## MSS318R Activities Report [Appendix to Final Report]

# Automatic retinal image analysis (ARIA) to predict coronary artery disease in HIV-infected individuals

### Background

People living with HIV (PLWH) had twice the risk of developing cardiovascular diseases than the HIVuninfected population. Coronary atherosclerosis was common in PLWH. The currently available risk prediction tools for cardiovascular disease are derived from the general population or predominantly Caucasian PLWH. These tools have poorer performance in PLWH of Asian ethnicity. Therefore, more accurate non-invasive tools for prediction of cardiovascular disease, particularly in Asian PLWH, are urgently needed.

It has long been recognized that retinal characteristics are associated with the development of coronary artery disease. Recently, more detailed evaluation of the complex retinal vasculature has been made feasible with advances in digitalized retinal imaging techniques.

This study aims to determine the prevalence of coronary artery disease and the performance of retinal image analysis of the retinal vasculature in predicting coronary atherosclerosis and coronary artery disease in a cohort of PLWH with risk factors for cardiovascular disease in Hong Kong.

#### Methodology

We performed a prospective cross-sectional study. PLWH followed up at Infectious Diseases Clinic at Prince of Wales Hospital with one or more cardiovascular risk factors were enrolled to the study. We performed coronary CT angiogram for detection of coronary atherosclerosis and obstructive coronary artery stenosis, and automatic retina image analysis to evaluate a wide spectrum of characteristics of retinal vasculature. We determined the association between retinal information and other traditional cardiovascular risk factors and coronary artery disease, using multivariable stepwise logistic regression analyses.

#### Results

One hundred and twenty participants were enrolled. Two did not perform retinal imaging due to death and loss to follow-up. Three other patients were excluded from analyses due to inadequate images taken from one or both eyes. One hundred and fifteen participants were included in the final analyses.

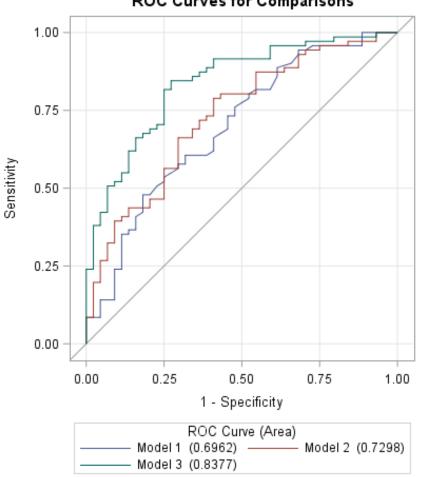
Seventy-one (62%) had coronary atherosclerosis, while 23 (20%) had obstructive coronary artery disease. Coronary atherosclerosis was associated with age and male gender, and a trend with higher triglycerides and lower HDL cholesterol. The retinal characteristics that were associated with coronary atherosclerosis included left and right adjusted central retinal arteriolar and venular equivalent and bifurcation coefficient of artery, and right venous asymmetry index. Table 1 showed the logistic regression analysis of the variables associated with coronary atherosclerosis using both traditional cardiovascular risk factors and retinal characteristics. Figure 1 and Table 2 showed the comparisons

between area under curves (AUC) for the three different models. Inclusion of both traditional cardiovascular risk factors and retinal characteristics had the best performance in predicting coronary atherosclerosis. Likewise, inclusion of both traditional cardiovascular risk factors and retinal characteristics had the best performance in predicting obstructive coronary artery disease.

95% CI for OR OR P Value Lower Upper Model 1: Inclusion of traditional cardiovascular risk factors only Sex 5.27 1.29 21.58 0.021 1.63 0.96 2.75 0.070 Age group Triglycerides 1.48 0.98 2.25 0.063 AUC: 0.6962 (95%CI: 0.596-0.7965) Model 2: Inclusion of retinal characteristics only 0.69 0.46 1.04 0.079 IMBCA 2.16 1.10 4.24 0.025 rCRVE 0.32 0.15 0.65 0.002 radjustedCRAE 0.58 0.36 0.92 0.020 rMVasymmetry AUC: 0.7298 (95%CI: 0.6356-0.8241) Model 3: Inclusion of traditional cardiovascular risk factors and retinal characteristics 16.92 3.05 93.91 0.0012 Sex 1.93 1.26 2.96 0.0026 Triglycerides 1.09 1.01 1.18 0.0209 **ICRVE** 0.39 0.18 0.84 0.0168 ladjustedCRAE 0.90 0.53 0.31 0.0183 IMBCA 0.50 0.29 0.84 0.0094 **IMAangle** 0.60 0.36 1.00 0.0515 **IExudates** 1.09 1.03 1.16 0.0046 **IAVR** 0.46 0.24 0.91 0.0255 radjustedCRAE 0.64 0.35 1.17 0.1495 rMBCA 0.60 0.36 1.01 0.0533 rTortuosity AUC: 0.8377 (95%CI: 0.7637-0.9117)

Table 1. Stepwise logistic regression analysis showing factors associated with coronary atheroslcerosis in different models.

Figure 1. AUC comparison between different models. (Model 1: Inclusion of traditional cardiovascular risk factors only; Model 2: Inclusion of retinal information only; Model 3: Inclusion of traditional cardiovascular risk factors and retinal information).



**ROC Curves for Comparisons** 

Table 2. Comparison of area under receiver operating characteristic curves (AUC) between different models using DeLong method.

Model comparison	P Value
Model 1 vs Model 2	0.617
Model 1 vs Model 3	0.001
Model 2 vs Model 3	0.037

#### Conclusion

Our study showed that inclusion of retinal characteristics was feasible and performed better in the prediction of coronary atherosclerosis and obstructive coronary artery disease in PLWH with cardiovascular risk factors than traditional cardiovascular risk factors.