

| sTable 1 Number, crude rate, age-standardized DALYs rate for overall DUDs in 2021 and percentage change from 1990 |                                      |                     |                         |                     |                         |                     |                        |                     |                      |                     |                      |                     |                      |                     |                      |                     |
|-------------------------------------------------------------------------------------------------------------------|--------------------------------------|---------------------|-------------------------|---------------------|-------------------------|---------------------|------------------------|---------------------|----------------------|---------------------|----------------------|---------------------|----------------------|---------------------|----------------------|---------------------|
| DALYs number                                                                                                      |                                      |                     | DALYs rate              |                     | Overall                 |                     | Opioid                 |                     | Cocaine              |                     | Amphetamine          |                     | Cannabis             |                     | Other drug           |                     |
| location                                                                                                          | Number                               | EAPC                | Crude rate              | EAPC                | ASR                     | EAPC                | ASR                    | EAPC                | ASR                  | EAPC                | ASR                  | EAPC                | ASR                  | EAPC                | ASR                  | EAPC                |
| Tropical Latin America                                                                                            | 162061.67(110807.96 - 213561.19)     | -2.1(-2.44 - 1.88)  | 106.23(72.64 - 139.99)  | -0.83(-1.17 - 0.61) | 102.09(71.29 - 133.35)  | -0.77(-1.03 - 0.59) | 41.29(27.16 - 55.97)   | 0.23(0.18 - 0.26)   | 21.2(13.77 - 31.77)  | -2.68(-3.25 - 2.19) | 24.06(13.51 - 37.6)  | 0.11(0.04 - 0.1)    | 13.91(7.96 - 22.24)  | 0.45(0.29 - 0.49)   | 1.63(1.24 - 2.14)    | -4.44(-4.95 - 3.94) |
| Global                                                                                                            | 8910603.44(7055603.29 - 10630912.05) | -1.78(-1.89 - 1.71) | 167.06(132.29 - 199.32) | -0.53(-0.64 - 0.46) | 166.44(132.55 - 198.4)  | -0.44(-0.53 - 0.37) | 103.69(81.83 - 122.75) | -0.9(-1.02 - 0.88)  | 10.91(7.86 - 14.84)  | -0.77(-1.13 - 0.53) | 29.63(19.51 - 43.52) | 1.12(0.95 - 1.28)   | 8.63(5.1 - 13.25)    | 0.14(0.13 - 0.1)    | 13.58(11.05 - 18.14) | 0.78(0.41 - 1.4)    |
| High-income North America                                                                                         | 1078039.6(831794.64 - 1308306.68)    | -5.77(-6.05 - 5.61) | 383.09(295.58 - 464.91) | -4.93(-5.22 - 4.78) | 352.05(270.73 - 427.06) | -5.19(-5.47 - 5.04) | 207.65(165.2 - 248.95) | -6.18(-6.29 - 6.08) | 67.79(45.53 - 97.79) | -2.48(-3.13 - 2.02) | 32.67(18.88 - 50.76) | -3.5(-4.54 - 2.81)  | 29.53(17.89 - 45.14) | 0.19(0.17 - 0.17)   | 14.41(12.77 - 16.45) | -4.48(-4.47 - 4.5)  |
| Central Latin America                                                                                             | 139214.36(105074.32 - 176664.83)     | -1.71(-1.8 - 1.64)  | 84.68(63.91 - 107.46)   | -0.33(-0.43 - 0.26) | 86.88(65.99 - 109.39)   | -0.06(-0.14 - 0.01) | 43.16(29.44 - 56.57)   | 0.22(0.15 - 0.22)   | 25.39(18.95 - 34.28) | -0.03(0 - 0)        | 7.43(4.46 - 11.21)   | -0.53(-0.8 - 0.36)  | 6.43(3.92 - 9.86)    | -0.68(-0.75 - 0.59) | 4.48(3.84 - 5.24)    | -0.78(-0.83 - 0.74) |
| South Asia                                                                                                        | 686910.37(536014.83 - 854810.86)     | -2.53(-2.56 - 2.49) | 62.82(49.02 - 78.18)    | -0.87(-0.9 - 0.83)  | 71.11(56.1 - 87.26)     | -0.33(-0.33 - 0.29) | 55.72(42.26 - 68.1)    | -0.33(-0.32 - 0.35) | 2.76(1.34 - 4.81)    | -0.17(-1.3 - 0.58)  | 2.02(1.33 - 3.13)    | -0.54(-0.67 - 0.23) | 7.59(4.48 - 12.05)   | 0.05(0.01 - 0.02)   | 3.02(2.1 - 5.66)     | -1.05(-1.62 - 0.3)  |

|                          | DALYs number                       |                     | DALYs rate              |                     | Overall                 |                     | Opioid                  |                     | Cocaine              |                     | Amphetamine           |                     | Cannabis            |                     | Other drug           |                     |
|--------------------------|------------------------------------|---------------------|-------------------------|---------------------|-------------------------|---------------------|-------------------------|---------------------|----------------------|---------------------|-----------------------|---------------------|---------------------|---------------------|----------------------|---------------------|
| location                 | Number                             | EAPC                | Crude rate              | EAPC                | ASR                     | EAPC                | ASR                     | EAPC                | ASR                  | EAPC                | ASR                   | EAPC                | ASR                 | EAPC                | ASR                  | EAPC                |
| High-income Asia Pacific | 166825.47(118067.02 - 220263.36)   | 0.29(0.17 - 0.4)    | 96.22(68.1 - 127.04)    | 0.51(0.39 - 0.61)   | 89.87(63.71 - 119.15)   | -0.01(-0.07 - 0.04) | 43.18(29.21 - 57.21)    | -0.04(-0.18 - 0.01) | 14.88(8.99 - 23.47)  | 0.03(0.07 - 0.04)   | 15.18(8.42 - 23.49)   | 0(-0.1 - 0.05)      | 13.69(7.79 - 22.47) | 0.04(0.04 - 0.01)   | 2.94(2.08 - 4.14)    | 0.1(0.04 - 0.18)    |
| East Asia                | 3614386.8(2897066.28 - 4299278.97) | 2.39(2.51 - 2.31)   | 296.88(237.96 - 353.14) | 3.02(3.14 - 2.94)   | 268.49(217.53 - 316.18) | 2.71(2.89 - 2.56)   | 153.51(120.43 - 181.16) | 3.41(3.54 - 3.24)   | 2.96(2.04 - 4.13)    | 2.19(2.37 - 2.16)   | 73.34(50.41 - 104.33) | 1.61(1.73 - 1.52)   | 4.85(2.82 - 7.75)   | -0.69(-0.64 - 0.71) | 33.83(25.87 - 48.59) | 3.79(3.69 - 4.35)   |
| Central Europe           | 124195.04(97749.04 - 151799.02)    | 0(-0.08 - 0.07)     | 99.28(78.14 - 121.35)   | -0.26(-0.34 - 0.2)  | 98.28(77.04 - 120.4)    | -0.47(-0.53 - 0.42) | 52.45(41.21 - 63.16)    | -0.66(-0.82 - 0.59) | 7.79(5.84 - 10.86)   | 0.67(1.09 - 0.43)   | 21.57(12.6 - 33.39)   | -0.7(-0.85 - 0.63)  | 10.36(6.26 - 16.38) | 0.12(0.07 - 0.23)   | 6.11(5.29 - 7.03)    | -0.06(-0.05 - 0.07) |
| Central Asia             | 92368.72(67553.27 - 118438.49)     | -1.73(-1.89 - 1.56) | 133.27(97.46 - 170.88)  | -0.7(-0.86 - 0.52)  | 135.73(100.13 - 171.36) | -0.5(-0.64 - 0.38)  | 104.53(75.93 - 132.93)  | -0.38(-0.53 - 0.27) | 5.11(3.74 - 7.06)    | -1.46(-1.63 - 1.18) | 16.12(9.58 - 24.82)   | -0.72(-0.98 - 0.51) | 5.56(2.96 - 9.47)   | -0.12(-0.05 - 0.08) | 4.41(3.46 - 5.48)    | -1.49(-1.66 - 1.34) |
| Eastern Europe           | 768334.28(614076.36 - 907539.44)   | -0.26(-0.47 - 0.2)  | 339.23(271.13 - 400.7)  | -0.56(-0.76 - 0.49) | 323.6(257.79 - 382.26)  | -0.71(-0.87 - 0.65) | 250.75(195.86 - 302.22) | -0.69(-0.9 - 0.61)  | 16.25(13.63 - 19.71) | 0.71(0.89 - 0.5)    | 28.68(18.13 - 42.77)  | -1.03(-1.46 - 0.71) | 9.2(5.29 - 14.86)   | -0.2(-0.1 - -0.15)  | 18.73(16.86 - 20.82) | -1.47(-1.5 - 1.42)  |



|                             | DALYs number          |                 | DALYs rate      |                  | Overall         |                   | Opioid          |                   | Cocaine           |                 | Amphetamine       |                   | Cannabis             |                   | Other drug           |                       |
|-----------------------------|-----------------------|-----------------|-----------------|------------------|-----------------|-------------------|-----------------|-------------------|-------------------|-----------------|-------------------|-------------------|----------------------|-------------------|----------------------|-----------------------|
| location                    | Number                | EAPC            | Crude rate      | EAPC             | ASR             | EAPC              | ASR             | EAPC              | ASR               | EAPC            | ASR               | EAPC              | ASR                  | EAPC              | ASR                  | EAPC                  |
|                             | -                     |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   | 1 - -             |                      |                   |                      |                       |
|                             | 6143.33)              |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   | 0.05)             |                      |                   |                      |                       |
| Australasia                 | 79156.5               |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   |                   |                      |                   |                      |                       |
|                             | 6(63933.22 -          | -1.95(-2.08 - - | 390.39(315.31 - | -0.6(-0.73 - -   | 365.23(294.85 - | -0.77(-0.88 - -   | 194.37(161.72 - | -0.18(-0.12 - -   | 28.88(17.39 -     | -0.41(-0.45 - - | 65.48(38.31 -     | -0.71(-1.08 - -   | 35.19(22.77 -        | 1.73(1.71 - 1.62) | 41.31(36.93 -        | -3.48(-3.45 - -       |
|                             | 93223.68)             | 1.88)           | 459.77)         | 0.53)            | 430.61)         | 0.73)             | 225.02)         | 0.21)             | 45.95)            | 0.35)           | 102.45)           | 0.44)             | 51.28)               |                   | 45.91)               | 3.48)                 |
| Southern Sub-Saharan Africa | 82678.2               |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   |                   |                      |                   |                      |                       |
|                             | 2(64117.82 -          | -1.11(-1.27 - - | 157.73(122.32 - | 0.26(0.1 - 0.36) | 175.43(138.47 - | 0.66(0.54 - 0.71) | 121.87(93.86 -  | 1.02(0.87 - 1.09) | 16.46(12.09 -     | -0.16(-0.09 - - | 18.29(10.99 -     | 0.14(-0.09 -      | 7.3(4.28 - 11.46)    | -0.44(-0.35 - -   | 11.51(9.54 -         | 0.1(-0.1 - 0.32)      |
|                             | 100809.1)             | 1.01)           | 192.32)         |                  | 210.37)         |                   | 147.22)         |                   | 22.78)            | 0.1)            | 27.48)            | 0.32)             |                      |                   | 14.07)               |                       |
| Southern Latin America      | 51692.8               |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   |                   |                      |                   |                      |                       |
|                             | 8(35509.65 -          | -1.34(-1.49 - - | 104.35(71.68 -  | -0.34(-0.49 - -  | 104.52(72.1 -   | -0.17(-0.3 - -    | 51.92(33.12 -   | 0.01(-0.28 -      | 32.28(19.05 -     | -0.27(-0.32 - - | 9.4(5.26 - 15.29) | -0.22(-0.39 - -   | 8.39(5.29 - 12.81)   | -0.77(-0.99 - -   | 2.53(1.69 - 3.61)    | -0.3(-0.48 - -        |
|                             | 68800.7)              | 1.26)           | 138.88)         | 0.27)            | 139.02)         | 0.11)             | 69.72)          | 0.09)             | 51.34)            | 0.14)           |                   | 0.13)             |                      |                   |                      | 0.2)                  |
| Western Europe              | 847230.94(685035.12 - | -1.02(-1.13 - - | 220.4(178.2 -   | -0.6(-0.72 - -   | 211.34(170.89 - | -0.86(-0.96 - -   | 136.19(113.6 -  | -0.86(-0.89 - -   | 20.73(13.42 -     | -0.56(-0.7 - -  | 23.53(14.04 -     | -1(-1.39 - -0.73) | 17.95(11.31 - 26.84) | 0.12(0.02 - 0.16) | 12.95(11.26 - 14.99) | -1.98(-2.11 - - 1.84) |
|                             | 1005491.98)           | 0.94)           | 261.57)         | 0.53)            | 250.97)         | 0.8)              | 159.53)         | 0.84)             | 30.4)             | 0.46)           | 36.51)            |                   |                      |                   |                      |                       |
|                             |                       |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   |                   |                      |                   |                      |                       |
| Central Sub-Saharan Africa  | 23760.5               |                 |                 |                  |                 |                   |                 |                   |                   |                 |                   |                   |                      |                   |                      |                       |
|                             | 9(17783.58 -          | -3.45(-3.37 - - | 43.23(32.36 -   | -0.57(-0.48 - -  | 52.77(39.86 -   | -0.37(-0.29 - -   | 39.45(28.8 -    | -0.43(-0.39 - -   | 1.96(1.24 - 3.24) | -0.45(-0.54 - - | 5.09(2.76 - 8.08) | -0.2(-0.27 - -    | 5.13(2.75 - 8.5)     | -0.05(-0.06 - -   | 1.15(0.65 - 1.69)    | -0.29(-0.5 - - 0.33)  |
|                             | 30833.37)             | 3.42)           | 56.1)           | 0.54)            | 67.88)          | 0.34)             | 51.84)          | 0.36)             |                   |                 |                   | 0.11)             |                      |                   |                      |                       |

| DALYs number         |              |                 | DALYs rate     |                | Overall        |                 | Opioid         |                 | Cocaine        |                | Amphetamine |                 | Cannabis    |                  | Other drug        |                     |
|----------------------|--------------|-----------------|----------------|----------------|----------------|-----------------|----------------|-----------------|----------------|----------------|-------------|-----------------|-------------|------------------|-------------------|---------------------|
| location             | Number       | EAPC            | Crude rate     | EAPC           | ASR            | EAPC            | ASR            | EAPC            | ASR            | EAPC           | ASR         | EAPC            | ASR         | EAPC             | ASR               | EAPC                |
| Andean Latin America | 30542.1      |                 |                |                |                |                 |                |                 |                |                |             |                 |             |                  |                   |                     |
|                      | 9(22396.51 - | -2.38(-2.56 - - | 80.39(58 .95 - | -0.61(-0.8 - - | 83.02(62 .22 - | -0.32(-0.45 - - | 43.69(29 .09 - | -0.18(-0.37 - - | 16.74(12 .96 - | -0.79(-0.8 - - | 9.17(5.14 - | -0.59(-0.95 - - | 7.07(4.09 - | 0.1(0.05 - 0.08) | 6.34(5.19 - 7.61) | 0.14(0.24 - - 0.02) |
|                      | 38683.36)    | 2.32)           | 101.82)        | 0.56)          | 104.32)        | 0.28)           | 59.24)         | 0.15)           | 21.58)         | 0.77)          | 14.16)      | 0.48)           | 11.19)      |                  |                   |                     |

Notes: ASR, age-standardized rate per 100000 residents; EAPC, estimated annual percent change (%); data in () indicates the uncertainty interval, it reflects the certainty of an estimate based on data availability, studies size and consistency across data sources.