

Supplementary Table e-2. Demographic and clinical characteristic considering genders (A) and Riluzole treatment (B) in the London and Oxford cohorts separately and combined.

| A. Gender | London | | Oxford | | Combined | |
|---|----------------------|------------------------|----------------------|----------------------|----------------------|-----------------------|
| | Male (n=67) | Female (n=36) | Male (n=45) | Female (n=19) | Male (n=112) | Female (n=55) |
| Blood NfL level [^] (pg/mL) | 82.3 (46.5, 150) | 142.9 (76.1, 218.8)** | 90 (53, 146) | 90 (59, 183) | 84.9 (50.8, 146.3) | 125.7 (62, 214.7)* |
| Age At Sampling [^] (years) | 63.6 (57.8, 68.3) | 69.7 (59.3, 74.2)* | 63 (53.5, 69) | 57 (54, 67) | 63.3 (55.2, 69) | 67 (55.3, 73) |
| Age At Onset [^] (years) | 61.9 (44, 66.1) | 67.7 (57.8, 73.8)** | 60.8 (51.1, 67.7) | 54.8 (51.2, 65.7) | 61.7 (51.1, 66.5) | 65.7 (53.2, 70.9)# |
| PRB [^] (points/month) | 0.449 (0.156, 0.853) | 0.684 (0.362, 1.471)** | 0.470 (0.232, 0.900) | 0.449 (0.275, 0.753) | 0.452 (0.181, 0.855) | 0.593 (0.313, 1.175)* |
| PRL [^] (points/month) | 0.521 (0.202, 0.893) | 0.770 (0.407, 1.326)* | 0.523 (0.226, 0.963) | 0.449 (0.272, 1.067) | 0.522 (0.210, 0.925) | 0.705 (0.354, 1.175)* |
| ALSFRS_R Score [^] | 39 (33, 44) | 33.5 (26.3, 37.8)*** | 35 (31, 39) | 36 (32, 37) | 38 (32, 42) | 35 (28, 37)*** |
| Diagnosis [n (%)] | | | | | | |
| Definite ALS | 13 (19.4%) | 6 (16.7%) | 13 (28.9%) | 7 (36.8%) | 26 (23.2%) | 13 (23.6%) |
| Probable ALS | 26 (38.8%) | 21 (58.3%) | 14 (31.1%) | 7 (36.8%) | 40 (35.7%) | 28 (50.9%) |
| Possible ALS | 21 (31.3%) | 6 (16.7%) | 18 (40%) | 5 (26.4%) | 39 (34.8%) | 11 (20%) |
| Lab-Supported probable ALS | 7 (10.5%) | 3 (8.3%) | 0 | 0 | 7 (6.3%) | 3 (5.5%) |
| | | | | | | |
| B. Use of Riluzole | London | | Oxford | | Combined | |

| | No (n=29; M/F: 21/8) | Yes (n=74; M/F: 46/28) | No (n=34; M/F: 24/10) | Yes (n=30; M/F: 21/9) | No (n=63; M/F: 45/18) | Yes (n=104; M/FL 67/37) |
|-----------------------------|----------------------|------------------------|-----------------------|-----------------------|-----------------------|-------------------------|
| Blood NfL level^ (pg/mL) | 66.8 (37.5, 139.2) | 114 (58.7, 178.8)## | 98.5 (50.3, 155.3) | 88.5 (57.5, 135.5) | 83 (42, 151) | 99.9 (59, 158.6) |
| Age At Sampling^ (years) | 64.4 (58.2, 69.9) | 66.1 (58.7, 72.5) | 66.5 (60.3, 70.8) | 56 (51.5, 65.3)** | 65.5 (58.6, 70) | 63.8 (54, 69.8) |
| Age At Onset^ (years) | 62.2 (54, 67.6) | 64.7 (56.5, 70.1) | 63.6 (56.3, 68.9) | 53.6 (50.0, 62.3)** | 62.9 (54.4, 68.6) | 61.8 (51.2, 67.9) |
| PRB^ (points/month) | 0.516 (0.127, 0.784) | 0.513 (0.246, 1.017) | 0.402 (.0220, 0.838) | 0.516 (0.321, 0.801) | 0.410 (0.169, 0.785) | 0.516 (0.267, 0.914) |
| PRL^ (points/month) | 0.521 (0.18, 0.913) | 0.661 (0.331, 1.088) | 0.387 (0.206, 0.839) | 0.623 (0.343, 1.009) | 0.449 (0.206, 0.888) | 0.643 (0.338, 1.067)* |
| ALSFRS_R Score^ | 39 (31.5, 43) | 37 (27.8, 41.3) | 36 (32, 39.3) | 35 (31, 38) | 37 (32, 41) | 36 (30.1, 40) |
| Diagnosis [n (%)] | | | | | | |
| Definite ALS | 3 (10.3%) | 16 (21.6%) | 9 (26.5%) | 11 (36.7%) | 12 (19%) | 27 (26.0%) |
| Probable ALS | 10 (34.5%) | 37 (50%) | 9 (26.5%) | 12 (40%) | 19 (30.2%) | 49 (47.0%) |
| Possible ALS | 12 (41.4%) | 15 (20.3%) | 16 (47%) | 7 (23.3%) | 28 (44.5%) | 22 (21.2%) |
| Lab-Supported probable ALS | 4 (13.8%) | 6 (8.1%) | 0 | 0 | 4 (6.3%) | 6 (5.8%) |

^: Median (IQR)

Comparison was made between the gender and within each cohort; Mann-Whitney U test or unpaired *t*-test was used where suitable.

*: $p < 0.05$; **: $p < 0.01$; ***: $p < 0.001$; #: $p = 0.054$; ##: $p = 0.057$