

SUPPLEMENTARY MATERIAL

Table S1: Adjusted comparison of viral suppression and change in CD4 cell counts at 24 and 48 weeks from ART initiation by first-line antiretroviral regimen, OT analyses, CoRIS cohort, 2018-2021

| | Viral suppression | | | | Change in CD4 cell count | | | |
|------------------|------------------------|------------------------|------------------------|------------------------|--|---|--|---|
| | 24 weeks | | 48 weeks | | 24 weeks | | 48 weeks | |
| | No./No. with data (%)* | Adjusted OR (95% CI)** | No./No. with data (%)* | Adjusted OR (95% CI)** | Mean change in CD4 cell count (95% CI) | Adjusted mean difference in CD4 cell count increase (95% CI)* | Mean change in CD4 cell count (95% CI) | Adjusted mean difference in CD4 cell count increase (95% CI)* |
| DTG/3TC | 234/254 (92.1) | 1 | 140/147 (95.2) | 1 | 208.3 (183.5; 233.1) | 0 | 262.5 (225.9; 299.0) | 0 |
| 3DR-InSTI | 831/989 (84.0) | 0.88 (0.57; 1.35) | 608/670 (90.7) | 0.69 (0.32; 1.46) | 202.7 (189.3; 216.2) | 1.88 (-26.38; 30.14) | 203.0 (188.7; 217.3) | 2.88 (-25.34; 31.11) |
| BIC/FTC/TAF | 518/623 (83.1) | 0.79 (0.52; 1.20) | 369/411 (89.8) | 0.66 (0.28; 1.57) | 201.3 (183.9; 218.6) | 1.73 (-28.62; 32.07) | 244.7 (222.2; 267.1) | -18.57 (-67.69; 30.56) |
| DTG/3TC/ABC | 145/175 (82.9) | 0.71 (0.39; 1.30) | 140/149 (94.0) | 1.07 (0.53; 2.14) | 196.2 (167.0; 225.4) | 1.81 (-38.38; 42.00) | 272.6 (235.4; 309.8) | 5.47 (-55.56; 66.49) |
| DRV/COBI/FTC/TAF | 83/113 (73.5) | 0.38 (0.24; 0.60) | 67/76 (88.2) | 0.57 (0.19; 1.66) | 184.8 (145.2; 224.4) | -8.35 (-54.67; 37.97) | 249.7 (197.4; 302.0) | -7.06 (-81.87; 67.75) |
| EVG/COBI/FTC/TAF | 63/69 (91.3) | 1.37 (0.57; 3.30) | 40/42 (95.2) | 1.02 (0.24; 4.32) | 185.5 (122.7; 248.3) | -0.23 (-48.95; 48.49) | 275.2 (190.0; 360.4) | 23.05 (-47.50; 93.60) |
| DTG+FTC/TDF | 105/122 (86.1) | 1.47 (0.64; 3.38) | 59/68 (86.8) | 0.69 (0.18; 2.58) | 228.5 (193.0; 264.0) | 18.75 (-26.25; 63.76) | 273.0 (219.8; 326.2) | -19.58 (-98.33; 59.17) |

DTG/3TC: dolutegravir/lamivudine; 3DR-InSTI: three-drug regimen including integrase inhibitor (BIC/FTC/TAF, DTG/3TC/ABC, EVG/COBI/FTC/TAF and DTG+FTC/TDF); BIC/FTC/TAF: bicitegravir/emtricitabine/tenofovir alafenamide; DTG/3TC/ABC: dolutegravir/lamivudine/abacavir; DRV/COBI/FTC/TAF: darunavir/cobicistat/emtricitabine/tenofovir alafenamide; EVG/COBI/FTC/TAF: elvitegravir/cobicistat/emtricitabine/tenofovir alafenamide; DTG+FTC/TDF: dolutegravir+emtricitabine/tenofovir disoproxil fumarate.

* Number of subjects achieving VS / Number of subjects with available data on VS (percentage of subjects achieving VS among those with available data)

**Adjusted for sex, age at ART initiation, transmission category, educational level, country of origin, CD4 cell count and viral load within 6 months previous to ART initiation, presence of hepatitis C virus antibodies, presence of hepatitis B virus surface antigen, and AIDS diagnosis at initiation of ART.

Appendix S1 Patient 1: 31-year-old male. At baseline, he had viral load 353,000 copies/ml, CD4 cell count 15/microl, he had no resistance to reverse transcriptase inhibitors and no integrase inhibitor resistance testing was available. He achieved undetectable viral load (VL) and had VF 3 and 10 months after the start of treatment, respectively. Resistance testing was performed at VF with no resistance to integrase inhibitors; reverse transcriptase could not be amplified. DTG/3TC was maintained until the last visit 19 months after ART initiation: during this time he had detectable viral load, always below 200 copies/ml.

Patient 2: 28-year-old male. At baseline, he had viral load 220 copies/ml, CD4 cell count 430/microl, and baseline resistance testing was not available. He achieved undetectability 2.5 months after the start of DTG/3TC. He switched to DRV/COB/TAF/FTC 4 months after ART initiation because of suspected adverse effect (anxiety), and he abandoned treatment 10 days later. He experienced VF 5 months after ART initiation (resistance testing was not performed at the time), and 10 days later DTG/3TC was reintroduced. Viral load was undetectable 3.5 months after the restart of DTG/3TC and has remained undetectable for 18 months until the last visit.

Patient 3: 35-year-old male. At baseline, he had VL 2,000/ml and CD4 cell count 375, and no resistance mutations to reverse transcriptase or integrase inhibitors. He achieved undetectability and had viral failure 2 and 6 months after starting DTG/3TC, respectively. He had low treatment compliance since the start of treatment.

Table S2: Adjusted comparison of viral suppression and change in CD4 cell counts at 24 and 48 weeks from ART initiation by first-line antiretroviral regimen in subgroups, CoRIS cohort, 2018-2021

S3a. Viral suppression

| | Viral suppression | | | |
|--|------------------------|------------------------|------------------------|------------------------|
| | 24 weeks | | 48 weeks | |
| | No./No. with data (%)* | Adjusted OR (95% CI)** | No./No. with data (%)* | Adjusted OR (95% CI)** |
| ART initiation with CD4 cell count < 200 cells/μL | | | | |
| DTG/3TC | 9/11 (81.8) | | 5/6 (83.3) | |
| BIC/FTC/TAF | 117/161 (72.7) | | 89/105 (84.8) | |
| DTG/3TC/ABC | 22/31 (71.0) | | 19/24 (79.2) | |
| DRV/COBI/FTC/TAF | 15/28 (53.6) | | 18/22 (81.8) | |
| EVG/COBI/FTC/TAF | 11/17 (64.7) | | 14/16 (87.5) | |
| DTG+FTC/TDF | 45/64 (70.3) | | 40/49 (81.6) | |
| ART initiation with HIV RNA VL > 100,000 copies/ml | | | | |
| DTG/3TC | 60/76 (78.9) | | 34/38 (89.5) | |
| BIC/FTC/TAF | 219/291 (75.3) | 1.30 (0.81; 2.09) | 170/198 (85.9) | 0.94 (0.30; 2.95) |
| DTG/3TC/ABC | 64/82 (78.0) | 1.44 (0.76; 2.75) | 65/73 (89.0) | 1.15 (0.27; 4.82) |
| DRV/COBI/FTC/TAF | 24/45 (53.3) | 0.56 (0.28; 1.11) | 25/34 (73.5) | 0.42 (0.11; 1.70) |
| EVG/COBI/FTC/TAF | 21/28 (75.0) | 1.28 (0.67; 2.45) | 24/27 (88.9) | 1.32 (0.35; 5.02) |
| DTG+FTC/TDF | 112/141 (79.4) | 1.63 (0.74; 3.59) | 92/107 (86.0) | 1.01 (0.37; 2.73) |
| ART initiation with HIV RNA VL > 500,000 copies/ml | | | | |
| DTG/3TC | 10/18 (55.6) | | 5/6 (83.3) | |
| BIC/FTC/TAF | 85/124 (68.5) | | 75/86 (87.2) | |
| DTG/3TC/ABC | 18/29 (62.1) | | 25/30 (83.3) | |
| DRV/COBI/FTC/TAF | 11/22 (50.0) | | 7/12 (58.3) | |
| EVG/COBI/FTC/TAF | 7/11 (63.6) | | 11/13 (84.6) | |
| DTG+FTC/TDF | 50/68 (73.5) | | 39/47 (83.0) | |
| ART initiation within 7 days of enrolment | | | | |
| DTG/3TC | 127/142 (89.4) | | 71/77 (92.2) | |
| BIC/FTC/TAF | 350/419 (83.5) | 1.14 (0.69; 1.87) | 268/302 (88.7) | 1.04 (0.35; 3.06) |
| DTG/3TC/ABC | 53/63 (84.1) | 1.10 (0.47; 2.57) | 50/54 (92.6) | 1.44 (0.43; 4.82) |
| DRV/COBI/FTC/TAF | 38/55 (69.1) | 0.42 (0.16; 1.08) | 34/40 (85.0) | 0.75 (0.23; 2.43) |
| EVG/COBI/FTC/TAF | 47/54 (87.0) | 1.42 (0.66; 3.05) | 47/50 (94.0) | 1.61 (0.46; 5.63) |
| DTG+FTC/TDF | 114/142 (80.3) | 1.07 (0.41; 2.77) | 91/106 (85.8) | 0.96 (0.42; 2.22) |
| Women | | | | |
| DTG/3TC | 18/20 (90.0%) | | 7/7 (100.0%) | |
| BIC/FTC/TAF | 68/75 (90.7%) | | 45/52 (86.5%) | |
| DTG/3TC/ABC | 11/13 (84.6%) | | 14/15 (93.3%) | |
| DRV/COBI/FTC/TAF | 11/15 (73.3%) | | 8/10 (80.0%) | |
| EVG/COBI/FTC/TAF | 8/9 (88.9%) | | 9/10 (90.0%) | |
| DTG+FTC/TDF | 24/30 (80.0%) | | 24/28 (85.7%) | |

* Number of subjects achieving VS / Number of subjects with available data on VS (percentage of subjects achieving VS among those with available data)

**Adjusted for sex, age at ART initiation, transmission category, educational level, country of origin, CD4 cell count and viral load within 6 months previous to ART initiation, presence of hepatitis C virus antibodies, presence of hepatitis B virus surface antigen, and AIDS diagnosis at initiation of ART.

ART: antiretroviral therapy; DTG/3TC: dolutegravir/lamivudine; BIC/FTC/TAF: bictegravir/emtricitabine/tenofovir alafenamide; DTG/3TC/ABC: dolutegravir/lamivudine/abacavir; DRV/COBI/FTC/TAF: darunavir/cobicistat/emtricitabine/tenofovir alafenamide; EVG/COBI/FTC/TAF: elvitegravir/cobicistat/emtricitabine/tenofovir alafenamide; DTG+FTC/TDF: dolutegravir+emtricitabine/tenofovir disoproxil fumarate.

S2b. Change in CD4 cell counts

| | Change in CD4 cell count | | | |
|--|--|---|--|---|
| | 24 weeks | | 48 weeks | |
| | Mean change in CD4 cell count (95% CI) | Adjusted mean difference in CD4 cell count increase (95% CI)* | Mean change in CD4 cell count (95% CI) | Adjusted mean difference in CD4 cell count increase (95% CI)* |
| ART initiation with CD4 cell count < 200 cells/μL | | | | |
| DTG/3TC | 155.5 (115.1; 196.0) | | 250.8 (168.4; 333.2) | |
| BIC/FTC/TAF | 150.5 (130.8; 170.2) | | 226.2 (195.7; 256.8) | |
| DTG/3TC/ABC | 173.3 (123.6; 223.1) | | 242.9 (182.6; 303.1) | |
| DRV/COBI/FTC/TAF | 135.1 (101.7; 168.5) | | 198.0 (147.5; 248.4) | |
| EVG/COBI/FTC/TAF | 184.6 (133.0; 236.2) | | 209.1 (152.9; 265.3) | |
| DTG+FTC/TDF | 168.9 (137.5; 200.2) | | 199.4 (167.8; 231.0) | |
| ART initiation with HIV RNA VL > 100,000 copies/ml | | | | |
| DTG/3TC | 225.1 (185.8; 264.4) | | 331.7 (271.0; 392.4) | |
| BIC/FTC/TAF | 216.9 (192.1; 241.8) | 26.69 (-15.86; 69.25) | 277.7 (246.5; 308.8) | -5.22 (-55.50; 45.07) |
| DTG/3TC/ABC | 237.7 (198.9; 276.5) | 44.79 (-7.18; 96.77) | 313.3 (262.0; 364.6) | 20.96 (-40.43; 82.34) |
| DRV/COBI/FTC/TAF | 197.3 (132.0; 262.7) | 9.67 (-65.41; 84.75) | 259.3 (186.8; 331.8) | -28.46 (-93.71; 36.79) |
| EVG/COBI/FTC/TAF | 222.2 (144.2; 300.1) | 30.74 (-62.50; 123.98) | 277.4 (174.8; 379.9) | -5.06 (-94.88; 84.75) |
| DTG+FTC/TDF | 250.5 (214.8; 286.3) | 48.55 (-4.11; 101.22) | 285.5 (239.3; 331.7) | -15.03 (-101.90; 71.85) |
| ART initiation with HIV RNA VL > 500,000 copies/ml | | | | |
| DTG/3TC | 268.9 (163.6; 374.1) | | 371.4 (107.1; 635.8) | |
| BIC/FTC/TAF | 252.6 (216.4; 288.8) | | 320.9 (269.3; 372.5) | |
| DTG/3TC/ABC | 258.8 (197.7; 320.0) | | 353.2 (257.5; 448.9) | |
| DRV/COBI/FTC/TAF | 279.0 (156.6; 401.5) | | 339.8 (189.1; 490.4) | |
| EVG/COBI/FTC/TAF | 180.6 (78.1; 283.0) | | 264.6 (137.5; 391.8) | |
| DTG+FTC/TDF | 283.1 (223.7; 342.5) | | 339.1 (265.4; 412.9) | |
| ART initiation within 7 days of enrolment | | | | |
| DTG/3TC | 209.5 (179.2; 239.9) | | 250.1 (201.9; 298.4) | |
| BIC/FTC/TAF | 208.4 (186.1; 230.7) | 9.02 (-26.33; 44.38) | 258.8 (233.1; 284.5) | 12.99 (-44.81; 70.80) |
| DTG/3TC/ABC | 259.6 (208.7; 310.4) | 54.25 (-6.23; 114.73) | 325.1 (267.3; 382.9) | 61.03 (-7.56; 129.63) |
| DRV/COBI/FTC/TAF | 191.8 (142.7; 241.0) | 1.73 (-71.98; 75.45) | 278.0 (210.7; 345.3) | 47.87 (-40.25; 135.99) |
| EVG/COBI/FTC/TAF | 181.2 (125.6; 236.8) | -8.69 (-78.65; 61.26) | 308.1 (246.5; 369.8) | 71.35 (4.34; 138.35) |
| DTG+FTC/TDF | 261.3 (223.2; 299.5) | 50.68 (-2.62; 103.99) | 302.6 (257.3; 347.8) | 39.25 (-35.38; 113.87) |
| Women | | | | |
| DTG/3TC | 174.9 (94.9; 255.0) | | 364.4 (28.1; 700.7) | |
| BIC/FTC/TAF | 225.4 (174.4; 276.5) | | 237.8 (185.2; 290.3) | |
| DTG/3TC/ABC | 288.1 (167.5; 408.7) | | 249.5 (155.7; 343.2) | |

| | | | | |
|------------------|----------------------|--|----------------------|--|
| DRV/COBI/FTC/TAF | 246.1 (171.1; 321.1) | | 190.3 (112.8; 267.8) | |
| EVG/COBI/FTC/TAF | 348.0 (98.2; 597.8) | | 254.6 (-28.5; 537.8) | |
| DTG+FTC/TDF | 233.2 (143.3; 323.1) | | 290.7 (190.8; 390.6) | |

**Adjusted for sex, age at ART initiation, transmission category, educational level, country of origin, CD4 cell count and viral load within 6 months previous to ART initiation, presence of hepatitis C virus antibodies, presence of hepatitis B virus surface antigen, and AIDS diagnosis at initiation of ART.*

ART: antiretroviral therapy; DTG/3TC: dolutegravir/lamivudine; BIC/FTC/TAF: bictegravir/emtricitabine/tenofovir alafenamide; DTG/3TC/ABC: dolutegravir/lamivudine/abacavir; DRV/COBI/FTC/TAF: darunavir/cobicistat/emtricitabine/tenofovir alafenamide; EVG/COBI/FTC/TAF: elvitegravir/cobicistat/emtricitabine/tenofovir alafenamide; DTG+FTC/TDF: dolutegravir+emtricitabine/tenofovir disoproxil fumarate.