

**Supplementary Table 1. Summary of adverse events in atogepant and placebo groups**

| Adverse events                             | Dose           | Studies reported (n) | The incidence rate in the atogepant group (event/total, %) | The incidence rate in the placebo group (event/total, %) |
|--|----------------|----------------------|--|--|
| Any TEAE                                   | 10.00 mg (QD)  | 2                    | 178/314, 56.70%  | 218/408, 53.40%  |
|  | 30.00 mg (QD)  | 2                    | 234/411, 57.00%  | 218/408, 53.40%  |
|  | 60.00 mg (QD)  | 3                    | 312/573, 54.45%  | 302/565, 53.45%  |
|  | 30.00 mg (BID) | 1                    | 52/86, 60.40%  | 92/186, 49.50%   |
|  | 60.00 mg (BID) | 1                    | 53/91, 58.24%  | 92/186, 49.50%   |
| Treatment-related TEAE                     | 10.00 mg (QD)  | 2                    | 68/314, 21.65%   | 50/408, 12.25%   |
|  | 30.00 mg (QD)  | 2                    | 73/411, 17.76%   | 50/408, 12.25%   |
|  | 60.00 mg (QD)  | 3                    | 100/573, 17.45%  | 64/565, 11.33%   |
|  | 30.00 mg (BID) | 1                    | 18/86, 20.90%  | 30/186, 16.10%   |
|  | 60.00 mg (BID) | 1                    | 24/91, 26.40%  | 30/186, 16.10%   |
| SAEs                                       | 10.00 mg (QD)  | 2                    | 3/314, 0.95%   | 4/408, 0.98%   |
|  | 30.00 mg (QD)  | 2                    | 2/411, 0.49%   | 4/408, 0.98%   |
|  | 60.00 mg (QD)  | 3                    | 6/573, 1.05%   | 4/565, 0.70%   |
|  | 30.00 mg (BID) | 1                    | 0/86, 0.00%  | 2/186, 1.07%   |
|  | 60.00 mg (BID) | 1                    | 0/91, 0.00%  | 2/186, 1.07%   |
| TEAEs leading to treatment discontinuation | 10.00 mg (QD)  | 1                    | 9/221, 4.10%   | 6/222, 2.70%   |
|  | 30.00 mg (QD)  | 1                    | 4/228, 1.80%   | 6/222, 2.70%   |
|  | 60.00 mg (QD)  | 2                    | 9/387, 2.33%   | 8/379, 2.11%   |
| Constipation                               | 10.00 mg (QD)  | 2                    | 19/314, 6.05%  | 5/408, 1.23%   |
|  | 30.00 mg (QD)  | 2                    | 26/411, 6.33%  | 5/408, 1.23%   |
|  | 60.00 mg (QD)  | 3                    | 41/573, 7.15%  | 9/565, 1.59%   |
|  | 30.00 mg (BID) | 1                    | 3/86, 3.49%  | 4/186, 2.15%   |
|  | 60.00 mg (BID) | 1                    | 4/91, 4.40%  | 4/186, 2.15%   |
| Nausea                                     | 10.00 mg (QD)  | 2                    | 16/314, 5.09%  | 13/408, 3.19%  |
|  | 30.00 mg (QD)  | 2                    | 23/411, 5.60%  | 13/408, 3.19%  |
|  | 60.00 mg (QD)  | 3                    | 47/573, 8.20%  | 18/565, 3.18%  |
|  | 30.00 mg (BID) | 1                    | 8/86, 9.30%  | 9/186, 4.84%   |
|  | 60.00 mg (BID) | 1                    | 9/91, 9.90%  | 9/186, 4.84%   |
| UTIs                                       | 10.00 mg (QD)  | 2                    | 5/314, 1.60%   | 12/408, 2.94%  |
|  | 30.00 mg (QD)  | 2                    | 20/411, 4.86%  | 12/408, 2.94%  |
|  | 60.00 mg (QD)  | 3                    | 18/573, 3.14%  | 16/565, 2.80%  |
|  | 30.00 mg (BID) | 1                    | 2/86, 2.33%  | 4/186, 2.15%   |
|  | 60.00 mg (BID) | 1                    | 3/91, 3.30%  | 4/186, 2.15%   |
| Nasopharyngitis                            | 10.00 mg (QD)  | 2                    | 7/314, 2.23%   | 12/408, 2.94%  |
|  | 30.00 mg (QD)  | 2                    | 19/411, 4.62%  | 12/408, 2.94%  |
|  | 60.00 mg (QD)  | 3                    | 30/573, 5.33%  | 24/565, 4.25%  |
|  | 30.00 mg (BID) | 1                    | 1/86, 1.16%  | 4/186, 2.15%   |
|  | 60.00 mg (BID) | 1                    | 3/91, 3.30%  | 4/186, 2.15%   |
| URTIs                                      | 10.00 mg (QD)  | 2                    | 15/314, 4.80%  | 25/408, 6.13%  |
|  | 30.00 mg (QD)  | 2                    | 27/411, 6.57%  | 25/408, 6.13%  |
|  | 60.00 mg (QD)  | 2                    | 19/417, 4.56%  | 25/408, 6.13%  |
|  | 30.00 mg (BID) | 1                    | 6/86, 6.97%  | 15/186, 8.06%  |
|  | 60.00 mg (BID) | 1                    | 6/91, 6.59%  | 15/186, 8.06%  |
| Fatigue                                    | 10.00 mg (QD)  | 2                    | 4/314, 1.27%   | 10/408, 2.45%  |
|  | 30.00 mg (QD)  | 2                    | 10/411, 2.43%  | 10/408, 2.45%  |

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**Supplementary Table 1. Continued**

| Adverse events                             | Dose           | Studies reported (n) | The incidence rate in the atogepant group (event/total, %) | The incidence rate in the placebo group (event/total, %) |
|--|----------------|----------------------|--|--|
| Increased blood creatine kinase level      | 60.00 mg (QD)  | 2                    | 14/417, 3.35%  | 10/408, 2.45%  |
|  | 30.00 mg (BID) | 1                    | 2/86, 2.33%  | 6/186, 3.23%   |
|  | 60.00 mg (BID) | 1                    | 9/91, 9.90%  | 6/186, 3.23%   |
|  | 10.00 mg (QD)  | 2                    | 9/314, 2.86%   | 5/408, 1.23%   |
|  | 30.00 mg (QD)  | 2                    | 5/411, 1.22%   | 5/408, 1.23%   |
|  | 60.00 mg (QD)  | 2                    | 9/417, 2.16%   | 5/408, 1.23%   |
|  | 30.00 mg (BID) | 1                    | 6/86, 6.97%  | 3/186, 1.60%   |
| Death                                      | 60.00 mg (BID) | 1                    | 2/91, 2.19%  | 3/186, 1.60%   |
|  | 10.00 mg (QD)  | 1                    | 0/221, 0.00%   | 0/222, 0.00%   |
|  | 30.00 mg (QD)  | 1                    | 0/228, 0.00%   | 0/222, 0.00%   |
| Somnolence                                 | 60.00 mg (QD)  | 2                    | 0/387, 0.00%   | 0/379, 0.00%   |
|  | 10.00 mg (QD)  | 1                    | 7/221, 3.20%   | 2/222, 0.90%   |
|  | 30.00 mg (QD)  | 1                    | 4/228, 1.80%   | 2/222, 0.90%   |
| Sinusitis                                  | 60.00 mg (QD)  | 1                    | 4/231, 1.70%   | 2/222, 0.90%   |
|  | 10.00 mg (QD)  | 1                    | 4/221, 1.80%   | 3/222, 1.40%   |
|  | 30.00 mg (QD)  | 1                    | 3/228, 1.30%   | 3/222, 1.40%   |
| Gastroenteritis                            | 60.00 mg (QD)  | 1                    | 5/231, 2.20%   | 3/222, 1.40%   |
|  | 10.00 mg (QD)  | 1                    | 2/221, 0.90%   | 4/222, 1.80%   |
|  | 30.00 mg (QD)  | 1                    | 5/228, 2.20%   | 4/222, 1.80%   |
| Increase alanine amino-transferase level   | 60.00 mg (QD)  | 1                    | 3/231, 1.30%   | 4/222, 1.80%   |
|  | 10.00 mg (QD)  | 1                    | 3/221, 1.40%   | 6/222, 2.70%   |
|  | 30.00 mg (QD)  | 1                    | 2/228, 0.90%   | 6/222, 2.70%   |
| Influenza                                  | 60.00 mg (QD)  | 1                    | 2/231, 0.90%   | 6/222, 2.70%   |
|  | 10.00 mg (QD)  | 1                    | 3/221, 1.40%   | 2/222, 0.90%   |
|  | 30.00 mg (QD)  | 1                    | 2/228, 0.90%   | 2/222, 0.90%   |
| Sinus congestion                           | 60.00 mg (QD)  | 1                    | 5/231, 2.20%   | 2/222, 0.90%   |
|  | 10.00 mg (QD)  | 1                    | 1/221, 0.50%   | 5/222, 2.30%   |
|  | 30.00 mg (QD)  | 1                    | 2/228, 0.90%   | 5/222, 2.30%   |
| Increased aspartate aminotransferase level | 60.00 mg (QD)  | 1                    | 4/231, 1.70%   | 5/222, 2.30%   |
|  | 10.00 mg (QD)  | 1                    | 2/221, 0.90%   | 6/222, 2.70%   |
|  | 30.00 mg (QD)  | 1                    | 2/228, 0.90%   | 6/222, 2.70%   |
| Anxiety                                    | 60.00 mg (QD)  | 1                    | 1/231, 0.40%   | 6/222, 2.70%   |
|  | 10.00 mg (QD)  | 1                    | 2/221, 0.90%   | 2/222, 0.90%   |
|  | 30.00 mg (QD)  | 1                    | 1/228, 0.40%   | 2/222, 0.90%   |
|  | 60.00 mg (QD)  | 1                    | 5/231, 2.20%   | 2/222, 0.90%   |

TEAE, treatment-related treatment-emergent adverse event; SAE, serious adverse event; UTI, urinary tract infection; URTI, upper respiratory tract infection; QD, once daily; BID, twice a day.