

**Data Supplement to Gerhard et al. Comparative mortality risks of antipsychotic medications in community-dwelling older adults. *Br J Psychiatry* doi: 10.1192/bjp.bp.112.122499**

**Table DS1 Dose conversion to chlorpromazine equivalencies**

| Antipsychotic Agent | Chlorpromazine (CPZ) equivalencies<br>(APM dose * FACTOR = 100 mg of CPZ)† | Chlorpromazine (CPZ) equivalencies<br>(APM dose * FACTOR = 100 mg of CPZ)‡ |
|---------------------|--|--|
| Haloperidol         | 50   | 54.3   |
| Aripiprazole        | 13.3   | 15.6   |
| Olanzapine          | 20   | 21.1   |
| Quetiapine          | 1.3  | 0.7  |
| Risperidone         | 50   | 75.8   |
| Ziprasidone         | 1.7  | 1.98   |

†Woods et al. (28)

‡Andreasen et al. (29)

**Table DS2. APM Dose Distributions Observed in the Study Population**

| Mg chlorpromazine equivalencies* (mg actual dose) |                   |               |            |               |             |               |             |               |        |               |        |               |         |               |
|---|-------------------|---------------|------------|---------------|-------------|---------------|-------------|---------------|--------|---------------|--------|---------------|---------|---------------|
| Percentile  | All APMs combined | Risperidone   | Olanzapine | Quetiapine    | Haloperidol | Aripiprazole  | Ziprasidone |               |        |               |        |               |         |               |
| 10  | 25 (n/a)          | 13 (0.25)     | 50 (2.5)   | 33 (25)       | 25 (0.5)    | 67 (5.0)      | 34 (20)     |               |        |               |        |               |         |               |
| 20  | 25 (n/a)          | 25 (0.5)      | 50 (2.5)   | 33 (25)       | 25 (0.5)    | 67 (5.0)      | 34 (20)     |               |        |               |        |               |         |               |
| 30  | 33 (n/a)          | 25 (0.5)      | 50 (2.5)   | 33 (25)       | 50 (1.0)    | 67 (5.0)      | 68 (40)     |               |        |               |        |               |         |               |
| 40  | 50 (n/a)          | 25 (0.5)      | 50 (2.5)   | 33 (25)       | 50 (1.0)    | 67 (5.0)      | 68 (40)     |               |        |               |        |               |         |               |
| 50  | 50 (n/a)          | 25 (0.5)      | 100 (5.0)  | 65 (50)       | 50 (1.0)    | 133 (10.0)    | 68 (40)     |               |        |               |        |               |         |               |
| 60  | 50 (n/a)          | 50 (1.0)      | 100 (5.0)  | 65 (50)       | 100 (2.0)   | 133 (10.0)    | 68 (40)     |               |        |               |        |               |         |               |
| 70  | 98 (n/a)          | 50 (1.0)      | 100 (5.0)  | 65 (50)       | 100 (2.0)   | 133 (10.0)    | 102 (60)    |               |        |               |        |               |         |               |
| 80  | 100 (n/a)         | 50 (1.0)      | 100 (5.0)  | 98 (75)       | 150 (3.0)   | 200 (15.0)    | 136 (80)    |               |        |               |        |               |         |               |
| 90  | 150 (n/a)         | 100 (2.0)     | 200 (10.0) | 130 (100)     | 250 (5.0)   | 200 (15.0)    | 204 (120)   |               |        |               |        |               |         |               |
| Dose group**                                      | Mg CPZ            | % of patients | Mg RIS     | % of patients | Mg OLA      | % of patients | Mg QUE      | % of patients | Mg HAL | % of patients | Mg ARI | % of patients | Mg ZIP  | % of patients |
| Low   | ]0,50[            | 33.1%         | ≤0.5       | 55.7%         | ≤2.5        | 48.3%         | ≤25         | 48.6%         | ≤1     | 52.2%         | ≤5     | 42.4%         | ≤20     | 24.9%         |
| Medium  | [50,75]           | 36.0%         | ]0.5,1]    | 29.5%         | ]2.5,5]     | 33.3%         | ]25,50]     | 27.8%         | ]1,4]  | 33.9%         | ]5,15[ | 33.0%         | ]20,60[ | 41.7%         |
| High  | >75               | 30.9%         | >1         | 14.8%         | >5          | 18.4%         | >50         | 23.5%         | >4     | 13.9%         | ≥15    | 24.6%         | ≥60     | 33.4%         |

\* using the conversion from Woods et al. (28)

\*\*for dose-response analyses

**Table DS3. Consistency of Prescribed Daily APM Dose Over Follow-up**

|                | Last dose same as<br>index dose (%) | Second dose same<br>as index dose (%)* |
|----------------|-------------------------------------|--|
| All study APMs | <b>82.98</b>                        | <b>88.16</b>                           |
| Risperidone    | 81.98                               | 87.52                                  |
| Olanzapine     | 83.92                               | 88.82                                  |
| Quetiapine     | 80.90                               | 87.21                                  |
| Haloperidol    | 87.22                               | 90.10                                  |
| Aripiprazole   | 84.56                               | 88.97                                  |
| Ziprasidone    | 87.14                               | 89.50                                  |

\* For those without second APM claim except the index claim before the censoring date, index dose is used as the second dose in the calculation.

**Table DS4. Death within 180 days after start of new antipsychotic medication\***

|                                 | Risperidone<br>n=49,333<br>pyrs=11,262 |                     | Olanzapine<br>n=44,296<br>pyrs=10,674 |                     | Quetiapine<br>n=26,130<br>pyrs=6,136 |                   | Haloperidol<br>n=13,159<br>pyrs=2,261 |                     | Aripiprazole<br>n=1,908<br>pyrs=429 |                  | Ziprasidone<br>n=1,567<br>pyrs=328 |                   |
|---------------------------------|--|---------------------|---------------------------------------|---------------------|--------------------------------------|-------------------|---------------------------------------|---------------------|-------------------------------------|------------------|------------------------------------|-------------------|
|                                 | N                                      | Rate<br>(95%CI)     | N                                     | Rate<br>(95%CI)     | N                                    | Rate<br>(95%CI)   | N                                     | Rate<br>(95%CI)     | N                                   | Rate<br>(95%CI)  | N                                  | Rate<br>(95%CI)   |
| <b>Non-cancer mortality</b>     | 1,654                                  | 14.7<br>(14.0-15.4) | 1,204                                 | 11.3<br>(10.6-11.9) | 594                                  | 9.7<br>(8.9-10.5) | 710                                   | 31.4<br>(29.1-33.7) | 25                                  | 5.8<br>(3.5-8.1) | 29                                 | 8.8<br>(5.6-12.1) |
| <b>Cause-specific mortality</b> |  |                     |                                       |                     |                                      |                   |                                       |                     |                                     |                  |                                    |                   |
| Circulatory system              | 897                                    | 8.0<br>(7.4-8.5)    | 685                                   | 6.4<br>(5.9-6.9)    | 330                                  | 5.4<br>(4.8-6.0)  | 399                                   | 17.7<br>(15.9-19.4) | 13                                  | 3.0<br>(1.4-4.7) | 17                                 | 5.2<br>(2.7-7.6)  |
| Cerebrovascular diseases        | 155                                    | 1.4<br>(1.2-1.6)    | 99                                    | 0.9<br>(0.7-1.1)    | 53                                   | 0.9<br>(0.6-1.1)  | 68                                    | 3.0<br>(2.3-3.7)    | n/a                                 | 0.2<br>(0.0-0.7) | n/a                                | 0.6<br>(0.0-1.5)  |
| Respiratory system              | 230                                    | 2.0<br>(1.8-2.3)    | 178                                   | 1.7<br>(1.4-1.9)    | 84                                   | 1.4<br>(1.1-1.7)  | 101                                   | 4.5<br>(3.6-5.3)    | n/a                                 | 1.4<br>(0.3-2.5) | n/a                                | 0.9<br>(0.0-2.0)  |
| Other                           | 527                                    | 4.7<br>(4.3-5.1)    | 341                                   | 3.2<br>(2.9-3.5)    | 180                                  | 2.9<br>(2.5-3.4)  | 210                                   | 9.3<br>(8.0-10.5)   | n/a                                 | 1.4<br>(0.3-2.5) | n/a                                | 2.7<br>(1.0-4.5)  |

\*: Rate expressed per 100 person-years (pyrs)

n/a: cell size under 10.

**Table DS5. Hazard ratios of death during different follow-up periods**

|                             | HR (95% CI)                 |                             |                              |                               |  |
|-----------------------------|-----------------------------|-----------------------------|------------------------------|-------------------------------|--|
|                             | Full Duration<br>(180 days) | Duration 1<br>(0 - 39 days) | Duration 2<br>(40 - 79 days) | Duration 3<br>(80 - 180 days) |  |
| <b>NON-CANCER MORTALITY</b> |                             |                             |                              |                               |  |
| Risperidone                 | 1.0 -                       | 1.0 -                       | 1.0 -                        | 1.0 -                         |  |
| Olanzapine                  | 0.99 (0.91-1.07)            | 0.98 (0.91-1.06)            | 1.02 (0.91-1.15)             | 0.96 (0.81-1.15)              |  |
| Quetiapine                  | 0.81 (0.73-0.89)            | 0.89 (0.80-0.98)            | 0.84 (0.72-0.97)             | 0.71 (0.56-0.90)              |  |
| Haloperidol                 | 1.45 (1.31-1.60)            | 1.42 (1.29-1.56)            | 1.23 (1.04-1.45)             | 1.04 (0.76-1.42)              |  |
| Aripiprazole                | 0.66 (0.42-1.02)            | 0.71 (0.46-1.10)            | 0.81 (0.43-1.52)             | 1.03 (0.48-2.23)              |  |
| Ziprasidone                 | 0.87 (0.59-1.30)            | 0.91 (0.61-1.34)            | 1.06 (0.59-1.89)             | 1.03 (0.46-2.34)              |  |

Note: Hazard ratios are hdPS-adjusted.

**Table DS6. Stratified analyses of death within 180 Days**

|                    | Haloperidol |             | Aripiprazole |             | Olanzapine |             | Quetiapine |             | Ziprasidone |             |
|--------------------|-------------|-------------|--------------|-------------|------------|-------------|------------|-------------|-------------|-------------|
|                    | HR          | 95%CI       | HR           | 95%CI       | HR         | 95%CI       | HR         | 95%CI       | HR          | 95%CI       |
| <b>DEMENTIA</b>    |             |             |              |             |            |             |            |             |             |             |
|                    | 1.28        | (1.09-1.49) | 0.56         | (0.26-1.19) | 0.94       | (0.83-1.06) | 0.84       | (0.72-0.97) | 0.89        | (0.50-1.58) |
| <b>NO DEMENTIA</b> |             |             |              |             |            |             |            |             |             |             |
|                    | 1.55        | (1.37-1.75) | 0.66         | (0.37-1.18) | 1.01       | (0.91-1.11) | 0.80       | (0.70-0.92) | 0.64        | (0.34-1.21) |
| <b>AGE 65-80</b>   |             |             |              |             |            |             |            |             |             |             |
|                    | 1.57        | (1.32-1.86) | 0.64         | (0.36-1.16) | 0.95       | (0.83-1.09) | 0.78       | (0.66-0.93) | 0.49        | (0.23-1.03) |
| <b>AGE ≥81</b>     |             |             |              |             |            |             |            |             |             |             |
|                    | 1.29        | (1.15-1.45) | 0.72         | (0.37-1.39) | 1.01       | (0.91-1.11) | 0.80       | (0.70-0.91) | 1.14        | (0.68-1.90) |

Note: Reference category for all comparisons is risperidone. All hazard ratios are adjusted using stratum specific high-dimensional propensity scores.