

Supplementary Material*

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* This supplementary material was provided by the authors to give readers further details on their article. The material was reviewed but not copyedited.

Supplement Table. Effect of Hydroxychloroquine Reported in Controlled Studies

| Name | Type | RoB | Unadjusted Absolute effect of HCQ vs. Control (95% CI) | SOE |
|--|--------|-----|---|-----|
| <i>All-Cause Mortality</i> | | | | |
| Abd-Elsalam ⁴ | RCT | H | HCQ: 6/97 vs. 5/97; RD 1.0% (-5.5% to 7.5%) | L |
| Cavalcanti ¹⁰ | | H | HCQ: 5/159 vs. 5/173; RD 0.3% (-3.4% to 3.9%) | |
| Cavalcanti ¹⁰ | | H | HCQ+AZ: 3/172 vs. 5/173; RD -1.2% (-4.3% to 2.0%) | |
| Mitjà ¹¹ | | SC | HCQ: 0/136 vs. 0/157; RD 0% (NA) | |
| Skipper ¹² | | H | HCQ: 1/201 vs. 1/194; RD -0.02% (-1.4% to 1.4%) | |
| Horby ¹³ | | SC | HCQ: 418/1561 vs. 788/3155; RD 1.8% (-0.9% to 4.5%) | |
| C-P Chen ¹⁴ | | SC | HCQ: 0/21 vs. 0/12; RD 0% (NA) | |
| J Chen ¹⁵ | | SC | HCQ: 0/15 vs. 0/15; RD 0% (NA) | |
| L Chen ¹⁶ | | H | HCQ: 0/15 vs. 0/12; RD 0% (NA) | |
| Sulaiman ⁵ | Cohort | S | HCQ: 7/1817 vs. 54/3724; RD -1.1% (-1.5% to -0.6%) † | L |
| Catteau ⁶ | | S | HCQ: 804/4542 vs. 957/3533; RD -9.4% (-11.2% to -7.5%) † | |
| Di Castelnovo ⁷ | | S | HCQ: 386/2634 vs. 190/817; RD -8.6% (-11.8% to -5.4%) † | |
| Ip ⁸ | | S | HCQ: 2/97 vs. 44/970; RD -2.5% (-5.6% to 0.6%) | |
| Kalligeros ⁹ | | S | HCQ±AZ: NA/36 vs. NA/72; aHR 0.82 (0.20 to 3.24)* | |
| Paccoud ¹⁷ | | S | HCQ: 3/38 vs. 6/46; RD -5.2% (-18.1% to 7.8%) | |
| Lecronier ¹⁸ | | C | HCQ: 9/38 vs. 9/22; RD -17.2% (-41.8% to 7.4%) | |
| Barbosa ¹⁹ | | C | HCQ: 4/31 vs. 1/32; RD 9.8% (-3.5% to 23.3%) | |
| Magagnoli ²⁰ | | S | HCQ±AZ: 27/97 vs. 18/158; RD 16.4% (6.2% to 26.6%) † | |
| Mallat ²¹ | | S | HCQ: 0/23 vs. 0/11 [0%]; RD 0% (NA) | |
| Membrillo ²² | | C | HCQ: 27/123 vs. 21/43; RD -26.9% (-43.5% to -10.3%) † | |
| Geleris ²³ | | M | HCQ±AZ: 157/811 vs. 75/565; RD 6.1% (2.2% to 10%) † | |
| Rosenberg ²⁴ | | S | HCQ±AZ: 54/271 vs. 28/221; RD 7.3% (0.8% to 13.7%) † | |
| Mahévas ²⁵ | | S | HCQ: 9/84 vs. 8/97; RD 2.5% (-6.1% to 11.1%) | |
| Ip ²⁶ | | S | HCQ: 383/1914 vs. 120/598; RD -0.1% (-3.7% to 3.6%) | |
| Sbidian ²⁷ | | M | HCQ: 111/623 vs. 830/3792; RD -4.1% (-7.4% to 0.8%) | |
| Singh ²⁸ | | S | HCQ: 104/910 vs. 109/910; RD -0.6% (-3.5% to 2.4%) | |
| Yu ²⁹ | | S | HCQ: 9/48 vs. 238/502; RD -29.7% (-40.5% to -16.8%) † | |
| Arshad ³⁰ | | S | HCQ±AZ: 162/1202 vs. 108/409; RD -12.9% (-17.6% to -8.2%) † | |
| <i>Composite of Intubation or Death</i> | | | | |
| Horby ¹³ | RCT | SC | HCQ: 388/1300 vs. 696/2623; RD 3.3% (0.3% to 6.3%) † | L |
| Geleris ²³ | Cohort | M | HCQ±AZ: 262/811 vs. 84/565; RD 17.4% (13.1% to 21.8%) † | I |
| <i>Composite of ICU Admission Within 7-Days or Death</i> | | | | |
| Paccoud ¹⁷ | Cohort | S | HCQ: [By 12 days] 13/38 vs. 16/46; RD -0.6% (-21.0% to 19.9%) | I |
| Mahévas ³¹ | | S | HCQ: 16/84 vs. 21/97; RD -2.6% (-14.3% to 9.1%) | |
| <i>ICU Admission</i> | | | | |
| Sulaiman ⁵ | Cohort | S | HCQ: 14/1817 vs. 56/3724; RD -0.7% (-1.3% to -0.2%) † | I |

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| Catteau ⁶ | | S | HCQ: 313/4539 vs. 96/3529; RD 4.2% (3.3% to 5.1%) † | |
| Ip ⁸ | | S | HCQ: 3/97 vs. 42/970; RD -1.3% (-4.9% to 2.4%) | |
| Rosenberg ²⁴ | | S | HCQ±AZ: 52/271 vs. 27/221; RD 7% (0.6% to 13.3%) † | |
| Survival without ICU Admission | | | | |
| Mahévas ²⁵ | Cohort | S | HCQ: 17/84 vs. 22/89; RD -4.5% (-16.9% to 7.9%) | I |
| Need of Mechanical Ventilation | | | | |
| Abd-Elsalam ⁴ | RCT | H | HCQ: 4/97 vs. 5/97; RD -1.0% (-6.9% to 4.9%) | |
| Cavalcanti ¹⁰ | | H | HCQ: 12/159 vs. 12/173; RD 0.6% (-5.0% to 6.2%) | L |
| Cavalcanti ¹⁰ | | H | HCQ+AZ: 19/172 vs. 12/173; RD 4.1% (-1.9% to 10.1%) | |
| Mitjà ¹¹ | | SC | HCQ: 0/136 vs. 0/157; RD 0% (NA) | |
| Horby ¹³ | | SC | HCQ: 118/1300 vs. 215/2623; RD 0.9% (-1.0% to 2.8%) | |
| Lecronier ¹⁸ | Cohort | C | HCQ: 5/38 vs. 2/22; RD 4.1% (-12.1% to 20.2%) | L |
| Magagnoli ²⁰ | | S | HCQ±AZ: 12/90 vs. 25/177; RD -0.8% (-9.5% to 7.9%) | |
| Mallat ²¹ | | S | HCQ: 0/23 vs. 0/11; RD 0% (NA) | |
| Geleris ²³ | | M | HCQ±AZ: 154/811 vs. 26/565; RD 14.4% (11.2% to 17.6%) † | |
| Rosenberg ²⁴ | | S | HCQ±AZ: 51/271 vs. 18/221; RD 10.7% (4.8% to 16.6%) † | |
| Singh ²⁸ | | S | HCQ: 46/910 vs. 57/910; RD -1.2% (-3.3% to 0.9%) | |
| Severe Disease Progression | | | | |
| J Chen ¹⁵ | RCT | SC | HCQ: 1/15 vs. 0/15; RD 6.7% (-6.0% to 19.3%) | I |
| L Chen ¹⁶ | | H | HCQ: 0/15 vs. 0/12; RD 0% (NA) | |
| Z Chen ³² | | SC | HCQ: 0/31 vs. 4/31; RD -12.9% (-24.7% to -1.1%) † | |
| Barbosa ¹⁹ | Cohort | C | HCQ: [Respiratory support level] +0.63±0.79 vs. +0.16±0.64 points; MD 0.47 (0.11 to 0.83) † | I |
| Mallat ²¹ | | S | HCQ: [High flow oxygen therapy] 0/23 vs. 0/11; RD 0% (NA) | |
| Mahévas ³¹ | | S | HCQ: [ARDS] 24/84 vs. 23/95; RD 4.4% (-8.6% to 17.3%) | |
| Hospitalization | | | | |
| Mitjà ¹¹ | RCT | SC | HCQ: 8/136 vs. 11/157; RD -1.1% (-6.7% to 4.5%) | L |
| Skipper ¹² | | H | HCQ: 4/212 vs. 10/211; RD: -2.9% (-6.3% to 0.6%) | |
| Sulaiman ⁵ | Cohort | S | HCQ: 171/1817 vs. 617/3724; RD -7.2% (-9.0% to -5.4%) † | I |
| Ip ⁸ | | S | HCQ: 21/97 vs. 305/970; RD -9.8% (-18.5% to -1.1%) † | |
| Komissarov ³³ | | C | HCQ: 7/33 vs. 0/10; RD 21.2% (7.3% to 35.2%) † | |
| Discharge from Hospital | | | | |
| Horby ¹³ | RCT | SC | HCQ: 941/1561 vs. 1982/3155; RD -2.5% (-5.5% to 0.4%) | L |
| Paccoud ¹⁷ | Cohort | S | HCQ: 21/38 vs. 26/46; RD -1.3% (-22.6% to 20.1%) | I |
| Mahévas ²⁵ | | S | HCQ: 67/84 vs. 71/89; RD 0% (-12% to 12%) | |
| Sbidian ²⁷ | | M | HCQ: 351/623 vs. 1507/3792; RD 16.6% (12.4% to 20.8%) | |
| Symptom Resolution | | | | |
| Skipper ¹² | RCT | H | HCQ: [At 14 days:] 152/201 vs. 135/194; RD 6.0% (-2.7% to 14.8%) | I |
| J Chen ¹⁵ | | SC | HCQ: [Fever] 1 vs. 1 day; MD 0 days (NA) | |
| L Chen ¹⁶ | | H | HCQ: [Days to recovery [Median(IQR)]]: 6 (3-8) vs. 7.5 (5-16.3) | |
| Z Chen ³² | | SC | HCQ: [Fever] 2.2±0.4 vs. 3.2±1.3 days; MD -1 day (-1.5 to -0.5) † [Cough] 2.0±0.2 vs. 3.1±1.5 days; MD -1.1 days (-1.6 to -0.6) † | |
| Tang ³⁴ | | H | HCQ: [Composite symptom resolution] 32/64 vs. 24/55; RD 6.4% (-11.6% to 24.3%) | |
| Abd-Elsalam ⁴ | | H | HCQ: [Complete recovery] 52/97 vs. 33/97; RD 19.5% (5.9% to 33.3%) | |
| Kalligeros ⁹ | Cohort | S | HCQ±AZ: [Days to improvement] MD +0.23 days (-1.87 to 2.33)** | I |

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| | | | HCQ±AZ: [Days to fever resolution] MD +1.27 days (-0.49 to 3.03)** | |
| Progression of Pulmonary Lesions on CT Scan | | | | |
| J Chen ¹⁵ | RCT | SC | HCQ: 5/15 vs. 7/15; RD -13.3% (-48.1% to 21.4%) | L |
| Z Chen ³² | | SC | HCQ: 2/31 vs. 9/31; RD -22.6% (-40.8% to -4.4%) † | |
| Improvement in Pulmonary Lesions on CT Scan | | | | |
| Z Chen ³² | RCT | SC | HCQ: 25/31 vs. 17/31; RD 25.8% (3.4% to 48.2%) † | I |
| Upper Respiratory Virological Clearance | | | | |
| C-P Chen ¹⁴ | RCT | SC | HCQ: [Day 14] 17/21 vs. 9/12; RD 6.0% (-23.8% to 35.7%) | I |
| J Chen ¹⁵ | | SC | HCQ: [Day 7] 13/15 vs. 14/15; RD -6.7% (-28% to 14.7%) [Day 14] 15/15 vs. 15/15; RD 0% (NA) | |
| L Chen ¹⁶ | | H | HCQ: [Day 10] 15/15 vs/ 12/12; RD 0% (NA) | |
| Tang ³⁴ | | H | HCQ: [Day 23] 53/75 vs. 56.75; RD -4% (-18.3% to 10.3%) | |
| C-P Chen ¹⁴ | Cohort | C | HCQ: [Day 14] 12/28 vs. 5/9; RD -12.7% (-50.0% to 24.6%) | I |
| Lecronier ¹⁸ | | C | HCQ: [Day 7] 7/26 vs. 2/14; RD 12.6% (-12.4% to 37.7%) | |
| Mallat ²¹ | | S | HCQ: [Day 14] 11/23 vs. 10/11; RD -43.1% (-69.6% to -16.5%) † | |
| Gautret ³⁵ | | C | HCQ±AZ: [Day 6] 14/20 vs. 2/16; RD 57.6% (31.8% to 83.3%) † | |

RoB: Risk of bias; HCQ: Hydroxychloroquine; SOE: Strength of evidence; 95%CI: 95% confidence interval; RCT: Randomized controlled trial; RD: Absolute risk difference; MD: Mean difference; ARDS: Acute respiratory distress syndrome. † denotes a statistically significant finding. Risk of Bias Codes: SC – some concerns, H– high, M – moderate, S –serious, C – critical, NI – no information; Strength of Evidence Codes: I – insufficient, L- low.

*adjusted hazard ratio, number of events per treatment group not provided.

**mean difference, mean values and standard deviation per treatment group not provided.