Effectiveness and respiratory adverse events following inactivated and mRNA COVID-19 vaccines in patients with COPD and asthma: A Chinese population-based study

**Running header:** COVID vaccines in asthma and COPD: Efficacy and exacerbations **Journal name:** Drug safety

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Supplementary Table 1 ICD-9-CM codes for COPD, asthma, comorbidities, respiratory failure, and BNF codes for medications

|       | _   | ~     |       |
|-------|-----|-------|-------|
| 16 11 | _9_ | .( 'N | codes |

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| Study cohort  |   |       |  |  |  |
|---|---|-------|--|--|--|
| COPD  | 490, 491, 492, 494, 495, 496                | -     |  |  |  |
| Asthma  | 493   | -     |  |  |  |
| Respiratory failure   | 518.81-518.84                               | -     |  |  |  |
| Bronchodilators   | -   | 3.1   |  |  |  |
| Corticosteroids (respiratory)   | -   | 3.2   |  |  |  |
|   |   |       |  |  |  |
| Comorbidities   |   |       |  |  |  |
| Heart failure   | 398.91, 402.01, 402.11, 402.91, 404.01,     | -     |  |  |  |
|   | 404.03, 404.11, 404.13, 404.91, 404.93, 428 |       |  |  |  |
| Hypertension  | 401-405, 437.2                              | -     |  |  |  |
| Diabetes  | 250   | -     |  |  |  |
| Stroke  | 430-438                                     | -     |  |  |  |
| IHD   | 410-414                                     | -     |  |  |  |
| PVD   | 441, 443.9, V43.4                           | -     |  |  |  |
| Atrial fibrillation   | 427.3, 427.4                                | -     |  |  |  |
| Renal failure   | 580-589                                     | -     |  |  |  |
| Ulcer disease   | 531-534                                     | -     |  |  |  |
| Liver disease   | 570-573, 456.0, 456.1, 456.2                | -     |  |  |  |
| Hip fracture  | 820   | -     |  |  |  |
| Parkinson disease   | 332.0                                       | -     |  |  |  |
| Medications   |   |       |  |  |  |
| Diuretics   | -   | 2.2   |  |  |  |
| Anti-arrhythmic drugs   | -   | 2.3   |  |  |  |
| Beta-blockers   | -   | 2.4   |  |  |  |
| Hypertension and heart  | -   | 2.5   |  |  |  |
| failure drugs   |   |       |  |  |  |
| Calcium channel blockers,   | -   | 2.6   |  |  |  |
| nitrates  |   |       |  |  |  |
| Anticoagulants  | -   | 2.8   |  |  |  |
| Antiplatelet drugs  | -   | 2.9   |  |  |  |
| Stable angina, ACS drugs  | -   | 2.10  |  |  |  |
| Lipid-lowering drugs  | -   | 2.12  |  |  |  |
| Antihistamines  | -   | 3.4   |  |  |  |
| Antipsychotics drugs  | -   | 4.2   |  |  |  |
| Antidepressant drugs  | -   | 4.3   |  |  |  |
| Antiepileptic drugs   | -   | 4.8   |  |  |  |
| Anti-Parkinson drugs  | _   | 4.9   |  |  |  |
| Anti-dementia drugs   | _   | 4.11  |  |  |  |
| Antibiotics   | _   | 5.1   |  |  |  |
| Antiviral   | _   | 5.3   |  |  |  |
| Antidiabetic drugs  | _   | 6.1   |  |  |  |
| Systematic steroids   | _   | 6.3.2 |  |  |  |
| Abbraviations: ICD: international coding disease: COPD: chronic obstructive nulmonary disease: RNE: |   |       |  |  |  |

Abbreviations: ICD: international coding disease; COPD: chronic obstructive pulmonary disease; BNF: British National formula; IHD: Ischaemic heart disease; PVD: peripheral vascular disease; ACS: acute coronary syndrome

Supplementary Table 2 Vaccine effectiveness of fourth doses of CoronaVac and BNT162b2 compared to non-vaccination on COVID-related death, hospitalisation, and severe complications

|                               | Total number of people | Number of events | IPTW-adjusted HR (95% CI) | VE (95% CI)                                  |
|-------------------------------|------------------------|------------------|---------------------------|--|
| COVID-related mortality       |                        |                  | (> )                      |  |
| Unvaccinated                  | 24624                  | 1077 (4.37)      | 1.00                      | 1.00   |
| CoronaVac                     | 10563                  | 4 (0.04)         | 0.03 (0.008, 0.1)         | 99.7% (90%, 99.92%)                          |
| BNT162b2                      | 6709                   | 0                | NA                        | NA   |
| C-C-B-B                       | 2344                   | 1 (0.04)         | 0.02(0.008, 0.07)         | 99.8% (99.3%, 99.92%)                        |
| B-B-C-C                       | 64                     | 0                | NA                        | NA   |
| B-B-C-B                       | 29                     | 0                | NA                        | NA   |
| C-C-B-C                       | 398                    | 0                | NA                        | NA   |
| C-C-B                         | 677                    | 0                | NA                        | NA   |
| B-B-B-C                       | 133                    | 1 (0.75)         | 0.83 (0.50, 1.39)         | 17% (-39%, 50%)                              |
| COVID-related hospitalisation |                        | , ,              | , ,                       | <b>,</b> , , , , , , , , , , , , , , , , , , |
| Unvaccinated                  | 24624                  | 1953 (7.93)      | 1.00                      | 1.00   |
| CoronaVac                     | 10563                  | 159 (1.51)       | 0.58(0.47, 0.72)          | 42% (28%, 53%)                               |
| BNT162b2                      | 6709                   | 67 (1.00)        | 0.42 (0.34, 0.53)         | 58% (47%, 66%)                               |
| C-C-B-B                       | 2344                   | 19 (0.81)        | 0.28 (0.22, 0.36)         | 72% (64%, 78%)                               |
| B-B-C-C                       | 64                     | 1 (1.56)         | 0.72 (0.44, 1.18)         | 28% (-18%, 56%)                              |
| B-B-C-B                       | 29                     | 1 (3.45)         | 1.32 (0.81, 2.16)         | -32% (-116%, 19%)                            |
| C-C-B-C                       | 398                    | 7 (1.76)         | 0.65 (0.53, 0.81)         | 35% (19%, 47%)                               |
| C-C-B                         | 677                    | 10 (1.48)        | 0.43 (0.34, 0.53)         | 57% (47%, 66%)                               |
| B-B-B-C                       | 133                    | 1 (0.75)         | 0.33 (0.20, 0.54)         | 67% (46%, 80%)                               |
| COVID-related severe          |                        |                  |                           |  |
| complications                 |                        |                  |                           |  |
| Unvaccinated                  | 24624                  | 165 (0.67)       | 1.00                      | 1.00   |
| CoronaVac                     | 10563                  | 4 (0.04)         | 0.18 (0.05, 0.73)         | 82% (27%, 99.5%)                             |
| BNT162b2                      | 6709                   | 2 (0.03)         | 0.75(0.37, 1.52)          | 25% (-52%, 63%)                              |
| C-C-B-B                       | 2344                   | 0                | NA                        | NA   |
| B-B-C-C                       | 64                     | 0                | NA                        | NA   |
| B-B-C-B                       | 29                     | 0                | NA                        | NA   |
| C-C-B-C                       | 398                    | 0                | NA                        | NA   |
| C-C-B                         | 677                    | 1 (0.15)         | 0.88 (0.48, 1.59)         | 12% (-59%, 52%)                              |
| B-B-B-C                       | 133                    | 1 (0.75)         | 1.02 (0.88, 1.38)         | -2% (-38%, 12%)                              |

Abbreviations: C-C-B-B: CoronaVac- CoronaVac-BNT162b2-BNT162b2; B-B-C-C: BNT162b2-BNT162b2- CoronaVac- CoronaVac; B-B-C-B: BNT162b2-BNT162b2- CoronaVac-bnt162B2; C-C-B-C: CoronaVac-CoronaVac-BNT162b2-CoronaVac; C-C-C-B: CoronaVac- CoronaVac- CoronaVac-BNT162b2-BNT162b2-BNT162b2-CoronaVac.

## Supplementary Table 3 Subgroup analysis for estimating the association of 28-day respiratory exacerbation after CoronaVac vaccination

|                  | Number of events | Total person-<br>years | Incidence rate (95% CI)/person-years) | Adjusted IRR (95% CI) |
|------------------|------------------|------------------------|---------------------------------------|-----------------------|
| Male             |                  |                        |                                       |                       |
| Baseline period  | 3634             | 6352.35                | 0.57 (0.56, 0.58)                     | 1.00                  |
| First dose       |                  |                        |                                       |                       |
| Day <b>0</b> -28 | 86               | 171.68                 | 0.50(0.43, 0.58)                      | 1.15 (0.92, 1.44)     |
| Day <b>0</b> -7  | 28               | 49.30                  | 0.57(0.45, 0.72)                      | 1.30 (0.89, 1.91)     |
| Day 8-28         | 58               | 122.37                 | 0.47(0.39, 0.57)                      | 1.08 (0.83, 1.42)     |
| Second dose      |                  |                        | ,                                     |                       |

|                  |      | 1.60.10 | 0.47 (0.00 0.70)      | 1.02 (0.00 1.01)   |
|------------------|------|---------|-----------------------|--------------------|
| Day <b>0</b> -28 | 73   | 162.48  | 0.45 (0.38, 0.53)     | 1.03 (0.80, 1.31)  |
| Day <b>0-</b> 7  | 23   | 46.52   | 0.49 (0.37, 0.66)     | 1.13 (0.75, 1.72)  |
| Day 8-28         | 50   | 115.95  | $0.43 \ (0.35, 0.53)$ | 0.98 (0.74, 1.31)  |
| Third dose       |      |         |                       |                    |
| Day <b>0</b> -28 | 59   | 134.37  | 0.44 (0.36, 0.53)     | 1.1 (0.84, 1.50)   |
| <b>Day 0-7</b>   | 10   | 38.83   | 0.26 (0.15, 0.44)     | 0.67 (0.36, 1.25)  |
| Day 8-28         | 49   | 95.54   | 0.51 (0.42, 0.62)     | 1.32 (0.97, 1.80)  |
| Fourth dose      |      |         |                       |                    |
| Day <b>0</b> -28 | 11   | 22.00   | 0.50(0.33, 0.76)      | 1.33 (0.88, 1.78)  |
| Day <b>0</b> -7  | 2    | 6.57    | 0.30(0.10, 0.97)      | 1.46 (0.36, 5.83)  |
| Day 8-28         | 9    | 15.43   | 0.58 (0.38, 0.89)     | 1.17 (0.91, 1.43)  |
| Female           |      |         |                       |                    |
| Baseline period  | 2031 | 3599.51 | 0.56 (0.55, 0.58)     | 1.00               |
| First dose       |      |         | ( , )                 |                    |
| Day 0-28         | 36   | 104.38  | 0.34 (0.26, 0.45)     | 0.72 (0.51, 1.02)  |
| Day 0-7          | 6    | 30.04   | 0.20 (0.10, 0.41)     | 0.42 (0.19, 0.95)  |
| Day 8-28         | 30   | 74.34   | 0.40 (0.31, 0.53)     | 0.85 (0.58, 1.23)  |
| Second dose      | 30   | / 1.54  | 0.10 (0.51, 0.55)     | 0.05 (0.50, 1.25)  |
| Day 0-28         | 39   | 100.84  | 0.39 (0.30, 0.49)     | 0.79 (0.56, 1.09)  |
| Day 0-28         | 12   | 28.89   | 0.42 (0.27, 0.64)     | 0.85 (0.48, 1.50)  |
| Day 8-28         | 27   | 71.95   | 0.38 (0.28, 0.51)     | 0.77 (0.52, 1.14)  |
| Third dose       | 21   | 71.93   | 0.36 (0.26, 0.31)     | 0.77(0.32, 1.14)   |
|                  | 35   | 82.73   | 0.42 (0.22, 0.54)     | 0.02 (0.64, 1.22)  |
| Day 0-28         |      |         | 0.42 (0.33, 0.54)     | 0.92 (0.64, 1.32)  |
| Day 0-7          | 9    | 23.91   | 0.38 (0.22 0.63)      | 0.83 (0.43, 1.62)  |
| Day 8-28         | 26   | 58.82   | 0.44 (0.33, 0.59)     | 097 (0.64, 1.45)   |
| Fourth dose      | 4    | 10.17   | 0.22 (0.15, 0.72)     | 0.72 (0.25.2.10)   |
| Day <b>0</b> -28 | 4    | 12.17   | 0.33 (0.15, 0.73)     | 0.73 (0.25, 2.19)  |
| Day <b>0-</b> 7  | 1    | 3.67    | 0.27 (0.05, 1.45)     | 0.65 (0.09, 4.67)  |
| Day 8-28         | 3    | 8.49    | 0.35 (0.14, 0.88)     | 0.83 (0.23, 2.94)  |
| Age<65           | 0=0  |         | 0.00 (0.00 0.00)      | 4.00               |
| Baseline period  | 878  | 1516.60 | 0.58 (0.55, 0.60)     | 1.00               |
| First dose       |      |         |                       |                    |
| Day 0-28         | 29   | 56.87   | $0.51\ (0.40,0.66)$   | 1.23 (0.83, 1.81)  |
| Day 0-7          | 11   | 16.25   | 0.68 (0.48, 0.95)     | 1.62 (0.88, 2.96)  |
| Day 8-28         | 18   | 40.62   | 0.44(0.31, 0.63)      | 1.06 (0.65, 1.72)  |
| Second dose      |      |         |                       |                    |
| Day 0-28         | 19   | 55.84   | 0.34 (0.24, 0.49)     | 0.82 (0.51, 1.32)  |
| Day 0-7          | 8    | 15.97   | 0.50(0.31, 0.82)      | 1.21 (0.59, 2.45)  |
| Day 8-28         | 11   | 39.87   | 0.28 (0.17, 0.46)     | 0.66 (0.36, 1.21)  |
| Third dose       |      |         |                       |                    |
| Day 0-28         | 24   | 48.94   | 0.49(0.37, 0.65)      | 1.35 (0.86, 2.11)  |
| Day 0-7          | 3    | 14.09   | 0.21 (0.08, 0.58)     | 0.59 (0.19, 1.86)  |
| Day 8-28         | 21   | 34.84   | 0.60 (0.46, 0.79)     | 1.68 (0.88, 2.48)  |
| Fourth dose      |      |         | · , , ,               |                    |
| Day 0-28         | 1    | 6.34    | 0.16 (0.03, 0.95)     | 1.44 (0.15, 13.50) |
| Day 0-7          | 1    | 1.86    | 0.54 (0.14, 2.04)     | 5.69 (0.61, 52.81) |
| Day 8-28         | 0    | NA      | NA                    | NA                 |
| $Age \ge 65$     | ~    | T // T  | T 17 T                | - 12 E             |
| Baseline period  | 4787 | 8335.26 | 0.57 (0.56, 0.59)     | 1.00               |
| First dose       | 1707 | 0333.20 | 0.57 (0.50, 0.57)     | 1.00               |
| 1 11 51 11050    |      |         |                       |                    |

| Day 0-28 93 219.19 0.42 (0.36, 0.50) 0.92 (0.74   | , 1.14)  |
|---|----------|
| Day 0-7 23 63.09 0.36 (0.26, 0.50) 0.79 (0.52   | , 1.20)  |
| Day 8-28 70 156.10 0.45 (0.38, 0.53) 0.97 (0.76   | , 1.24)  |
| Second dose   |          |
| Day 0-28 93 207.48 0.45 (0.39, 0.52) 0.95 (0.77   | (1.19)   |
| Day 0-7 27 59.44 0.45 (0.34, 0.60) 0.97 (0.66   | (1.43)   |
| Day 8-28 66 148.03 0.45 (0.37, 0.53) 0.95 (0.74   |          |
| Third dose  | , ,      |
| Day 0-28 70 168.16 0.42 (0.35, 0.50) 0.96 (0.74   | , 1.24)  |
| Day 0-7 16 48.65 0.33 (0.22, 0.49) 0.77 (0.47   |          |
| Day 8-28 54 78.13 0.69 (0.60, 0.80) 1.05 (0.79  |          |
| Fourth dose   | , -,     |
| Day 0-28 14 27.83 0.50 (0.35, 0.73) 1.54 (0.81  | 2.91)    |
| Day 0-7 2 8.38 0.24 (0.07, 0.80) 0.77 (0.20   |          |
| Day 8-28 12 19.44 0.62 (0.43, 0.88) 1.98 (0.99  | . ,      |
| People with COPD  | , 5.75)  |
| Baseline period 3078 5353.59 0.57 (0.56, 0.59) 1.00   | )        |
| First dose  | ,        |
| Day 0-28 57 135.36 0.42 (0.35, 0.51) 1.02 (0.77   | 1 3/1)   |
| Day 0-28 37 133.30 0.42 (0.33, 0.31) 1.02 (0.77) Day 0-7 11 38.95 0.28 (0.17, 0.47) 0.68 (0.38) |          |
| Day 8-28 46 96.40 0.48 (0.39, 0.59) 1.15 (0.85  |          |
| Second dose 90.40 90.40 0.46 (0.39, 0.39) 1.13 (0.83  | , 1.50)  |
| Day 0-28 58 128.19 0.45 (0.37, 0.55) 1.09 (0.82   | 1.44)    |
| Day 0-28 36 128.19 0.43 (0.37, 0.33) 1.09 (0.82   |          |
|   |          |
|   | , 1.34)  |
| Third dose  Dev. 0.28 (0.20, 0.40)  | 1 20)    |
| Day 0-28 39 102.52 0.38 (0.30, 0.49) 0.95 (0.66   | . ,      |
| Day 0-7 6 29.68 0.20 (0.10, 0.41) 0.52 (0.23  |          |
| Day 8-28 33 72.84 0.45 (0.35, 0.58) 1.16 (0.79  | (, 1.72) |
| Fourth dose   | 4 17)    |
| Day 0-28 6 16.11 0.37 (0.20, 0.70) 1.49 (0.53   |          |
| Day 0-7 0 NA NA NA  |          |
| Day 8-28 6 11.17 0.54 (0.31, 0.93) 2.09 (0.75   | , 5.83)  |
| People with asthma  |          |
| Baseline period 1030 1786.81 0.58 (0.55, 0.60) 1.00   | )        |
| First dose  | 4 = 6    |
| Day 0-28 24 56.06 0.43 (0.32, 0.58) 1.14 (0.74  |          |
| Day 0-7 8 16.08 0.50 (0.30, 0.81) 1.32 (0.65  |          |
| Day 8-28 16 39.98 0.40 (0.27, 0.58) 1.06 (0.63  | , 1.78)  |
| Second dose   |          |
| Day 0-28 23 53.98 0.43 (0.31, 0.58) 1.09 (0.71  |          |
| Day 0-7 10 15.47 0.65 (0.45, 0.93) 1.65 (0.87   |          |
| Day 8-28 13 38.50 0.34 (0.22, 0.53) 0.86 (0.49)   | , 1.49)  |
| Third dose  |          |
| Day 0-28 20 45.12 0.44 (0.32, 0.61) 1.21 (0.73  |          |
| Day 0-7 6 13.05 0.46 (0.26, 0.83) 1.28 (0.55  |          |
| Day 8-28 14 32.07 0.44 (0.29, 0.65) 1.20 (0.68  | , 2.13)  |
| Fourth dose   |          |
| Day 0-28 3 6.56 0.46 (0.20, 1.05) 1.65 (0.48  | 5.72     |
| Day 0-7 1 1.94 0.52 (0.13, 2.02) 1.98 (0.23,  | 16.93)   |

1.48 (0.37, 5.81)

Day 8-28 2 4.62 0.43 (0.15, 1.23) 1.48 (0.3 Abbreviations: N, number; IRR, incidence rate ratio; CI, confidence interval; NA, nonapplicable

## Supplementary Table 4 Subgroup analysis for estimating the association of 28-day respiratory exacerbation after BNT162b2 vaccination $^1$

|                  | Number of events | Total person-<br>years | Incidence rate (95% CI)/person-years) | Adjusted IRR (95% CI) |
|------------------|------------------|------------------------|---------------------------------------|-----------------------|
| Male             | CVCIIIS          | y cuis                 | CI), person years)                    | CI)                   |
| Baseline period  | 2673             | 4714.95                | 0.57 (0.55, 0.58)                     | 1.00                  |
| First dose       | 2013             | 1711.55                | 0.57 (0.55, 0.50)                     | 1.00                  |
| Day 0-28         | 39               | 75.29                  | 0.52 (0.42, 0.64)                     | 1.06 (0.77, 1.47)     |
| Day 0-20 Day 0-7 | 9                | 26.17                  | 0.34 (0.20, 0.58)                     | 0.70 (0.36, 1.37)     |
| Day 8-28         | 30               | 49.12                  | 0.61 (0.49, 0.76)                     | 1.25 (0.87, 1.81)     |
| Second dose      | 30               | 49.12                  | 0.01 (0.49, 0.70)                     | 1.23 (0.67, 1.61)     |
| Day 0-28         | 51               | 88.96                  | 0.57 (0.48, 0.69)                     | 1.17 (0.88, 1.57)     |
| •                |                  |                        |                                       |                       |
| Day 0-7          | 13               | 25.44                  | 0.51 (0.35, 0.75)                     | 1.05 (0.60, 1.82)     |
| Day 8-28         | 38               | 63.52                  | 0.60 (0.49, 0.73)                     | 1.21 (0.87, 1.69)     |
| Third dose       | 2.5              | <b>55</b> 0.5          | 0.45 (0.26 0.50)                      | 1 05 (0 54 1 54)      |
| Day 0-28         | 35               | 77.05                  | 0.45 (0.36, 0.58)                     | 1.07 (0.74, 1.54)     |
| Day 0-7          | 9                | 22.08                  | 0.41 (0.25, 0.67)                     | 0.97 (0.49, 1.90)     |
| Day 8-28         | 26               | 54.97                  | 0.47 (0.36, 0.63)                     | 1.11 (0.74, 1.67)     |
| Fourth dose      |                  |                        |                                       |                       |
| Day 0-28         | 7                | 12.50                  | 0.56 (0.34, 0.92)                     | 1.53 (0.69, 3.39)     |
| Day 0-7          | 4                | 3.66                   | 1.09 (0.91, 1.17)                     | 2.11 (0.88, 3.34)     |
| Day 8-28         | 3                | 8.84                   | 0.34 (0.14, 0.85)                     | 1.05 (0.35, 3.19)     |
| Female           |                  |                        |                                       |                       |
| Baseline period  | 1595             | 2789.94                | 0.57 (0.55, 0.59)                     | 1.00                  |
| First dose       |                  |                        | , , ,                                 |                       |
| Day 0-28         | 26               | 58.04                  | 0.45 (0.34, 0.60)                     | 0.94 (0.63, 1.40)     |
| Day 0-7          | 10               | 20.00                  | 0.50 (0.32, 0.78)                     | 1.05 (0.56, 1.97)     |
| Day 8-28         | 16               | 38.05                  | 0.42 (0.29, 0.61)                     | 0.88 (0.53, 1.44)     |
| Second dose      | - 0              |                        | (**->, ***-)                          | (,)                   |
| Day 0-28         | 42               | 68.33                  | 0.61 (0.51, 0.74)                     | 1.28 (0.93, 1.77)     |
| Day 0-7          | 11               | 19.54                  | 0.56 (0.38, 0.83)                     | 1.17 (0.64, 2.14)     |
| Day 8-28         | 31               | 48.79                  | 0.64 (0.51, 0.79)                     | 1.32 (0.91, 1.91)     |
| Third dose       | 31               | 70.77                  | 0.04 (0.31, 0.77)                     | 1.32(0.71, 1.71)      |
|                  | 21               | 60.43                  | 0.25 (0.25, 0.40)                     | 0.96 (0.55, 1.22)     |
| Day 0-28         |                  |                        | 0.35 (0.25, 0.49)                     | 0.86 (0.55, 1.33)     |
| Day 0-7          | 7                | 17.35                  | 0.40 (0.23, 0.71)                     | 0.99 (0.46, 2.11)     |
| Day 8-28         | 14               | 43.08                  | 0.32 (0.21, 0.50)                     | 0.80 (0.47, 1.36)     |
| Fourth dose      |                  |                        |                                       | <b>3</b> 7.4          |
| Day 0-28         | 0                | 37.                    | 37.                                   | NA                    |
| Day 0-7          | 0                | NA                     | NA                                    | NA                    |
| Day 8-28         | 0                | NA                     | NA                                    | NA                    |
| Age<65years      |                  |                        |                                       |                       |
| Baseline period  | 1171             | 2061.14                | 0.57 (0.55, 0.59)                     | 1.00                  |
| First dose       |                  |                        |                                       |                       |
| Day 0-28         | 34               | 68.72                  | 0.49(0.39, 0.63)                      | 1.03 (0.73, 1.46)     |
| Day 0-7          | 12               | 23.76                  | 0.51 (0.34, 0.75)                     | 1.05 (0.59, 1.87)     |
| Day 8-28         | 22               | 44.96                  | 0.49(0.36, 0.66)                      | 1.02 (0.67, 1.56)     |
| Second dose      |                  |                        | •                                     |                       |
| Day 0-28         | 58               | 82.05                  | 0.71 (0.61, 0.81)                     | 1.44 (0.75, 2.13)     |
| Day 0-7          | 14               | 23.46                  | 0.60(0.43, 0.83)                      | 1.24 (0.73, 2.11)     |

| Day 8-28                      | 44             | 58.59               | 0.74 (0.65, 0.87)        | 1.55 (0.78, 2.32)                   |
|-------------------------------|----------------|---------------------|--------------------------|-------------------------------------|
| Third dose                    |                |                     |                          |                                     |
| Day 0-28                      | 28             | 73.79               | 0.38 (0.28, 0.51)        | 0.90 (0.61, 1.33)                   |
| Day 0-7                       | 10             | 21.13               | 0.47 (0.30, 0.74)        | 1.12 (0.58, 2.13)                   |
| Day 8-28                      | 18             | 52.65               | 0.34 (0.24, 0.50)        | 0.80 (0.50, 1.29)                   |
| Fourth dose                   |                |                     |                          |                                     |
| Day 0-28                      | 3              | 6.49                | 0.46 (0.20, 1.06)        | 1.53 (0.41, 5.80)                   |
| Day 0-7                       | 2              | 1.90                | 1.05 (0.92, 1.18)        | 3.43 (0.67, 17.68)                  |
| Day 8-28                      | 1              | 4.58                | 0.22 (0.04, 1.24)        | 0.66 (0.10, 4.66)                   |
| $Age \ge 65years$             |                |                     | ( , , ,                  | ( , , ,                             |
| Baseline period               | 3097           | 5443.75             | 0.57 (0.56, 0.58)        | 1.00                                |
| First dose                    |                |                     | ( , )                    |                                     |
| Day 0-28                      |                |                     |                          | 0.98 (0.68, 1.41)                   |
| Day 0-7                       | 7              | 22.41               | 0.31 (0.17, 0.58)        | 0.64 (0.30, 1.36)                   |
| Day 8-28                      | 24             | 42.21               | 0.57 (0.44, 0.74)        | 1.16 (0.77, 1.75)                   |
| Second dose                   | 2.             | 12.21               | 0.57 (0.11, 0.71)        | 1110 (0177, 1170)                   |
| Day 0-28                      |                |                     |                          | 0.94 (0.66, 1.34)                   |
| Day 0-7                       | 10             | 21.52               | 0.46 (0.30, 0.73)        | 0.95 (0.50, 1.78)                   |
| Day 8-28                      | 25             | 53.72               | 0.47 (0.35, 0.62)        | 0.94 (0.63, 1.41)                   |
| Third dose                    | 23             | 33.12               | 0.47 (0.55, 0.02)        | 0.54 (0.05, 1.41)                   |
| Day 0-28                      |                |                     |                          | 1.06 (0.71, 1.58)                   |
| Day 0-7                       | 6              | 18.30               | 0.33 (0.17, 0.63)        | 0.80 (0.35, 1.80)                   |
| Day 8-28                      | 22             | 45.39               | 0.48 (0.36, 0.65)        | 1.17 (0.75, 1.82)                   |
| Fourth dose                   | 22             | 43.37               | 0.40 (0.50, 0.05)        | 1.17 (0.75, 1.02)                   |
| Day 0-28                      |                |                     |                          | 0.86 (0.31, 2.36)                   |
| Day 0-28                      | 2              | 3.64                | 0.55 (0.22, 1.39)        | 1.58 (0.42, 5.88)                   |
| Day 8-28                      | 2              | 8.67                | 0.23 (0.07, 0.78)        | 0.74 (0.18, 3.00)                   |
| People with COPD              | 2              | 0.07                | 0.23 (0.07, 0.70)        | 0.74 (0.10, 5.00)                   |
| Baseline period               | 2031           | 3587.39             | 0.57 (0.55, 0.58)        | 1.00                                |
| First dose                    | 2031           | 3301.37             | 0.57 (0.55, 0.56)        | 1.00                                |
| Day 0-28                      | 22             | 41.08               | 0.54 (0.40, 0.71)        | 1.23 (0.79, 1.91)                   |
| Day 0-28                      | 7              | 14.13               | 0.50 (0.29, 0.84)        | 1.15 (0.54, 2.44)                   |
| Day 8-28                      | 15             | 26.95               | 0.56 (0.40, 0.78)        | 1.13 (0.34, 2.44) 1.27 (0.75, 2.14) |
| Second dose                   | 13             | 20.93               | 0.30 (0.40, 0.76)        | 1.27(0.75, 2.14)                    |
|                               | 29             | 47.62               | 0.56 (0.49, 0.76)        | 1.41 (0.95, 2.08)                   |
| Day 0-28                      |                | 47.62               | , , ,                    |                                     |
| Day 0-7                       | 6              | 13.62               | 0.44 (0.24, 0.80)        | 1.02 (0.45, 2.30)                   |
| Day 8-28                      | 23             | 34.00               | 0.68 (0.54, 0.85)        | 1.55 (0.98, 2.41)                   |
| Third dose                    | 1.5            | 20.71               | 0.20 (0.25, 0.56)        | 1 00 (0 57, 1 72)                   |
| Day 0-28                      | 15             | 39.71               | 0.38 (0.25, 0.56)        | 1.00 (0.57, 1.73)                   |
| Day 0-7                       | 6              | 11.38               | 0.57 (0.30, 0.91)        | 1.39 (0.60, 3.24)                   |
| Day 8-28                      | 9              | 28.33               | 0.32 (0.19, 0.54)        | 0.83 (0.42, 1.63)                   |
| Fourth dose                   | 4              | <b>7</b> 00         | 0.60.60.20.1.10          | 2 11 (0 00 10 01)                   |
| Day 0-28                      | 4              | 5.89                | 0.68 (0.39, 1.18)        | 3.11 (0.89, 10.81)                  |
| Day 0-7                       | 3              | 1.79                | 1.68 (1.25, 2.11)        | 8.54 (0.77, 16.31)                  |
| Day 8-28                      | 1              | 4.10                | 0.24 (0.04, 1.34)        | 1.49 (0.20, 11.28)                  |
| <sup>1</sup> No cases after 2 | X days followi | ng each dose in the | e asthma group. Abbrevia | tions: N. niimber:                  |

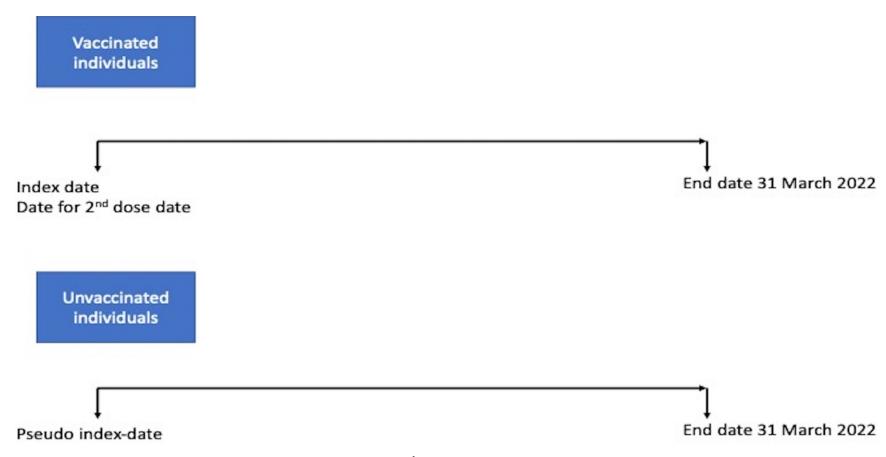
<sup>&</sup>lt;sup>1</sup> No cases after 28 days following each dose in the asthma group. Abbreviations: N, number; IRR, incidence rate ratio; CI, confidence interval; NA, non-applicable.

## Supplementary Table 5 Subgroup analysis for the association of two doses of CoronaVac and BNT162b2 on the risk of COVID-related mortality, COVID-related hospitalisation, and COVID-related severe complications

|                     | Total number of | Number of        | IPTW-adjusted           | VE (95% CI)    |
|---------------------|-----------------|------------------|-------------------------|----------------|
|                     | people          | events           | HR (95% CI)             |                |
| Male                | C               | OVID-related mo  | ortality                |                |
| Unvaccinated        | 12938           | 685 (5.29)       | 1.00                    | 1.00           |
| CoronaVac           | 51774           | 232 (0.45)       | 0.23 (0.19, 0.27)       | 77% (73%, 81%) |
| BNT162b2            | 38502           | ` /              |                         |                |
| Female              | 38302           | 51 (0.13)        | 0.09 (0.07, 0.11)       | 91% (89%, 93%) |
| Unvaccinated        | 11686           | 421 (2.60)       | 1.00                    | 1.00           |
| CoronaVac           |                 | 431 (3.69)       | 0.18 (0.15, 0.22)       | 82% (78%, 85%) |
|                     | 56649           | 159 (0.28)       |                         |                |
| BNT162b2            | 44821           | 12 (0.03)        | 0.03 (0.02, 0.05)       | 97% (95%, 98%) |
| Age<65              | 10100           | (4 (0 (2)        | 1.00                    | 1.00           |
| Unvaccinated        | 10190           | 64 (0.63)        | 1.00                    | 1.00           |
| CoronaVac           | 51684           | 26 (0.05)        | 0.23 (0.14, 0.36)       | 77% (64%, 86%) |
| BNT162b2            | 60265           | 12 (0.02)        | 0.09 (0.05, 0.17)       | 91% (83%, 95%) |
| $Age \ge 65$        | 1.440.4         | 1050 (5.00)      | 1.00                    | 1.00           |
| Unvaccinated        | 14434           | 1052 (7.29)      | 1.00                    | 1.00           |
| CoronaVac           | 56739           | 365 (0.64)       | 0.25 (0.22, 0.29)       | 75% (71%, 78%) |
| BNT162b2            | 23058           | 51 (0.22)        | $0.08 \ (0.07, 0.10)$   | 92% (90%, 93%) |
| People with COPD    |                 | /                | 4.00                    | 4.00           |
| Unvaccinated        | 3399            | 252 (7.41)       | 1.00                    | 1.00           |
| CoronaVac           | 12344           | 83 (0.67)        | 0.26 (0.19, 0.36)       | 74% (64%, 81%) |
| BNT162b2            | 4451            | 11 (0.25)        | $0.10 \ (0.06, \ 0.15)$ | 90% (85%, 94%) |
| People with asthma  |                 |                  |                         |                |
| Unvaccinated        | 2467            | 57 (2.31)        | 1.00                    | 1.00           |
| CoronaVac           | 11640           | 21 (0.18)        | 0.19 (0.10, 0.34)       | 81% (66%, 90%) |
| BNT162b2            | 9467            | 6 (0.06)         | 0.11 (0.06, 0.21)       | 89% (79%, 94%) |
| Respiratory failure |                 |                  |                         |                |
| Unvaccinated        | 1443            | 122 (8.45)       | 1.00                    | 1.00           |
| CoronaVac           | 3021            | 29 (0.96)        | 0.31 (0.19, 0.51)       | 69% (49%, 81%) |
| BNT162b2            | 1231            | 6 (0.49)         | 0.15 (0.08, 0.27)       | 85% (73%, 92%) |
|                     | COV             | ID-related hospi | talisation              |                |
| Male                |                 |                  |                         |                |
| Unvaccinated        | 12938           | 1125 (8.70)      | 1.00                    | 1.00           |
| CoronaVac           | 51774           | 2155 (4.16)      | 0.91 (0.88, 0.94)       | 9% (6%, 12%)   |
| BNT162b2            | 38502           | 806 (2.09)       | 0.62(0.58, 0.67)        | 38% (33%, 42%) |
| Female              |                 |                  |                         |                |
| Unvaccinated        | 11686           | 828 (7.09)       | 1.00                    | 1.00           |
| CoronaVac           | 56649           | 1720 (3.40)      | 0.90(0.81, 0.99)        | 10% (1%, 19%)  |
| BNT162b2            | 44821           | 628 (1.40)       | 0.56(0.52, 0.61)        | 44% (39%, 48%) |
| Age<65              |                 | • • •            |                         | , , ,          |
| Unvaccinated        | 10190           | 248 (2.43)       | 1.00                    | 1.00           |
| CoronaVac           | 51684           | 884 (1.71)       | 0.91 (0.72, 1.12)       | 9% (-12%, 28%) |
| BNT162b2            | 60265           | 657 (1.09)       | 0.90 (0.88, 0.92)       | 10% (8%, 12%)  |
| $Age \ge 65$        |                 | <b>\</b> /       | , , ,                   | · / /          |
| Unvaccinated        | 14434           | 1705 (11.80)     | 1.00                    | 1.00           |

| CoronaVac           | 56739 | 2991 (5.27)                | 0.72 (0.62, 0.83)                     | 28% (17%, 38%)                               |
|---------------------|-------|----------------------------|---------------------------------------|--|
| BNT162b2            | 23058 | 777 (3.37)                 | 0.67 (0.62, 0.71)                     | 33% (29%, 38%)                               |
| People with COPD    |       |                            |                                       |  |
| Unvaccinated        | 3399  | 422 (12.42)                | 1.00                                  | 1.00   |
| CoronaVac           | 12344 | 723 (5.86)                 | $0.73 \ (0.69, 0.80)$                 | 27% (20%, 31%)                               |
| BNT162b2            | 4451  | 150 (3.37)                 | 0.67 (0.58 0.77)                      | 33% (23%, 42%)                               |
| People with asthma  |       |                            |                                       |  |
| Unvaccinated        | 2467  | 137 (5.55)                 | 1.00                                  | 1.00   |
| CoronaVac           | 11640 | 349 (3.00)                 | 0.82(0.70, 0.96)                      | 18% (4%, 30%)                                |
| BNT162b2            | 9467  | 147 (1.55)                 | 0.71 (0.58, 0.86)                     | 29% (14%, 44%)                               |
| Respiratory failure |       |                            |                                       |  |
| Unvaccinated        | 1443  | 194 (13.44)                | 1.00                                  | 1.00   |
| CoronaVac           | 3021  | 212 (7.02)                 | 0.78 (0.65, 0.91)                     | 22% (9%, 55%)                                |
| BNT162b2            | 1231  | 56 (4.55)                  | 0.72 (0.56, 0.92)                     | 28% (8%, 44%)                                |
|                     | COVI  | <b>D-</b> related severe c |                                       | <b>,</b> , ,                                 |
| Male                |       |                            | •                                     |  |
| Unvaccinated        | 12938 | 109 (0.84)                 | 1.00                                  | 1.00   |
| CoronaVac           | 51774 | 129 (0.25)                 | 0.71 (0.55, 0.92)                     | 29% (8%, 45%)                                |
| BNT162b2            | 38502 | 45 (0.12)                  | 0.38 (0.28, 0.51)                     | 62% (49%, 72%)                               |
| Female              |       |                            |                                       |  |
| Unvaccinated        | 11686 | 56 (0.48)                  | 1.00                                  | 1.00   |
| CoronaVac           | 56649 | 54 (0.10)                  | 0.57 (0.38, 0.86)                     | 43% (14%, 72%)                               |
| BNT162b2            | 44821 | 16 (12.7)                  | 0.33 (0.22, 0.52)                     | 67% (48%, 78%)                               |
| Age<65              |       | •                          | , , , , , , , , , , , , , , , , , , , | ,  |
| Unvaccinated        | 10190 | 23 (0.23)                  | 1.00                                  | 1.00   |
| CoronaVac           | 51684 | 36 (0.07)                  | 0.83(0.72, 0.94)                      | 17% (6%, 28%)                                |
| BNT162b2            | 60265 | 24 (0.04)                  | 0.53 (0.33, 0.85)                     | 47% (15%, 67%)                               |
| $Age \ge 65$        |       |                            |                                       |  |
| Unvaccinated        | 14434 | 142 (0.98)                 | 1.00                                  | 1.00   |
| CoronaVac           | 56739 | 147 (0.26)                 | 0.73(0.57, 0.95)                      | 27% (5%, 43%)                                |
| BNT162b2            | 23058 | 37 (0.16)                  | 0.42 (0.32, 0.56)                     | 58% (44%, 68%)                               |
| People with COPD    |       | ` ,                        | , ,                                   | <b>,</b> , , , , , , , , , , , , , , , , , , |
| Unvaccinated        | 3399  | 29 (0.85)                  | 1.00                                  | 1.00   |
| CoronaVac           | 12344 | 31 (0.25)                  | 0.70 (0.51, 0.89)                     | 30% (11%, 49%)                               |
| BNT162b2            | 4451  | 11 (0.25)                  | 0.67 (0.39, 0.91)                     | 33% (9%, 61%)                                |
| People with asthma  |       | ,                          |                                       |  |
| Unvaccinated        | 2467  | 13 (0.53)                  | 1.00                                  | 1.00   |
| CoronaVac           | 11640 | 11 (0.09)                  | 0.43 (0.19, 0.99)                     | 67% (1%, 81%)                                |
| BNT162b2            | 9467  | 5 (0.05)                   | 0.21 (0.08, 0.58)                     | 79% (42%, 92%)                               |
| Respiratory failure |       | ` '                        |                                       | , , ,  |
| Unvaccinated        | 1443  | 25 (1.73)                  | 1.00                                  | 1.00   |
| CoronaVac           | 3021  | 8 (0.26)                   | 0.41 (0.16, 0.67)                     | 59% (33%, 84%)                               |
| BNT162b2            | 1231  | 5 (0.41)                   | 0.54 (0.24, 0.86)                     | 46% (14%, 76%)                               |

Abbreviations: N, number; IPTW, inverse probability treatment weight; HR, hazard ratio; VE, vaccine effectiveness; CI, confidence interval; COPD, chronic obstructive pulmonary disease; COVID, coronavirus disease; NA, non-applicable.



Supplementary Figure 1: Study design for the cohort study.<sup>1</sup>

<sup>1</sup>Pseudo index date for unvaccinated individuals is defined as follows: Randomly selected unvaccinated individuals with the same age and sex were matched to each vaccinated individual, assigning the second dose vaccination date from vaccinated individuals to matched unvaccinated individuals, serving as the pseudo-index date for the unvaccinated individuals. People with COPD and asthma were defined before the index date for vaccinated individuals and the pseudo-index date for unvaccinated individuals. People in the unvaccinated individuals who died before the pseudo-index date were excluded.