

# NON-HIGH-DENSITY LIPOPROTEIN CHOLESTEROL LEVEL LINKED WITH INCIDENCE OF CHRONIC KIDNEY DISEASE IN HIV INFECTED PATIENTS

Masaki Hara<sup>1,2</sup> Minoru Ando, <sup>1,2</sup> Naoki Yanagisawa, <sup>1,2</sup> Ken Tsuchiya<sup>2</sup>, Kosaku Nitta<sup>2</sup>

<sup>1</sup>Division of Nephrology and Infectious Diseases, Department of Medicine, Tokyo Metropolitan, Komagome Hospital, Bunkyo-Ku, Tokyo, Japan, and <sup>2</sup>Department IV of Internal Medicine, Tokyo Women's Medical University, Shinjuku-Ku, Tokyo, Japan

## INTRODUCTION

- HIV-infected patients receiving antiretroviral therapy (cART) have a high likelihood of dyslipidemia.
- However, it has not been fully understood whether the presence of dyslipidemia predisposes HIV-infected patients to kidney disease.

## OBJECTIVES

- Correlation between non HDL-C and incident CKD.

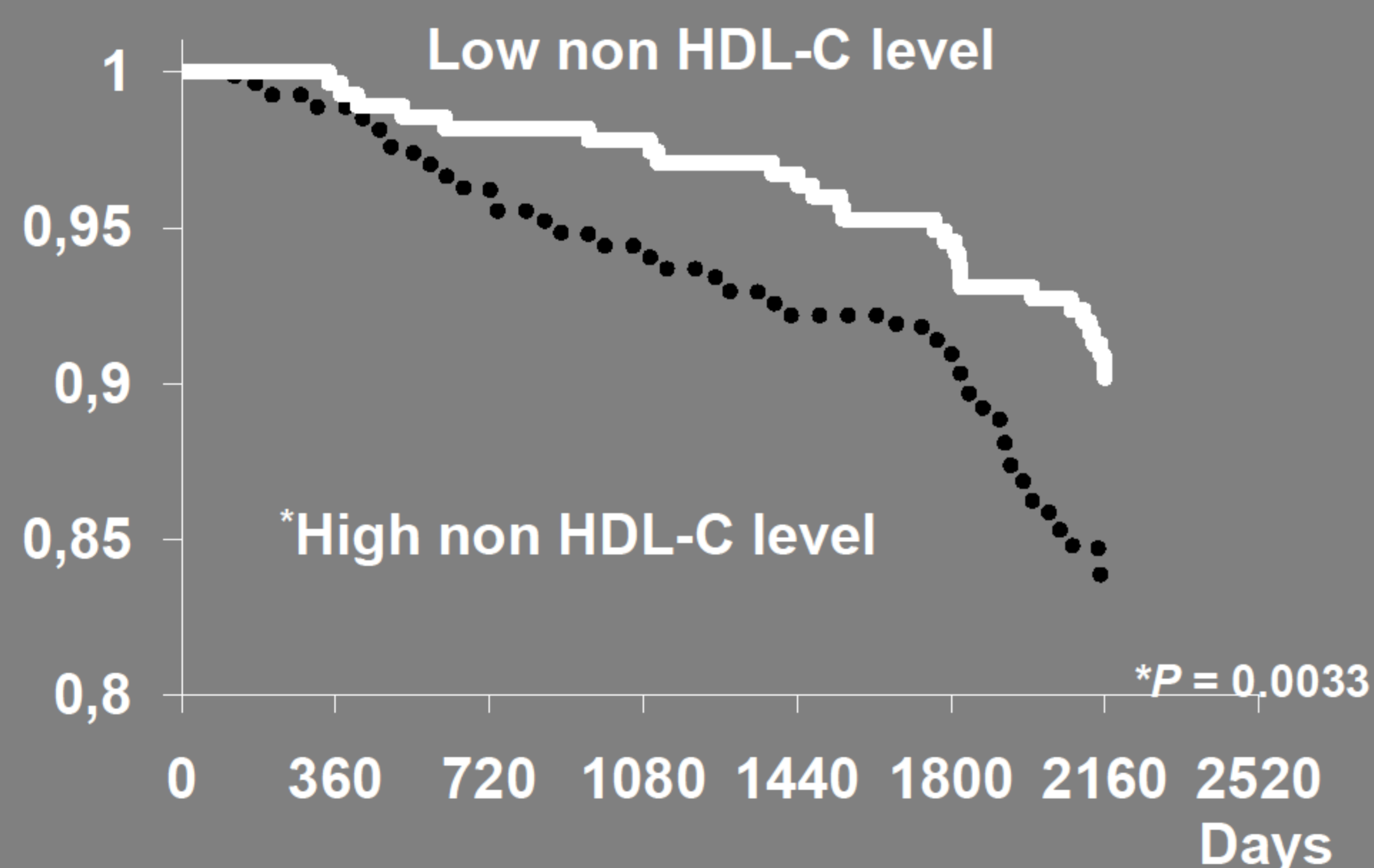
## RESULTS

Table 1. Demographics and laboratory characteristics

|                                   | High non HDL-C (N = 303) | Low non HDL-C (N = 302) | P value  |
|-----------------------------------|--------------------------|-------------------------|----------|
| Age (y)                           | 48 ± 11                  | 43 ± 11                 | <0.0001* |
| Men (%)                           | 91.0                     | 90.2                    | 0.7815   |
| Japanese (%)                      | 93.3                     | 94.4                    | 0.6137   |
| Prevalence of HT (%)              | 15.4                     | 19.0                    | 0.2812   |
| Prevalence of DM (%)              | 8.33                     | 2.62                    | 0.0021*  |
| Prevalence of hepatitis C (%)     | 2.00                     | 4.59                    | 0.1096   |
| cART+ (%)                         | 92.8                     | 87.4                    | 0.0410*  |
| PI+ (%)                           | 80.7                     | 74.8                    | 0.0967   |
| Duration of cART use (y)          | 5 ± 3                    | 4 ± 3                   | 0.0005*  |
| UACR (mg/g)                       | 114.9 ± 745.9            | 34.8 ± 102.9            | 0.0163*  |
| eGFR (ml/min/1.73m <sup>2</sup> ) | 87.1 ± 15.1              | 90.2 ± 17.7             | 0.0737   |
| CD4 (cells/μL)                    | 445 ± 216                | 389 ± 188               | 0.0013*  |
| HIV-RNA (< 50 copies/mL, %)       | 86.7                     | 76.7                    | 0.0016*  |
| Non HDL-C (mg/dL)                 | 170.9 ± 32.0             | 107.0 ± 18.6            | <0.0001* |

HDL-C, high-density lipoprotein cholesterol; HT, hypertension; DM, diabetes mellitus; cART, combination anti retroviral therapy; PI, protease inhibitor; UACR, urinary albumin creatinine ratio; eGFR, estimated glomerular filtration rate; HIV, human immune deficiency virus-1. Asterisk (\*) indicates that the parameter has statistical significance.

Figure 2. 6-year cumulative survival rate of incident CKD stratified by median value of non HDL-C level



## METHODS

Longitudinal study for six-years.

### Subjects:

•A total of 605 HIV-infected patients who had eGFR ≥ 60ml/min/m<sup>2</sup>.

### Methods:

- Incident CKD: eGFR less than 60 ml/min/1.73m<sup>2</sup> during follow-up period,
- Multivariable linear regression analysis*: factors associated with non HDL-C.
- Kaplan Meier analysis*: Cumulative survival rate.
- Cox proportional hazards regression model*: Mortality HR with 95% CI, adjusted for age, sex, prevalence of hypertension, prevalence of diabetes mellitus, presence or absence of cART, duration of cART use, albuminuria, eGFR, and high HIV viral load (≥ 50 copies/mL) at baseline.

Table 2. Factors associated with non HDL-C

| Factor                       | Multivariable linear regression analysis |                |         |
|------------------------------|--|----------------|---------|
|                              | β ± SE                                   | Standardized β | P-value |
| Age                          | 0.375 ± 0.174                            | 0.101          | 0.0317* |
| Sex, men                     | 1.897 ± 2.944                            | 0.027          | 0.5198  |
| Prevalence of HT             | -0.879 ± 2.308                           | -0.016         | 0.7032  |
| Prevalence of DM             | 13.48 ± 3.942                            | 0.151          | 0.0007* |
| cART+                        | 12.28 ± 7.148                            | 0.074          | 0.0862  |
| Average duration of cART use | 0.499 ± 0.599                            | 0.036          | 0.4526  |
| eGFR                         | -0.185 ± 0.108                           | -0.073         | 0.0903  |
| UACR                         | 0.009 ± 0.003                            | 0.120          | 0.0060* |
| CD 4                         | 0.026 ± 0.009                            | 0.133          | 0.0042* |
| High HIV viral load+         | -1.004 ± 2.943                           | -0.015         | 0.7330  |

HDL-C, high-density lipoprotein cholesterol; HT, hypertension; DM, diabetes mellitus; cART, combination anti retroviral therapy; eGFR, estimated glomerular filtration rate; UACR, urinary albumin creatinine ratio; HIV, human immune deficiency virus-1; β, regression coefficient; SE, standard error. Asterisk (\*) indicates that the parameter has statistical significance.

Figure 1. Correlation between non HDL-C and albuminuria

There were significantly positive correlation between non HDL-C and albuminuria.

Hypothesis : diagram of relationship between them

Table 3. Cox hazard analysis for incident CKD

| Variable                     | Univariable analysis |          | Multivariable analysis |          |
|------------------------------|----------------------|----------|------------------------|----------|
|                              | HR (95% CI)          | P-value  | HR (95% CI)            | P-value  |
| High non HDL-C               | 1.99 (1.26-3.22)     | 0.0031*  | 1.63 (1.02-2.67)       | 0.0423*  |
| Age, years                   | 1.05 (1.03-1.07)     | <0.0001* | 1.02 (0.99-1.04)       | 0.1188   |
| Sex, men                     | 1.28 (0.61-3.31)     | 0.5450   |                        |          |
| Prevalence of HT             | 1.99 (1.18-3.24)     | 0.0110*  | 1.44 (0.85-2.37)       | 0.1734   |
| Prevalence of DM             | 2.47 (1.09-4.83)     | 0.0314*  | 1.49 (0.61-3.27)       | 0.3609   |
| cART+                        | 1.42 (0.67-3.65)     | 0.3901   |                        |          |
| Average duration of cART use | 1.01 (0.94-1.08)     | 0.8772   |                        |          |
| Albuminuria                  | 1.002 (1.001-1.003)  | <0.0001* | 1.001 (1.000-1.002)    | 0.0127*  |
| eGFR                         | 0.92 (0.91-0.94)     | <0.0001* | 0.93 (0.91-0.95)       | <0.0001* |
| CD 4                         | 0.998 (0.997-0.999)  | 0.0070*  | 0.998 (0.997-0.999)    | 0.0026*  |
| High HIV viral load          | 0.90 (0.53-1.62)     | 0.9661   |                        |          |

HDL-C, high-density lipoprotein cholesterol; CKD, chronic kidney disease; HT, hypertension; DM, diabetes mellitus; cART, combination anti retroviral therapy; eGFR, estimated glomerular filtration rate; HR, hazard ratio; CI, confidence interval. Asterisk (\*) indicates that the parameter has statistical significance.

## SUMMARY AND DISCUSSION

- ❑ We confirmed that positive correlation between non HDL-C and albuminuria.
  - ⇒ Dyslipidemia may cause glomerulosclerosis and interstitial fibrosis through glomerular endothelial dysfunction, podocyte injury, and mesangial damage.
- ❑ Correlation between non HDL-C and incident CKD.
  - Non HDL-C was a significant factor about incident CKD.
  - ⇒ There are several reported that high non-HDL-C and low HDL-C was associated with renal impairment.

## CONCLUSIONS

HIV-infected patients with increased non-HDL-C level are more likely at high risk of future CKD. The intervention effect of lipid lowering therapy on incident CKD will be warranted in them.