computer-aided digital chest X-ray system improved the timeliness of tuberculosis (TB) diagnoses, reducing the median time to TB treatment initiation from 11 days under standard care to just 1 day [25]. In South Africa, the CheXaid deep learning algorithm achieved 79% accuracy in diagnosing TB from chest X-rays in PLWH, outperforming human clinicians (65% accuracy) and proving particularly valuable in resource-limited settings [46].

In the United States, a study used deep neural networks to classify cognitive impairment and frailty in PLWH, achieving accuracies of 82–86% and 75%, respectively, while identifying critical cerebral blood flow patterns [38]. In a subsequent study, ML models were used to analyse resting state networks of the brain in 297 virologically suppressed PLWH and 1509 healthy controls, identifying that the salience and parietal memory networks effectively distinguished HIV status, while features of the frontal parietal network were associated with cognitive impairment, with minor variability in predictive strength observed across different age groups [37].

3.4 | Testing and treatment support

AI chatbots have shown substantial potential in enhancing HIV services in Malaysia, Hong Kong and Singapore by providing accessible, stigma-free and personalized support. In Malaysia, 93% of users found chatbots useful, and 79% expressed willingness to continue using them for HIV testing and related services [40]. A focus group with 31 MSM in Malaysia highlighted that chatbots addressing HIV testing were well-received for their anonymity and ability to overcome systemic barriers like stigma and discrimination [23]. In Hong Kong, a trial involving 528 participants demonstrated that chatbots effectively promoted self-testing uptake and provided counselling comparable to traditional methods [60]. In Singapore, chatbots offered accurate, comprehensive health information while reducing barriers to care, addressing stigma, and improving access to HIV prevention and treatment [61].

3.5 | ML techniques

Various AI techniques have been extensively applied to enhance HIV care and research by addressing critical clinical and public health challenges (Table 2). Artificial neural networks (ANNs) [9, 12, 27, 29, 30, 35–38] and RF [9–12, 14, 16–20, 27, 29, 32, 33, 35, 39, 41, 51, 55] models have been utilized to predict treatment outcomes, viral load (VL) suppression and long-term health trends. SVM [20, 22, 30, 33, 35, 41, 49, 51, 52, 54] and LR [9–11, 18, 22, 41, 48, 49, 51] have been applied to identify factors influencing clinic attendance, immune function and comorbidities. Deep learning models, including convolutional neural networks [25, 46, 50], assist in diagnosing co-infections like TB through chest X-rays, while bidirectional recurrent neural networks [21] capture temporal trends in VL and CD4 counts. Natural language processing [23, 26, 40, 60] has enabled the development of chatbots to promote HIV self-testing and enhance patient engagement. Specialized techniques like YOLOX Nano [47] for interpreting HIV self-test images, clustering methods for identifying socio-behavioural predictors and gradient boosting [20, 44, 45] for predicting patient retention and ART interruptions, further highlight the capabilities of AI in HIV care. These applications showcase the versatility of AI in optimizing diagnosis, treatment and care delivery for PLWH.

4 | DISCUSSION

The current review underscores the significant potential of predictive ML applications in advancing HIV care, aligning

Table 2. Machine learning techniques

Specific ML technique	Description	Applications
Artificial neural networks (ANNs) [9, 12, 27, 29, 30, 35-38]	Deep learning models inspired by biological neurons; excel in complex, non-linear pattern recognition and are widely used in medical prediction tasks.	Predicting viral suppression and treatment outcomes [9, 12, 30]; predicting long-term health outcomes [19, 27]; guiding resource allocation during high-risk periods [36]; ART-drug and demographic analysis [30]; predicting changes in immune function [35]; classifying cognitive impairment and frailty in PLWH [37, 38]; predicting and visualizing outcomes of HIV index testing [32]; predicting HIV/AIDS knowledge [29].
Random forest (RF) [9-12, 14, 16-20, 27, 29, 32, 33, 35, 39, 41, 51, 55]	Ensemble method using multiple decision trees; offers high accuracy, resistance to overfitting and handles missing data well.	Predicting viral suppression [17], CD4 count and treatment outcomes [9, 12, 14, 18, 20, 27, 41, 51]; stratifying patients for personalized care; predicting long-term health outcomes [19]; trivalent inactivated influenza vaccine immunogenicity in children living with HIV using in vitro gene expression testing [33]; carotid artery plaques in PLWH [11]; changes in immune function [35]; predicting and visualizing outcomes of HIV index testing [32]; linking smoking-related DNA changes to HIV outcomes [55]; predicting HIV/AIDS knowledge [29]; depressive symptoms [39]; peripheral neuropathy [10]; and clinic attendance [22].
Support vector machines (SVMs) [20, 22, 30, 33, 35, 41, 49, 51, 52, 54]	Supervised ML model for classification and regression; effective in high-dimensional spaces and binary outcomes.	Predicting viral suppression, CD4 count and treatment outcomes [20, 30, 35, 41, 49, 51]; trivalent inactivated influenza vaccine immunogenicity in children living with HIV using in vitro gene expression testing [33]; identified visual field defects by CD4 levels; linking smoking-related DNA changes to HIV outcomes [55]; predicting BSI among PLWH [52]; multitarget inhibitors for HIV/HCV coinfection [54]; and clinic attendance [22].
Logistic regression (LR) [9-11, 18, 22, 41, 48, 49, 51]	Interpretable model for binary classification; often used as a baseline for comparison.	Predicting viral suppression and treatment outcomes [9, 18, 31, 41, 49]; carotid artery plaques in PLWH [11]; HIV/AIDS knowledge [29]; peripheral neuropathy [10]; sexual behaviour [48]; and clinic attendance [22].

(Continued)

Table 2. (Continued)

Specific ML technique	Description	Applications
Least absolute shrinkage and selection operator (LASSO regression) [14, 22, 53, 55, 62]	Regularization method for sparse models; performs variable selection and reduces overfitting.	Useful in risk factor modelling; handles collinearity well; performance improves with large, structured datasets.
Elastic net regression [12, 22, 33]	Hybrid of ridge and LASSO regression; balances prediction accuracy and variable selection.	Combines LASSO and ridge strengths; reliable for variable selection; moderate performance in multi-factor modelling.
Convolutional neural networks (CNNs) [25, 46, 50]	Specialized deep learning models for image data; state-of-the-art in radiology and diagnostic imaging.	High specificity in imaging tasks; effective in TB/HIV screening and test classification; state-of-the-art in vision tasks.
Bidirectional recurrent neural networks (BRNNs) [21]	Neural networks that process sequential data forward and backward; ideal for time-series clinical outcomes.	Handles sequential data well; effective for resistance prediction; limited but strong use case for temporal data.
Gradient boosting (GB) [13, 34]	Ensemble method using decision trees trained sequentially; achieves strong performance in complex datasets.	Consistently high performance in ART interruption and patient outcome prediction; interpretable boosting mechanism.
Auto-regressive integrated moving average (ARIMA) [36]	Time series forecasting model; effective for predicting incidence and trends in public health.	Good for time series forecasting; useful in incidence trend modelling; best in linear trends with stationary data.
Natural language processing (NLP) [23, 26, 40, 60]	AI technique for analysing and understanding human language; used in chatbots and patient interaction tools.	Effective for text and chatbot interventions; promising engagement tool in HIV prevention and testing campaigns.

(Continued)

Table 2. (Continued)

Specific ML technique	Description	Applications
Decision tree models [15, 19, 20, 27, 29, 41, 51]	Simple, interpretable models using rule-based splits; helpful for initial exploratory analysis.	Exploring associations between demographic and clinical variables; providing interpretable insights for public health interventions [27]; predicting HIV outcomes [15, 20, 51]; HIV/AIDS knowledge [29]; and HIV outcomes [41].
Logistic model tree (LMT) [19]	Combines logistic regression with decision tree structure for interpretable classification.	Classifying anti-HIV-1 peptides based on their sequence-related properties [19].
Cox proportional hazards model (Cox regression) [25]	Statistical model for time-to-event outcomes; useful in retention, mortality and survival analysis.	Modelling survival rates and longitudinal health trends; identifying patterns in patient retention and mortality over time [25].
Bayesian modelling, expert system [27, 58]	Probabilistic models using prior knowledge and observed data; supports therapeutic drug monitoring.	Developing a computer-based system to model and interpret plasma ART concentrations for therapeutic drug monitoring (TDM) [58]; predicting HIV outcomes [27].
Linear regression (LR) [28]	Simple model for continuous outcomes; useful for associations but limited for non-linear patterns.	Identifying correlations between immunologic signatures, clinical parameters and HIV DNA levels [28].
Pairwise controlled manifold approximation projection (PaCMAP) [28]	Dimensionality reduction technique; useful for visualizing high-dimensional clustering in immune data.	Reducing the dimensionality of immunophenotyping data to identify clusters of participants with distinct HIV reservoirs [28].
Global optimal sparse decision trees (GOSDT) [28]	Sparse and interpretable decision trees optimized globally; supports rule-based classification.	Predicting immunophenotypes by identifying sparse, interpretable decision rules linking immune and clinical features [28].
Leave-one-covariate-out inference (LOCO) [28]	Statistical method to evaluate the influence of each covariate on model output; supports model transparency.	Assessing the importance of individual covariates in predicting immunophenotypes and HIV DNA levels [28].
Object detection (YOLOX Nano) [47]	Object detection model for interpreting images; used in HIV self-testing result classification.	Interpreting HIV self-testing images [47].
Agglomerative hierarchical clustering [43]	Unsupervised clustering method; groups similar cases and visualizes socio-behavioural HIV patterns.	Clustered countries by socio-behavioural HIV predictors [43].

Abbreviations: ANNs, artificial neural networks; ARIMA, auto-regressive integrated moving average; BRNN, bidirectional recurrent neural networks; CNNs, convolutional neural networks; GB, gradient boosting; GOSDT, global optimal sparse decision trees; LASSO, least absolute shrinkage and selection operator; LMT, logistic model tree; LOCO, leave-one-covariate-out inference; LR, logistic regression; NLP, natural language processing; PaCMAP, pairwise controlled manifold approximation projection; RF, random forest; SVMs, support vector machines; YOLOX Nano, You Only Look Once, a deep learning object detection model.

with broader analyses of ML in healthcare. Reviews, such as Obermeyer et al. on predictive analytics in general healthcare, and Topol on AI in clinical medicine, emphasize ML's versatility in improving patient outcomes through early intervention and resource optimization [63, 64]. Similarly, our analysis illustrates how ML models, using routine patient data, predict virological suppression, retention in care and clinical outcomes with moderate to high accuracy.

Previous reviews have noted ML's capacity to address critical healthcare gaps, such as predicting patients' risk of disease progression, treatment interruption, adverse drug reactions, and loss to follow-up and optimizing treatment pathways [65–67]. For example, Topol discusses ML's role in personalized medicine, which aligns with this review's findings on predicting individual patient outcomes, such as virological failure and immune function recovery [64]. The capacity of ML to address retention in care parallels findings in primary care reviews, where predictive models have improved appointment adherence and patient engagement strategies [63].

The use of AI in diagnostics, as highlighted in our analysis, mirrors broader findings in imaging and pathology. For example, the application of AI-assisted diagnostic tools in TB and BSI detection aligns with reviews in oncology and radiology, which documented the enhancement of diagnostic accuracy and efficiency through deep learning models [68]. These advancements underscore ML's versatility in integrating clinical data and imaging for timely and precise diagnosis.

Previous reviews have identified challenges in the implementation of AI within clinical practice, including poor data quality, algorithmic bias and limited scalability of models across diverse settings [67, 69]. Issues such as the need for high-quality training datasets, ethical considerations in model deployment and integration into clinical workflows have been extensively documented [64]. Furthermore, the introduction of AI into HIV care presents significant regulatory and ethical challenges, particularly concerning data privacy and potential biases in AI models [69, 70]. Ensuring patient confidentiality is paramount, as AI systems often require access to sensitive health information, raising concerns about data security and unauthorized use [71]. Moreover, AI models trained on non-representative datasets may perpetuate existing health disparities, leading to biased outcomes in diagnosis and treatment [72].

To address the identified challenges, practical steps, including embedding equity into AI design from the outset by ensuring inclusive data, applying bias-aware modelling strategies, auditing decision outcomes and institutionalizing fairness through policy, can help prevent the perpetuation of health inequities [73]. Moreover, clear regulatory frameworks and ethical guidelines should be developed to safeguard patient data privacy and support responsible AI deployment in HIV care [70, 74]. Strengthening collaborations between clinicians, data scientists and policymakers is essential for aligning AI innovations with frontline healthcare needs [63].

This synthesis reinforces the broader consensus that ML has the potential to transform healthcare delivery across various disciplines. While specific to HIV care, the findings reflect universal themes in ML applications, emphasizing the need for ongoing research and collaboration to maximize the bene-

fits of these technologies in achieving better health outcomes globally.

AI and ML have the potential to revolutionize HIV care, particularly in sub-Saharan Africa, where the epidemic remains most severe, and health systems face significant challenges. AI is transforming HIV care by enabling early identification of high-risk patients, strengthening retention strategies, accelerating TB/HIV co-infection diagnosis and expanding access to self-testing, with the potential of driving better health outcomes in the most affected regions. As these technologies continue to evolve, their integration into healthcare systems could optimize resource allocation, enhance early intervention strategies and improve overall patient outcomes. However, to maximize their impact, AI-driven solutions must be tailored to local contexts, address structural barriers and ensure equitable access, ultimately strengthening HIV care delivery in the regions that need it most.

Looking ahead, the next phase for AI in HIV care and research should focus on evaluating the long-term impact of AI interventions on patient outcomes, health system efficiency and equity in HIV care, particularly in under-resourced settings. Moreover, the development of locally relevant implementation frameworks grounded in ethical, regulatory and infrastructural realities will be essential to guide the responsible and sustainable integration of AI into HIV programmes.

The present review has several limitations. The heterogeneity of studies makes direct comparisons challenging due to variations in design, population and AI methods. Additionally, publication bias may influence findings, as negative results are less frequently reported. Only studies published in English were included due to resource constraints, which may have led to the exclusion of relevant studies published in other languages.

5 | CONCLUSIONS

Our review highlights the transformative potential of ML applications in HIV treatment and care, demonstrating their effectiveness in predicting critical outcomes such as retention in care, virological treatment response and clinical complications. By leveraging routine patient data, ML models have proven valuable in optimizing resource allocation, improving diagnostic accuracy and enabling targeted interventions, particularly in resource-limited settings. While significant advancements have been achieved, challenges such as data quality, algorithmic bias and integration into clinical workflows persist. Addressing these limitations through robust research, ethical considerations and scalable implementations will be crucial for maximizing the impact of ML on HIV care.

The findings of this work reinforce the broader potential of ML across healthcare disciplines, aligning with global efforts to enhance patient outcomes and achieve public health goals, including the UNAIDS 95-95-95 targets. Future research should focus on refining ML algorithms, fostering interdisciplinary collaborations and ensuring equitable access to AI-driven innovations in HIV treatment and beyond.

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COMPETING INTERESTS

The authors declare that they have no competing interests related to this study.

AUTHOR CONTRIBUTIONS

SN conceptualized the study, conducted the research, performed data extraction and analysis, wrote the manuscript and coordinated the overall study. EMM and ML reviewed the study, contributed to the study design, and assisted in defining the selection and exclusion criteria. JS reviewed the study with a specific focus on AI methods and contributed to refining the AI-related methodology. LM designed the search strategy, conducted the literature search and reviewed relevant studies. TS conducted the quality appraisal of the selected studies, ensuring methodological rigour. TR reviewed the study, contributed to the study design, and played a key role in refining the selection and exclusion criteria.

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DISCLAIMER

The views and opinions expressed in this study are those of the authors and do not necessarily reflect the official policy or position of any affiliated institution or organization.

DATA AVAILABILITY STATEMENT

All data used in this systematic review are derived from previously published studies, which are cited in the manuscript.

REFERENCES

- van Schalkwyk C, Mahy M, Johnson LF, Imai-Eaton JW. Updated data and methods for the 2023 UNAIDS HIV estimates. *J Acquir Immune Defic Syndr*. 2024;95(1S):e1–e4. <https://doi.org/10.1097/qai.0000000000003344>
- Du-Harpur X, Watt FM, Luscombe NM, Lynch MD. What is AI? Applications of artificial intelligence to dermatology. *Br J Dermatol*. 2020;183(3):423–30. <https://doi.org/10.1111/bjd.18880>
- Marcus JL, Sewell WC, Balzer LB, Krakower DS. Artificial intelligence and machine learning for HIV prevention: emerging approaches to ending the epidemic. *Curr HIV/AIDS Rep*. 2020;17:171–79.
- Contreras I, Vehi J. Artificial intelligence for diabetes management and decision support: literature review. *J Med Internet Res*. 2018;20(5):e10775.
- Kumar Y, Koul A, Singla R, Ijaz MF. Artificial intelligence in disease diagnosis: a systematic literature review, synthesizing framework and future research agenda. *J Amb Intell Hum Comput*. 2023;14(7):8459–86.
- Tsoi K, Yiu K, Lee H, Cheng HM, Wang TD, Tay JC, et al. Applications of artificial intelligence for hypertension management. *J Clin Hypertens*. 2021;23(3):568–74.
- Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *BMJ*. 2021;372:n71.
- Zeng H, Zhao Y, Meng S, Tang X, Guo H, Wang Y, et al. Exploring HIV prevention strategies among street-based female sex workers in Chongqing, China. *Int J Environ Res Public Health*. 2015;12(1):855–70. <https://doi.org/10.3390/ijerph120100855>

- Jeni S, Friedman J, Johnna S, Harris E, Bailey L. Predictive analytics using machine learning to identify ART clients at health system level at greatest risk of treatment interruption in Mozambique and Nigeria. *J Acquir Immune Defic Syndr*. 2022;90(2):154–60. <https://doi.org/10.1097/QAI.0000000000002947>
- Tu W, Johnson E, Fujiwara E, Gill MJ, Kong L, Power C. Predictive variables for peripheral neuropathy in treated HIV type 1 infection revealed by machine learning. *AIDS*. 2021;35(11):1785–93. <https://doi.org/10.1097/QAD.0000000000002955>
- Roy CM-H, Durand M, Chartrand-Lefebvre C, Gilles S, Cécile T, Cloutier G. Associative prediction of carotid artery plaques based on ultrasound strain imaging and cardiovascular risk factors in people living with HIV and age-matched control subjects of the CHACS Cohort. *J Acquir Immune Defic Syndr*. 2022;91(1):91–100. <https://doi.org/10.1097/QAI.0000000000003016>
- Domínguez-Rodríguez S, Serna-Pascual M, Oletto A, Barnabas S, Zuidewind P, Dobbels E, et al. Machine learning outperformed logistic regression classification even with limit sample size: a model to predict pediatric HIV mortality and clinical progression to AIDS. *PLoS One*. 2022;17(10):e0276116. <https://doi.org/10.1371/journal.pone.0276116>
- Federico M, Jovana M, Licia G, Michela B, Laura S, Gianluca C, et al. A machine learning approach to predict weight change in ART-experienced people living with HIV. *J Acquir Immune Defic Syndr*. 2023;94(5):474–81. <https://doi.org/10.1097/QAI.0000000000003302>
- Esber AL, Dear NF, King D, Francisco LV, Sing'oei V, Owuoth J, et al. Achieving the third 95 in sub-Saharan Africa: application of machine learning approaches to predict viral failure. *AIDS*. 2023;37(12):1861–70. <https://doi.org/10.1097/QAD.0000000000003646>
- Fahey KA, Wei L, Njau PF, Shabani S, Kwilasa S, Maokola W, et al. Machine learning with routine electronic medical record data to identify people at high risk of disengagement from HIV care in Tanzania. *PLOS Glob Public Health*. 2022;2(9):e0000720. <https://doi.org/10.1371/journal.pgph.0000720>
- Kagendi N, Mwau M. A machine learning approach to predict HIV viral load hotspots in Kenya using real-world data. *Health Data Sci*. 2023;3:1–10. <https://doi.org/10.34133/hds.0019>
- Mamo DN, Yilma TM, Fekadie M, Sebastian Y, Bizuayehu T, Melaku MS, et al. Machine learning to predict virological failure among HIV patients on antiretroviral therapy in the University of Gondar Comprehensive and Specialized Hospital, in Amhara Region, Ethiopia, 2022. *BMC Med Inf Decis Making*. 2023;23:1–20. <https://doi.org/10.1186/s12911-023-02167-7>
- Maskew M, Sharpey-Schafer K, De Voux L, Crompton T, Bor J, Rennick M, et al. Applying machine learning and predictive modeling to retention and viral suppression in South African HIV treatment cohorts. *Sci Rep*. 2022;12(1):12715. <https://doi.org/10.1038/s41598-022-16062-0>
- Poorinmohammad N, Mohabatkhar H. A comparison of different machine learning algorithms for the prediction of anti-HIV-1 peptides based on their sequence-related properties. *Int J Pept Res Ther*. 2015;21(1):57–62. <https://doi.org/10.1007/s10989-014-9432-x>
- Seboka BT, Yehualashet DE, Tesfa GA. Artificial intelligence and machine learning based prediction of viral load and CD4 status of people living with HIV (PLWH) on anti-retroviral treatment in Gedeo Zone public hospitals. *Int J Gen Med*. 2023;16:435–51. <https://doi.org/10.2147/IJGM.S397031>
- Steiner MC, Gibson KM, Crandall KA. Drug resistance prediction using deep learning techniques on HIV-1 sequence data. *Viruses*. 2020;12(5):560. <https://doi.org/10.3390/v12050560>
- Xu X, Fairley CK, Chow EPF, Lee D, Aung ET, Zhang L, et al. Using machine learning approaches to predict timely clinic attendance and the uptake of HIV/STI testing post clinic reminder messages. *Sci Rep*. 2022;12(1):8757. <https://doi.org/10.1038/s41598-022-12033-7>
- Peng ML, Wickersham JA, Altice FL, Shrestha R, Azwa I, Zhou X, et al. Formative evaluation of the acceptance of HIV prevention artificial intelligence chatbots by men who have sex with men in Malaysia: Focus Group Study. *JMIR Format Res*. 2022;6(10):e42055. <https://doi.org/10.2196/42055>
- Murnane PM, Ayieko J, Vittinghoff E, Gandhi M, Chaplain K, Beteniko M, et al. Machine learning algorithms using routinely collected data do not adequately predict viremia to inform targeted services in postpartum women living with HIV. *J Acquir Immune Defic Syndr*. 2021;88(5):439–47. <https://doi.org/10.1097/QAI.0000000000002800>
- MacPherson P, Webb EL, Kamchedzera W, Joekes E, Mjoli G, Lalloo DG, et al. Computer-aided X-ray screening for tuberculosis and HIV testing among adults with cough in Malawi (the PROSPECT study): a randomised trial and cost-effectiveness analysis. *PLoS Med*. 2021;18(9):e1003752. <https://doi.org/10.1371/journal.pmed.1003752>

26. Koh MCY, Ngiam JN, Yong J, Tambyah PA, Archuleta S. The role of an artificial intelligence model in antiretroviral therapy counselling and advice for people living with HIV. *HIV Med.* **2024**;25:59–67. <https://doi.org/10.1111/hiv.13604>
27. Olatosi B, Sun X, Chen S, Zhang J, Liang C, Weissman S, et al. Application of machine-learning techniques in classification of HIV medical care status for people living with HIV in South Carolina. *AIDS.* **2021**;35:S19–S28. <https://doi.org/10.1097/QAD.0000000000002814>
28. Semenova L, Wang Y, Falcinelli S, Archin N, Cooper-Volkheimer AD, Margolis DM, et al. Machine learning approaches identify immunologic signatures of total and intact HIV DNA during long-term antiretroviral therapy. **2023**. <https://doi.org/10.1101/2023.11.16.567386>
29. Aybar-Flores A, Talavera A, Espinoza-Portilla E. Predicting the HIV/AIDS knowledge among the adolescent and young adult population in Peru: application of quasi-binomial logistic regression and machine learning algorithms. *Int J Environ Res Public Health.* **2023**;20(7):5318. <https://doi.org/10.3390/ijerph20075318>
30. Bala K, Etikan I, Usman AG, Abba SI. Artificial-intelligence-based models coupled with correspondence analysis visualization on ART—cases from Gombe State, Nigeria: a comparative study. *Life.* **2023**;13(3):715. <https://doi.org/10.3390/life13030715>
31. Bisaso KR, Karungi SA, Kiragga A, Castelnuovo JKM. A comparative study of logistic regression based machine learning techniques for prediction of early virological suppression in antiretroviral initiating HIV patients. *BMC Med Inf Decis Making.* **2018**;18:23. <https://doi.org/10.1186/s12911-018-0659-x>
32. Chikusi H, Leo J, Kajjage S. Machine learning model for prediction and visualization of HIV index testing in northern Tanzania. *Int J Adv Comput Sci Appl.* **2022**;13(2):391–99. <https://doi.org/10.14569/IJACSA.2022.0130246>
33. Cotugno N, Santilli V, Pascucci GR, Manno EC, De Armas L, Pallikkuth S, et al. Artificial intelligence applied to in vitro gene expression testing (IVIGET) to predict trivalent inactivated influenza vaccine immunogenicity in HIV infected children. *Front Immunol.* **2020**;11:559590. <https://doi.org/10.3389/fimmu.2020.559590>
34. Rachel E, Carstens J, Sue LR, Tonderai M, Eisenstein M, Keiser O, et al. Validation and improvement of a machine learning model to predict interruptions in antiretroviral treatment in South Africa. *J Acquir Immune Defic Syndr.* **2023**;92(1):42–49. <https://doi.org/10.1097/QAI.0000000000003108>
35. Li B, Li M, Song Y, Lu X, Liu D, He C, et al. Construction of machine learning models to predict changes in immune function using clinical monitoring indices in HIV/AIDS patients after 9.9-years of antiretroviral therapy in Yunnan, China. *Front Cell Infect Microbiol.* **2022**;12:867737. <https://doi.org/10.3389/fcimb.2022.867737>
36. Li Z, Li Y. A comparative study on the prediction of the BP artificial neural network model and the ARIMA model in the incidence of AIDS. *BMC Med Inf Decis Making.* **2020**;20:1–13. <https://doi.org/10.1186/s12911-020-01157-3>
37. Luckett PH, Paul RH, Kayla H, Lee JJ, Shimony JS, Meeker KL, et al. Modeling the effects of HIV and aging on resting-state networks using machine learning. *J Acquir Immune Defic Syndr.* **2021**;88(4):414–19. <https://doi.org/10.1097/QAI.0000000000002783>
38. Luckett P, Paul RH, Navid J, Cooley SA, Wisch JK, Boerwinkle AH, et al. Deep learning analysis of cerebral blood flow to identify cognitive impairment and frailty in persons living with HIV. *J Acquir Immune Defic Syndr.* **2019**;82(5):496. <https://doi.org/10.1097/QAI.0000000000002181>
39. Marathe G, Moodie EEM, Brouillette M-J, Cox J, Cooper C, Delaunay CL, et al. Predicting the presence of depressive symptoms in the HIV-HCV co-infected population in Canada using supervised machine learning. *BMC Med Res Method.* **2022**;22:1–11. <https://doi.org/10.1186/s12874-022-01700-y>
40. Cheah MH, Gan YN, Altice FL, Wickersham JA, Shrestha R, Salleh NAM, et al. Testing the feasibility and acceptability of using an artificial intelligence chatbot to promote HIV testing and pre-exposure prophylaxis in Malaysia: mixed methods study. *JMIR Hum Factors.* **2024**;11. <https://doi.org/10.2196/52055>
41. Mulyadi WJ, Qomariyah NN. Using machine learning to analyse the effect of antiretroviral therapy (ART) on people with HIV. **2023**.
42. Mutai CK, McSharry PE, Ngaruye I, Musabanganji E. Use of machine learning techniques to identify HIV predictors for screening in sub-Saharan Africa. *BMC Med Res Method.* **2021**;21:1–11. <https://doi.org/10.1186/s12874-021-01346-2>
43. Mutai CK, McSharry PE, Ngaruye I, Musabanganji E. Use of unsupervised machine learning to characterise HIV predictors in sub-Saharan Africa. *BMC Infect Dis.* **2023**;23:1–13. <https://doi.org/10.1186/s12879-023-08467-7>
44. Paul RH, Cho KS, Belden AC, Mellins CA, Malee KM, Robbins RN, et al. Machine-learning classification of neurocognitive performance in children with perinatal HIV initiating de novo antiretroviral therapy. *AIDS.* **2020**;34(5):737–48. <https://doi.org/10.1097/QAD.0000000000002471>
45. Paul RH, Cho KS, Luckett P, Strain JF, Belden AC, Bolzenius JD, et al. Machine learning analysis reveals novel neuroimaging and clinical signatures of frailty in HIV. *J Acquir Immune Defic Syndr.* **2020**;84(4):414. <https://doi.org/10.1097/QAI.0000000000002360>
46. Pranav R, Chloe OC, Amit S, Nishit A, Li J, Amirhossein K, et al. CheXaid: deep learning assistance for physician diagnosis of tuberculosis using chest X-rays in patients with HIV. *NPJ Digital Med.* **2020**;3(1):115. <https://doi.org/10.1038/s41746-020-00322-2>
47. Roche S, Ekwunife O, Mendonca R, Kwach B, Omollo V, Zhang S, et al. Measuring the performance of computer vision artificial intelligence to interpret images of HIV self-testing results. *Front Public Health.* **2024**;12:1334881. <https://doi.org/10.3389/fpubh.2024.1334881>
48. Andresen S, Balakrishna S, Mugglin C, Schmidt AJ, Braun DL, Marzel A, et al. Unsupervised machine learning predicts future sexual behaviour and sexually transmitted infections among HIV-positive men who have sex with men. *PLoS Comput Biol.* **2022**;18(10):e1010559. <https://doi.org/10.1371/journal.pcbi.1010559>
49. Shi M, Lin J, Wei W, Qin Y, Meng S, Chen X, et al. Machine learning-based in-hospital mortality prediction of HIV/AIDS patients with *Talaromyces marneffei* infection in Guangxi, China. *PLoS Negl Trop Dis.* **2022**;16(5):e0010388. <https://doi.org/10.1371/journal.pntd.0010388>
50. Turb v, Herbst C, Mngomezulu T, Meshkinfamfar S, Dlamini N, Mhlongo T, et al. Deep learning of HIV field-based rapid tests. *Nat Med.* **2021**;27(7):1165–70. <https://doi.org/10.1038/s41591-021-01384-9>
51. Wang B, Liu F, Deveaux L, Ash A, Gerber B, Allison J, et al. Predicting adolescent intervention non-responsiveness for precision HIV prevention using machine learning. *AIDS Behav.* **2023**;27(5):1392–402. <https://doi.org/10.1007/s10461-022-03874-4>
52. Wu L, Xia D, Xu K. Multi-clinical factors combined with an artificial intelligence algorithm diagnosis model for HIV-infected people with bloodstream infection. *Infect Drug Resist.* **2023**;16:6085–97. <https://doi.org/10.2147/IDR.S423709>
53. Yang X, Zhang J, Chen S, Weissman S, Olatosi B, Li X. Utilizing electronic health record data to understand comorbidity burden among people living with HIV: a machine learning approach. *AIDS.* **2021**;35:S39–S51. <https://doi.org/10.1097/QAD.0000000000002736>
54. Yu W, Li W, Du T, Hong Z, Lin J. Targeting HIV/HCV coinfection using a machine learning-based multiple quantitative structure-activity relationships (multiple QSAR) method. *Int J Mol Sci.* **2019**;20(14):3572. <https://doi.org/10.3390/ijms20143572>
55. Zhang X, Hu Y, Aouizerat BE, Peng G, Marconi VC, Corley MJ, et al. Machine learning selected smoking-associated DNA methylation signatures that predict HIV prognosis and mortality. *Clin Epigenet.* **2018**;10:155. <https://doi.org/10.1186/s13148-018-0591-z>
56. Roche SD, Ekwunife OI, Mendonca R, Kwach B, Omollo V, Zhang S, et al. Measuring the performance of computer vision artificial intelligence to interpret images of HIV self-testing results. *Front Public Health.* **2024**;12:1334881. <https://doi.org/10.3389/fpubh.2024.1334881>
57. Wang D, Larder B, Revell A, Montaner J, Harrigan R, De Wolf F, et al. A comparison of three computational modelling methods for the prediction of virological response to combination HIV therapy. *Artif Intell Med.* **2009**;47(1):63–74. <https://doi.org/10.1016/j.artmed.2009.05.002>
58. Goicoechea M, Vidal A, Capparelli E, Rigby A, Kemper C, Diamond C, et al. A computer-based system to aid in the interpretation of plasma concentrations of antiretrovirals for therapeutic drug monitoring. *Antivir Ther.* **2007**;12(1):55–62.
59. Maskew M, Sharpey-Schafer K, De Voux L, Crompton T, Bor J, Rennie M, et al. Applying machine learning and predictive modeling to retention and viral suppression in South African HIV treatment cohorts. *Sci Rep.* **2022**;12(1):12715. <https://doi.org/10.1038/s41598-022-16062-0>
60. Chen S, Zhang Q, Chan C-K, Fuk-Yuen Y, Chidgey A, Yuan F, et al. Evaluating an innovative HIV self-testing service with web-based, real-time counseling provided by an artificial intelligence chatbot (HIVtest-chatbot) in increasing HIV self-testing use among Chinese men who have sex with men: protocol for a non-inferiority randomized controlled trial. *JMIR Res Protoc.* **2023**;12:e48447. <https://doi.org/10.2196/48447>
61. Sharpey-Schafer K, De Voux L, Crompton T, Bor J, Rennie M. The role of an artificial intelligence model in antiretroviral therapy counselling and advice for people living with HIV. *HIV Med.* **2024**; <https://doi.org/10.1111/hiv.13604>
62. Matta J, Singh V, Auten T, Sanjel P. Inferred networks, machine learning, and health data. *PLoS One.* **2023**;18(1):e0280910. <https://doi.org/10.1371/journal.pone.0280910>
63. Obermeyer Z, Emanuel EJ. Predicting the future—big data, machine learning, and clinical medicine. *N Engl J Med.* **2016**;375(13):1216–19. <https://doi.org/10.1056/NEJMp1606181>

64. Topol E. Deep medicine: how artificial intelligence can make healthcare human again. Hachette UK; 2019.
65. Dixon D, Sattar H, Moros N, Kesireddy SR, Ahsan H, Lakkimsetti M, et al. Unveiling the influence of AI predictive analytics on patient outcomes: a comprehensive narrative review. *Cureus*. 2024;16(5):e59954. <https://doi.org/10.7759/cureus.59954>
66. Abhadiomhen SE, Nzeakor EO, Oyibo K. Health risk assessment using machine learning: systematic review. *Electronics*. 2024;13(22):4405.
67. Alowais SA, Alghamdi SS, Alsuhebany N, Alqahtani T, Alshaya AI, Almohareb SN, et al. Revolutionizing healthcare: the role of artificial intelligence in clinical practice. *BMC Med Educ*. 2023;23(1):689. <https://doi.org/10.1186/s12909-023-04698-z>
68. Litjens G, Kooi T, Bejnordi BE, Setio AAA, Ciampi F, Ghafoorian M, et al. A survey on deep learning in medical image analysis. *Med Image Anal*. 2017;42:60–88.
69. Norori N, Hu Q, Aellen FM, Faraci FD, Tzovara A. Addressing bias in big data and AI for health care: a call for open science. *Patterns* (N Y). 2021;2(10):100347. <https://doi.org/10.1016/j.patter.2021.100347>
70. Garrett R, Kim S, Young SD. Ethical considerations for artificial intelligence applications for HIV. *AI*. 2024;5(2):594–601.
71. Murdoch B. Privacy and artificial intelligence: challenges for protecting health information in a new era. *BMC Med Ethics*. 2021;22(1):122. <https://doi.org/10.1186/s12910-021-00687-3>
72. Dankwa-Mullan I. Health equity and ethical considerations in using artificial intelligence in public health and medicine. *Prevent Chron Dis*. 2024;21:E64.
73. Agarwal R, Bjarnadottir M, Rhue L, Dugas M, Crowley K, Clark J, et al. Addressing algorithmic bias and the perpetuation of health inequities: an AI bias aware framework. *Health Policy Technol*. 2023;12(1):100702.
74. Topol EJ. High-performance medicine: the convergence of human and artificial intelligence. *Nat Med*. 2019;25(1):44–56.

SUPPORTING INFORMATION

Additional information may be found under the Supporting Information tab for this article:

File S1: Search strategy

File S2: Quality appraisal
PRISMA 2020 Checklist