HE²RO Policy Brief

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APPLYING MACHINE LEARNING TO LABORATORY DATA: PREDICTING SUPPRESSION OF NEXT HIV VIRAL LOAD IN SOUTH AFRICA

Background

During 2018, South Africa was estimated to have more than seven million people living with HIV[1], representing the largest single country epidemic[2] and treatment program. [3] In September 2016, the National Department of Health revised its treatment guidelines to extend the availability of ART to all people living with HIV, irrespective of CD4 cell count and stage of disease[3]. This policy, widely referred to as "treat all" or "universal test and treat" (UTT) holds promise to offer substantial advancements not only in the health of those living with HIV [4-5], but also in the country's efforts to meet 95-95-95 targets. However, implementation of a policy like UTT requires a rapid scale-up and expansion of the ART program on a country-wide level; a shift that is often challenging in these settings. By expanding eligibility, UTT has eliminated "pre-ART" care for most patients.[6] In many cases, this translates into more rapid initiation of ART with fewer required clinic visits prior to dispensing drugs. In many cases, the pre-ART care cascade is compressed into a single visit with same day initiation (SDI) of treatment. Though this increases uptake of ART, retention of new patients may have suffered. The problem of patient loss to follow-up (LTFU) within the public sector in South Africa has been well documented.[7] The potential for improved health through expanded ART availability will only be realized if individuals sustain engagement in HIV care.

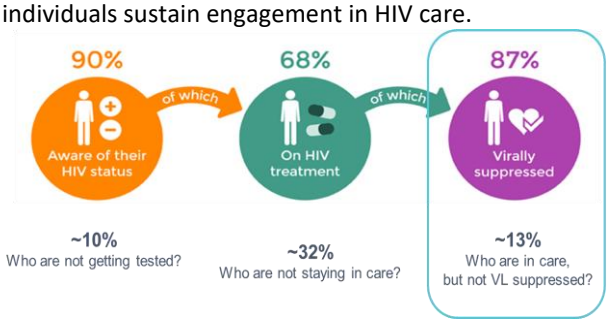

Figure 1: SA HIV care cascade, 2018 (avert.org)

To optimize South Africa's HIV response and reach targets of 95% tested, 95% treated, and 95% virally suppressed, numbers of patients initiating and successfully maintaining viral suppression on antiretroviral therapy must increase. In 2018, just 53% of people living with HIV (PLWH) in South Africa were virally suppressed.[1] While much effort and resources have been focused on tracing those LTFU and returning them to care, very little prior work has successfully addressed identifying those most at risk of poor treatment outcomes while still engaged in care.

Methods

We applied machine learning and modelling algorithms developed by Palindrome, data science implementers, (https://www.palindromedata.com) to de-identified HIV programmatic data collected from public sector treatment facilities based in two districts supported by Right to Care between 2015 and 2019. We included data for patients all patients who had accessed HIV care, initiated treatment and were retained through to virologic monitoring. HIV viral load (VL) suppression at next VL test was selected as primary outcome as it is an established clinical treatment outcome and objectively defined (diagnostically measurable reading) and thus made for a good target outcome to build confidence around the approach. High (>1000) Viral Loads that followed shortly (<6 months) after a previously high Viral Load were excluded from analyses due to the high probability of also being high.

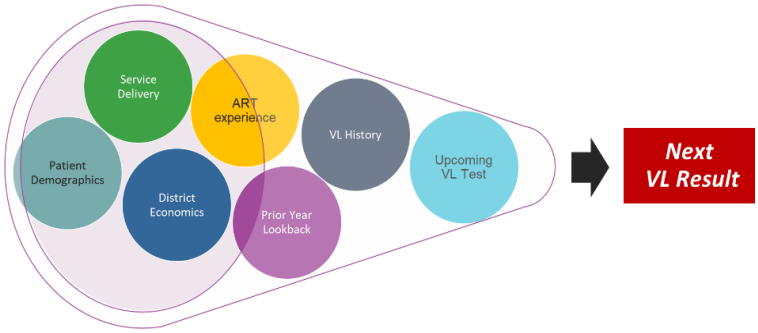

Figure 2: Framework for modelling compound effects

As the likelihood of viral load suppression is impacted by multiple components of the patient's treatment journey (Figure 2), demographic, clinical, behavioral (e.g. visit patterns) and laboratory data were investigated as potential predictor variables of VL result at next visit. Multiple models were created using various combinations of predictor variables and classification algorithms. These were then tested against unseen data to identify the optimal balance between predictive power and implementation feasibility. The final models were built using a random forest classifier and combined features. Models were evaluated using receiver operating characteristic (ROC) curves which assess the performance of each model's predictions against a test set of unseen data with known outcomes. The area under the curve (AUC) measures how well a variable can classify into two groups – in this case VL suppressed or unsuppressed. AUC values range between 0.5 (poor classifier) and 1.0 (excellent classifier).

Results

We included data on 688,614 VL results, during the study period 1 January 2016 – July 2019. We tested >50 potential input features per patient in 7 different models using multiple combinations of input features and classification algorithms. Each model was tested against unseen data to identify optimal predictive performance. Model results ranged from AUC of 0.57 for the poorest performing model (included gender and age at ART start only) to an AUC of 0.739 for the best performing model (Figure 3). Practically, this means the model correctly anticipated whether the VL result at next test would be suppressed or unsuppressed in approximately 3 out of 4 patients. The model consistently achieved an accuracy of 75% per month over the most recent months of patient data in 2019 suggesting it is both historically accurate but also relevant to patients currently accessing care.

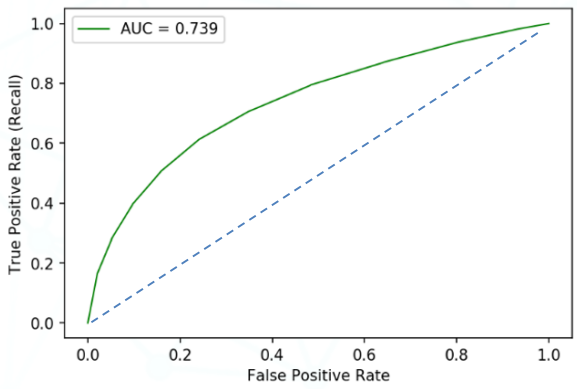

Figure 3. ROC curve for final model

Several patient characteristics were found to contribute a larger importance in the Random Forest model in terms of predicting their risk of having an unsuppressed VL at next visit (Figure 4). These included: age at ART initiation, most recent VL result value, time on ART, pattern of previous missed visits and month they accessed treatment. As a reminder, the Random Forest method observes the correlation of these features *in combination* – as such the figure should be read as a group, rather than an ordered list of priority or individual causation.

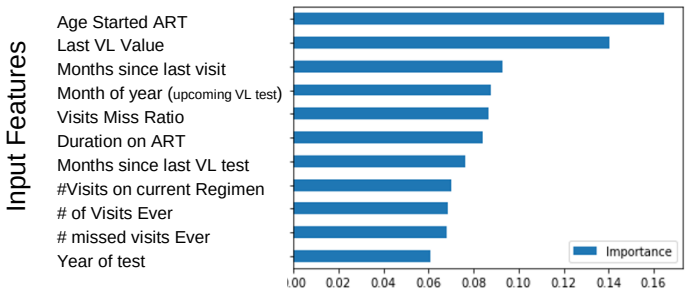

Figure 4. The top 11 features with largest predictive importance in final model, from the original 50

Policy relevance

As South Africa increases efforts towards 95-95-95 goals, knowing which patients require additional services and interventions in order to achieve successful treatment outcomes at each step of the cascade is critical. Our model was able to effectively separate high-risk from low-risk patients using a combination of clinical, laboratory and behavioral (visit patterns) data. Early detection of patients at high risk of becoming virologically unsuppressed has implications not only for the individual patient's health, but also for the risk of onward transmission of the virus and impact on breaking transmission chains. Potential operational application of these results could include the ability to score patients into risk categories at each visit and triage their care accordingly – high risk patients get prioritized to receive intensive intervention at point of care while low risk patients are expedited through the visit. On-going work will also continue to develop the model and explore other predicted outcomes such as risk of disengaging from care at next scheduled visit.

Leveraging predictive models to better understand the risk of individuals will allow for health care services to better triage patients, improving efficiency and resource utilization. By prioritizing those most at-risk, clinics can realize better health outcomes without additional investments in data collection and staff. Moreover, by anticipating future issues before any visible signs are present (e.g. an unsuppressed VL), clinics can intervene pro-actively while patients are still accessible, engaged in health services and provide targeted services earlier.

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