Supplemental Online Content

Walsh PS, Schnadower D, Zhang Y, Ramgopal S, Shah SS, Wilson PM. Association of early oseltamivir with improved outcomes in hospitalized children with influenza, 2007-2020. *JAMA Pediatr*. Published online September 19, 2022. doi:10.1001/jamapediatrics.2022.3261

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This supplemental material has been provided by the authors to give readers additional information about their work.

eTable 1: Covariate Distributions in Unweighted and Propensity Score Weighted Samples

| Covariates Distributions in Unweighted and PS Weighted Samples | | | | | |
|--|-------------------------|-------------------|-------------------------|-------------------|--|
| | Unweighted Samples | | PS Weighted Samples | | |
| Characteristics | No Early Oseltamivir | Early Oseltamivir | No Early Oseltamivir | Early Oseltamivir | |
| Age | | | | | |
| < 2 | 41.0% (9258) | 35.0% (11624) | 33.7% (11190) | 35.0% (11624) | |
| 2 - 5 | 21.0% (4755) | 20.5% (6806) | 20.8% (6903) | 20.5% (6806) | |
| ≥ 5 | 38.0% (8579) | 44.5% (14777) | 45.5% (15113) | 44.5% (14777) | |
| Race | | | | | |
| Black | 22.1% (5000) | 25.1% (8336) | 24.8% (8220) | 25.1% (8336) | |
| White | 54.2% (12245) | 49.6% (16461) | 49.9% (16577) | 49.6% (16461) | |
| Other | 18.4% (4150) | 20.4% (6782) | 20.3% (6756) | 20.4% (6782) | |
| Unknown | 5.3% (1197) | 4.9% (1628) | 5.0% (1654) | 4.9% (1628) | |
| Insurance | Insurance | | | | |
| Public | 61.4% (13872) | 64.6% (21466) | 64.5% (21403) | 64.6% (21466) | |
| Private | 32.2% (7279) | 30.8% (10225) | 31.1% (10341) | 30.8% (10225) | |
| Other | 4.4% (1003) | 3.1% (1041) | 3.1% (1036) | 3.1% (1041) | |
| Unknown | 1.9% (438) | 1.4% (475) | 1.3% (428) | 1.4% (475) | |
| ccc | | | | | |
| CCC, Yes | 46.8% (10581) | 48.1% (15979) | 48.7% (16171) | 48.1% (15979) | |
| Asthma | Asthma | | | | |
| Asthma, Yes | 9.4% (2118) | 12.6% (4170) | 12.2% (4050) | 12.6% (4170) | |
| Early ICU | | | | | |
| Early ICU, Yes | 24.0% (5429) | 28.9% (9610) | 28.1% (9347) | 28.9% (9610) | |
| Ventilation | | | | | |
| Ventilation, Yes | 13.4% (3032) | 17.6% (5846) | 17.0% (5633) | 17.6% (5846) | |
| Early Antibiotic | | | | | |
| Early Antibiotic, Yes | 65.4% (14764) | 67.8% (22527) | 68.2% (22640) | 67.8% (22527) | |

Abbreviations: CCC, complex chronic condition; ICU, intensive care unit.
^aOther includes those identified as other, Asian, American Indian/Alaskan/Native American, or NativeHawaiian/Pacific Islander.

eTable 2: Characteristics of children treated with early, late, and no oseltamivir during hospitalization.

| Characteristic | No Oseltamivir | Late Oseltamivir | Early Oseltamivir |
|----------------------------|----------------|------------------|-------------------|
| Number | 18,494 | 4,098 | 33,207 |
| Age Categories | | | |
| < 2 | 8027 (43.4) | 1231 (30.0) | 11624 (35.0) |
| 2 - 5 | 3853 (20.8) | 902 (22.0) | 6806 (20.5) |
| ≥ 5 | 6614 (35.8) | 1965 (48.0) | 14777 (44.5) |
| Gender | | | |
| Male | 10337 (55.9) | 2254 (55.0) | 18715 (56.4) |
| Female | 8153 (44.1) | 1842 (44.9) | 14491 (43.6) |
| Race | | | |
| Black | 4021 (21.7) | 979 (23.9) | 8336 (25.1) |
| White | 10214 (55.2) | 2031 (49.6) | 16461 (49.6) |
| Other | 3243 (17.5) | 907 (22.1) | 6782 (20.4) |
| Unknown | 1016 (5.5) | 181 (4.4) | 1628 (4.9) |
| Insurance Type | | | |
| Public | 11278 (61.0) | 2594 (63.3) | 21466 (64.6) |
| Private | 6093 (32.9) | 1186 (28.9) | 10225 (30.8) |
| Other | 846 (4.6) | 157 (3.8) | 1041 (3.1) |
| Unknown | 277 (1.5) | 161 (3.9) | 475 (1.4) |
| Region | | | |
| Midwest | 4352 (23.5) | 919 (22.4) | 8082 (24.3) |
| Northeast | 1992 (10.8) | 456 (11.1) | 3237 (9.7) |
| South | 7988 (43.2) | 1582 (38.6) | 12817 (38.6) |
| West | 4162 (22.5) | 1141 (27.8) | 9071 (27.3) |
| CCC | | | |
| No | 10704 (57.9) | 1307 (31.9) | 17228 (51.9) |
| Yes | 7790 (42.1) | 2791 (68.1) | 15979 (48.1) |
| History of Asthma (no CCC) | | | |
| No | 16643 (90.0) | 3831 (93.5) | 29037 (87.4) |
| Yes | 1851 (10.0) | 267 (6.5) | 4170 (12.6) |
| Early ICU | | | |
| No | 14263 (77.1) | 2900 (70.8) | 23597 (71.1) |
| Yes | 4231 (22.9) | 1198 (29.2) | 9610 (28.9) |

Abbreviations: CCC, complex chronic condition; ICU, intensive care unit.

^aOther includes those identified as other, Asian, American Indian/Alaskan/Native American, or NativeHawaiian/Pacific Islander.

eTable 3: Covariate Distributions in Weighted Cohort by Age Group

| Group/Covariates Age: < 2 Race Insurance CCC | Black White Other Unknown Public Private Other | 25.7% (2881) 45.4% (5086) 23.1% (2583) 5.7% (640) 72.5% (8112) 23.5% (2628) | 24.2% (2817) 46.6% (5419) 22.9% (2667) 6.2% (721) 72.9% (8472) |
|--|--|--|--|
| Race | White Other Unknown Public Private | 45.4% (5086) 23.1% (2583) 5.7% (640) 72.5% (8112) 23.5% (2628) | 46.6% (5419) 22.9% (2667) 6.2% (721) 72.9% (8472) |
| Insurance | White Other Unknown Public Private | 45.4% (5086) 23.1% (2583) 5.7% (640) 72.5% (8112) 23.5% (2628) | 46.6% (5419) 22.9% (2667) 6.2% (721) 72.9% (8472) |
| | Other Unknown Public Private | 23.1% (2583) 5.7% (640) 72.5% (8112) 23.5% (2628) | 22.9% (2667) 6.2% (721) 72.9% (8472) |
| | Unknown Public Private | 5.7% (640) 72.5% (8112) 23.5% (2628) | 6.2% (721) 72.9% (8472) |
| | Public Private | 72.5% (8112) 23.5% (2628) | 72.9% (8472) |
| | Private | 23.5% (2628) | ` ' |
| ccc | | , , | 23.0% (2675) |
| CCC | | 2.9% (322) | 2.7% (310) |
| CCC | Unknown | 1.2% (129) | 1.4% (167) |
| | CCC, Yes | 40.5% (4533) | 33.2% (3854) |
| Asthma | Asthma, Yes | 6.4% (712) | 5.1% (597) |
| Early ICU | Early ICU, Yes | 35.2% (3941) | 27.9% (3247) |
| Ventilation | Ventilation, Yes | 21.3% (2383) | 17.0% (1975) |
| Early Antibiotic | Early Antibiotic, Yes | 67.0% (7498) | 63.7% (7399) |
| Age: 2 - 5 | Larry / massesse, 100 | 01.070 (1.00) | (1000) |
| Race | Black | 24.9% (1717) | 25.0% (1701) |
| | White | 48.2% (3325) | 49.7% (3382) |
| | Other | 21.7% (1501) | 20.5% (1392) |
| | Unknown | 5.2% (360) | 4.9% (331) |
| Insurance | Public | 65.7% (4536) | 64.7% (4406) |
| inouranoo | Private | 29.6% (2047) | 30.1% (2047) |
| | Other | 3.3% (225) | 3.6% (243) |
| | Unknown | 1.4% (95) | 1.6% (110) |
| CCC | CCC, Yes | 48.9% (3377) | 50.6% (3442) |
| Asthma | Asthma, Yes | 14.9% (1030) | 14.8% (1006) |
| Early ICU | Early ICU, Yes | 24.2% (1668) | 27.8% (1891) |
| Ventilation | Ventilation, Yes | 14.5% (1003) | 16.7% (1136) |
| Early Antibiotic | Early Antibiotic, Yes | 70.0% (4833) | 68.6% (4666) |
| | , , | (111) | (111, |
| Race | Black | 24.0% (3622) | 25.8% (3818) |
| | White | 54.0% (8165) | 51.8% (7660) |
| | Other | 17.7% (2672) | 18.4% (2723) |
| | Unknown | 4.3% (654) | 3.9% (576) |
| nsurance | Public | 57.9% (8754) | 58.1% (8588) |
| | Private | 37.5% (5667) | 37.2% (5503) |
| | Other | 3.2% (489) | 3.3% (488) |
| | Unknown | 1.3% (204) | 1.3% (198) |
| CCC | CCC, Yes | 54.7% (8261) | 58.8% (8683) |
| Asthma | Asthma, Yes | 15.3% (2309) | 17.4% (2567) |
| Early ICU | Early ICU, Yes | 24.7% (3737) | 30.3% (4472) |
| Ventilation | Ventilation, Yes | 14.9% (2246) | 18.5% (2735) |
| Early Antibiotic | Early Antibiotic, Yes | 68.2% (10308) | 70.8% (10462) |

Abbreviations: CCC, complex chronic condition; ICU, intensive care unit.

aOther includes those identified as other, Asian, American Indian/Alaskan/Native American, or NativeHawaiian/Pacific Islander.

eTable 4: Covariate Distributions in Weighted Cohort by Presence of a Complex **Chronic Condition**

| | | No Early | |
|------------------|-----------------------|---------------|-------------------|
| Group/Covariates | Categories | Oseltamivir | Early Oseltamivir |
| CCC: No | · | <u>.</u> | |
| Age | < 2 | 39.1% (6657) | 45.1% (7770) |
| | 2 - 5 | 20.7% (3526) | 19.5% (3364) |
| | ≥ 5 | 40.2% (6853) | 35.4% (6094) |
| Race | Black | 24.7% (4203) | 23.9% (4118) |
| | White | 50.0% (8513) | 49.2% (8477) |
| | Other | 19.1% (3251) | 20.7% (3558) |
| | Unknown | 6.3% (1069) | 6.2% (1075) |
| Insurance | Public | 61.4% (10460) | 62.7% (10807) |
| | Private | 33.7% (5741) | 32.3% (5568) |
| | Other | 3.6% (620) | 3.6% (614) |
| | Unknown | 1.3% (214) | 1.4% (239) |
| Asthma | Asthma, Yes | 23.8% (4050) | 24.2% (4170) |
| Early ICU | Early ICU, Yes | 21.8% (3721) | 27.8% (4796) |
| Ventilation | Ventilation, Yes | 11.9% (2033) | 15.2% (2624) |
| Early Antibiotic | Early Antibiotic, Yes | 63.0% (10738) | 61.1% (10524) |
| CCC: Yes | | | |
| Age | < 2 | 28.0% (4533) | 24.1% (3854) |
| | 2 - 5 | 20.9% (3377) | 21.5% (3442) |
| | ≥ 5 | 51.1% (8261) | 54.3% (8683) |
| Race | Black | 24.8% (4018) | 26.4% (4218) |
| | White | 49.9% (8064) | 50.0% (7984) |
| | Other | 21.7% (3505) | 20.2% (3224) |
| | Unknown | 3.6% (585) | 3.5% (553) |
| Insurance | Public | 67.7% (10942) | 66.7% (10659) |
| | Private | 28.4% (4600) | 29.1% (4657) |
| | Other | 2.6% (416) | 2.7% (427) |
| | Unknown | 1.3% (213) | 1.5% (236) |
| Early ICU | Early ICU, Yes | 34.8% (5626) | 30.1% (4814) |
| Ventilation | Ventilation, Yes | 22.3% (3600) | 20.2% (3222) |
| Early Antibiotic | Early Antibiotic, Yes | 73.6% (11902) | 75.1% (12003) |

Abbreviations: CCC, complex chronic condition; ICU, intensive care unit.
^aOther includes those identified as other, Asian, American Indian/Alaskan/Native American, or NativeHawaiian/Pacific Islander.

eTable 5: Covariate Distributions in Weighted Cohort by Asthma History

| | | No Early | |
|--------------------|-----------------------|---------------|-------------------|
| Group/Covariates | Categories | Oseltamivir | Early Oseltamivir |
| History of Asthma: | No | | |
| Age | < 2 | 35.9% (10479) | 38.0% (11027) |
| | 2 - 5 | 20.1% (5873) | 20.0% (5800) |
| | ≥ 5 | 43.9% (12805) | 42.0% (12210) |
| Race | Black | 23.6% (6887) | 23.9% (6941) |
| | White | 50.6% (14751) | 50.5% (14651) |
| | Other | 21.0% (6119) | 20.9% (6063) |
| | Unknown | 4.8% (1399) | 4.8% (1382) |
| Insurance | Public | 64.7% (18855) | 65.4% (18980) |
| | Private | 30.9% (9016) | 30.1% (8728) |
| | Other | 3.1% (904) | 3.1% (898) |
| | Unknown | 1.3% (381) | 1.5% (431) |
| CCC | CCC, Yes | 55.5% (16171) | 55.0% (15979) |
| Early ICU | Early ICU, Yes | 28.7% (8374) | 27.9% (8112) |
| Ventilation | Ventilation, Yes | 17.4% (5060) | 17.1% (4976) |
| Early Antibiotic | Early Antibiotic, Yes | 69.0% (20132) | 69.5% (20187) |
| History of Asthma: | Yes | | |
| Age | < 2 | 17.6% (712) | 14.3% (597) |
| | 2 - 5 | 25.4% (1030) | 24.1% (1006) |
| | ≥ 5 | 57.0% (2309) | 61.6% (2567) |
| Race | Black | 32.9% (1333) | 33.5% (1395) |
| | White | 45.1% (1826) | 43.4% (1810) |
| | Other | 15.7% (637) | 17.2% (719) |
| | Unknown | 6.3% (255) | 5.9% (246) |
| Insurance | Public | 62.9% (2548) | 59.6% (2486) |
| | Private | 32.7% (1325) | 35.9% (1497) |
| | Other | 3.3% (132) | 3.4% (143) |
| | Unknown | 1.1% (46) | 1.1% (44) |
| Early ICU | Early ICU, Yes | 24.0% (972) | 35.9% (1498) |
| Ventilation | Ventilation, Yes | 14.1% (572) | 20.9% (870) |
| Early Antibiotic | Early Antibiotic, Yes | 61.9% (2508) | 56.1% (2340) |

Abbreviations: CCC, complex chronic condition; ICU, intensive care unit.

^aOther includes those identified as other, Asian, American Indian/Alaskan/Native American, or NativeHawaiian/Pacific Islander.

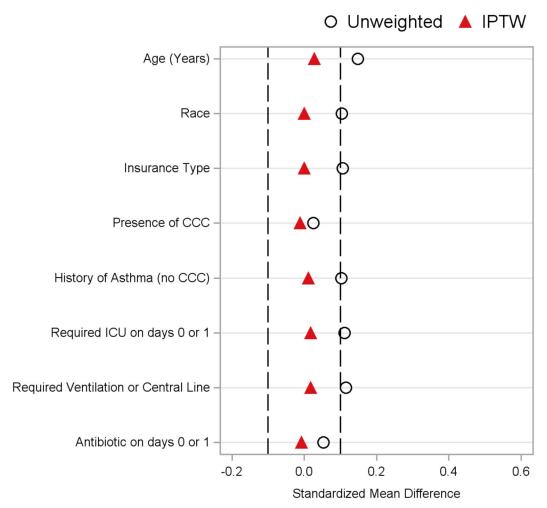
eTable 6: Covariate distributions in weighted cohort by early ICU requirement

| No Early | | | |
|------------------|-----------------------|---------------|-------------------|
| Group/Covariates | Categories | Oseltamivir | Early Oseltamivir |
| Early ICU: No | · | | |
| Age | < 2 | 30.4% (7250) | 35.5% (8377) |
| | 2 - 5 | 21.9% (5235) | 20.8% (4915) |
| | ≥ 5 | 47.7% (11376) | 43.7% (10305) |
| Race | Black | 24.5% (5834) | 25.9% (6103) |
| | White | 50.3% (12004) | 48.3% (11399) |
| | Other | 20.1% (4788) | 20.9% (4928) |
| | Unknown | 5.2% (1235) | 4.9% (1167) |
| Insurance | Public | 63.2% (15089) | 65.6% (15486) |
| | Private | 32.2% (7680) | 29.9% (7051) |
| | Other | 3.3% (780) | 3.0% (714) |
| | Unknown | 1.3% (312) | 1.5% (346) |
| CCC | CCC, Yes | 44.2% (10545) | 47.3% (11165) |
| Asthma | Asthma, Yes | 12.9% (3078) | 11.3% (2672) |
| Ventilation | Ventilation, Yes | 3.6% (853) | 3.8% (900) |
| Early Antibiotic | Early Antibiotic, Yes | 64.7% (15440) | 64.7% (15273) |
| Early ICU: Yes | · | · | |
| Age | < 2 | 42.2% (3941) | 33.8% (3247) |
| | 2 - 5 | 17.8% (1668) | 19.7% (1891) |
| | ≥ 5 | 40.0% (3737) | 46.5% (4472) |
| Race | White | 48.9% (4573) | 52.7% (5062) |
| | Black | 25.5% (2386) | 23.2% (2233) |
| | Other | 21.1% (1968) | 19.3% (1854) |
| | Unknown | 4.5% (419) | 4.8% (461) |
| Insurance | Public | 67.5% (6314) | 62.2% (5980) |
| | Private | 28.5% (2661) | 33.0% (3174) |
| | Other | 2.7% (256) | 3.4% (327) |
| | Unknown | 1.2% (116) | 1.3% (129) |
| CCC | CCC, Yes | 60.2% (5626) | 50.1% (4814) |
| Asthma | Asthma, Yes | 10.4% (972) | 15.6% (1498) |
| Ventilation | Ventilation, Yes | 51.1% (4780) | 51.5% (4946) |
| Early Antibiotic | Early Antibiotic, Yes | 77.0% (7199) | 75.5% (7254) |

Abbreviations: CCC, complex chronic condition; ICU, intensive care unit.
^aOther includes those identified as other, Asian, American Indian/Alaskan/Native American, or NativeHawaiian/Pacific Islander.

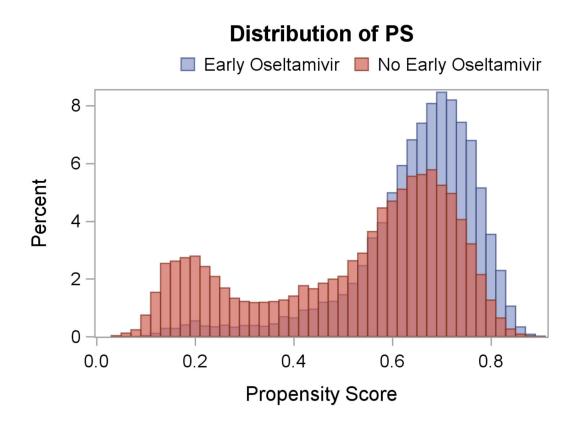
eFigure 1: Love plot of standardized mean differences of covariates for unweighted and IPTW cohort in the primary analysis.

Standardized Mean Difference

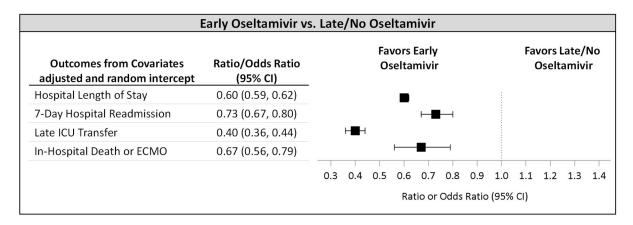


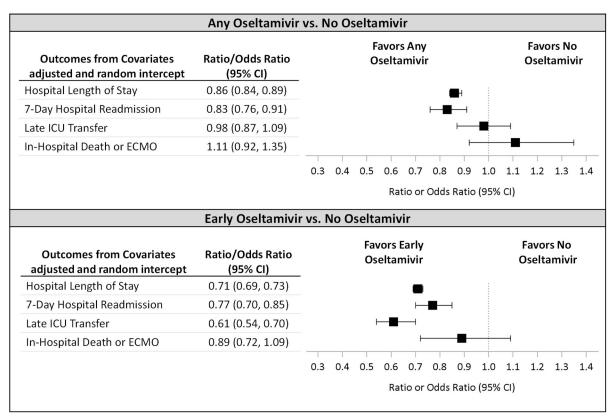
IPTW, inverse probability treatment weighting; CCC, complex chronic condition; ICU, intensive care unit.

eFigure 2: Distributions of propensity scores for treatment with early oseltamivir.



eFigure 3: Results of covariate-adjusted mixed effects models of oseltamivir use and outcomes.





Upper panel: Analysis of early oseltamivir compared to no early oseltamivir.

Lower panel: Analyses of any oseltamivir treatment vs. none and early oseltamivir vs. none (excluding late oseltamivir).