**Supplementary Table 1. Relationship between virological markers/pathological features and intrahepatic cccDNA**

|  |  |  |  |
| --- | --- | --- | --- |
|  | **cccDNA-negative**  **(*n*=13)** | **cccDNA-positive**  **(*n*=35)** | ***p-*value** |
| ALT (U/L), median (IQR) | 29.0 (17.5, 42.0) | 24.0 (20.0, 30.0) | 0.508 |
| Intrahepatic cccDNA log10 copies/cell, mean (SD) | ND | -1.1 (0.7) | NA |
| HBcrAg log10 IU/mL, mean (SD) | 3.7 (0.6) | 3.5 (0.5) | 0.529 |
| pgRNA, (copies/mL) | ND | ND | NA |
| HBsAb |  |  | 0.179 |
| > 1000 | 2 (15.4%) | 3 (8.6%) |  |
| 2–1000 | 7 (53.8%） | 28 (80.0%) |  |
| < 2 | 4 (30.8%） | 4 (11.4%) |  |
| Inflammation |  |  | 0.069 |
| G0-2 | 11 (84.6%) | 35 (100%) |  |
| G3-4 | 2 (15.4%) | 0 (0%) |  |
| Fibrosis |  |  | 0.011 |
| S0–2 | 8 (61.5%) | 33 (94.3%) |  |
| S3–4 | 5 (38.5%) | 2 (5.7%) |  |
| HBsAg immunohistochemistry (+/-) | 3/10 | 9/26 | 1.000 |
| HBcAg immunohistochemistry (+/–) | 0/13 | 0/35 | 1.000 |

ALT, alanine aminotransferase; cccDNA, covalently closed circular DNA; HBcAg, hepatitis B core antigen; HBcrAg, hepatitis B core related antigen; HBsAb, hepatitis B surface antibody; HBsAg, hepatitis B surface antigen; IQR, interquartile range; NA, not available; ND, not detected; pgRNA, pre-genomic RNA; SD, standard deviation.