

THE OFFICIAL JOURNAL OF THE AMERICAN SOCIETY OF CLINICAL PSYCHOPHARMACOLOGY

**Supplementary Material** 

- Article Title: Treatment Response With Esketamine Nasal Spray Plus an Oral Antidepressant in Patients With Treatment-Resistant Depression Without Evidence of Early Response: A Pooled Post Hoc Analysis of the TRANSFORM Studies
- Author(s): Ibrahim Turkoz, PhD; Ella Daly, MD; Jaskaran Singh, MD; Xiwu Lin, PhD; Yevgen Tymofyeyev, PhD; David Williamson, PhD; Giacomo Salvadore, MD; Abigail I. Nash, MD, PhD; Matthew Macaluso, DO; Samuel T. Wilkinson, MD; and J. Craig Nelson, MD
- DOI Number: https://doi.org/10.4088/JCP.20m13800

## List of Supplementary Material for the article

- 1. <u>Table 1</u> Safety Summary by Treatment Group During the Double-Blind Induction Phase
- 2. <u>Table 2</u> Study Completion/Withdrawal Information for Day 2 and Days 2 and 8 Nonresponders During the Double-Blind Induction Phase

## **Disclaimer**

This Supplementary Material has been provided by the author(s) as an enhancement to the published article. It has been approved by peer review; however, it has undergone neither editing nor formatting by in-house editorial staff. The material is presented in the manner supplied by the author.

© Copyright 2021 Physicians Postgraduate Press, Inc.

## Supplementary Table 1. Safety Summary by Treatment Group During the Double-

## **Blind Induction Phase**

|                                 | ESK + AD   | PBO + AD   |
|---------------------------------|------------|------------|
|                                 | n = 343    | n = 222    |
| TEAEs in ≥5% of subjects, n (%) |            |            |
| Overall                         | 299 (87.2) | 143 (64.4) |
| Anxiety                         | 30 (8.8)   | 12 (5.4)   |
| Blood pressure increased        | 30 (8.8)   | 5 (2.3)    |
| Diarrhea                        | 23 (6.7)   | 13 (5.9)   |
| Dissociation                    | 92 (26.8)  | 8 (3.6)    |
| Dizziness                       | 81 (23.6)  | 15 (6.8)   |
| Dizziness postural              | 22 (6.4)   | 1 (0.45)   |
| Dry mouth                       | 19 (5.5)   | 7 (3.2)    |
| Dysgeusia                       | 65 (19.0)  | 30 (13.5)  |
| Fatigue                         | 24 (7.0)   | 11 (5.0)   |
| Feeling drunk                   | 18 (5.3)   | 1 (0.5)    |
| Headache                        | 70 (20.4)  | 38 (17.1)  |
| Hypoesthesia                    | 37 (10.8)  | 3 (1.4)    |
| Hypoesthesia oral               | 37 (10.8)  | 3 (1.4)    |
| Insomnia                        | 29 (8.5)   | 16 (7.2)   |
| Nausea                          | 97 (28.3)  | 19 (8.6)   |
| Paresthesia                     | 43 (12.5)  | 4 (1.8)    |

| Paresthesia oral      | 19 (5.5)  | 3 (1.4)  |  |
|-----------------------|-----------|----------|--|
| Sedation              | 19 (5.5)  | 2 (0.90) |  |
| Somnolence            | 60 (17.5) | 20 (9.0) |  |
| Throat irritation     | 23 (6.7)  | 9 (4.1)  |  |
| Vertigo               | 77 (22.5) | 5 (2.3)  |  |
| Vision blurred        | 31 (9.0)  | 3 (1.4)  |  |
| Vomiting              | 32 (9.3)  | 4 (1.8)  |  |
| Serious TEAEs, n (%)  |           |          |  |
| Overall               | 3 (0.9)   | 1 (0.5)  |  |
| Depression            | 1 (0.3)   | 0        |  |
| Headache              | 1 (0.3)   | 0        |  |
| Multiple injuries     | 1 (0.3)   | 0        |  |
| Road traffic accident | 1 (0.3)   | 0        |  |
| Vertigo positional    | 0         | 1 (0.5)  |  |

AD, antidepressant; AE, adverse event; ESK, esketamine nasal spray; PBO, placebo

nasal spray, TEAE, treatment-emergent adverse event.

|                    | Overall    |            | Day 2 Nonresponders |            | Days 2 and 8 Nonresponders |            |
|--------------------|------------|------------|---------------------|------------|----------------------------|------------|
|                    | ESK+AD     | AD+PBO     | ESK+AD              | AD+PBO     | ESK+AD                     | AD+PBO     |
|                    | n = 343    | n = 222    | n = 263             | n = 184    | n = 235                    | n = 170    |
| Completed, n (%)   | 306 (89.2) | 206 (92.8) | 233 (88.6)          | 172 (93.5) | 213 (90.6)                 | 163 (95.9) |
| Withdrawn, n (%)   | 37 (10.8)  | 16 (7.2)   | 30 (11.4)           | 12 (6.5)   | 22 (9.4)                   | 7 (4.1)    |
| Adverse event      | 16 (4.7)   | 3 (1.4)    | 14 (5.3)            | 3 (1.6)    | 8 (3.4)                    | 2 (1.2)    |
| Lack of efficacy   | 4 (1.2)    | 0          | 4 (1.5)             | 0          | 3 (1.3)                    | 0          |
| Lost to follow-up  | 2 (0.6)    | 1 (0.5)    | 1 (0.4)             | 1 (0.5)    | 1 (0.4)                    | 1 (0.6)    |
| Withdrawal by      | 8 (2.3)    | 7 (3.2)    | 5 (1.9)             | 6 (3.3)    | 4 (1.7)                    | 3 (1.8)    |
| subject            |            |            |                     |            |                            |            |
| Protocol violation | 2 (0.6)    | 2 (0.9)    | 2 (0.8)             | 0          | 2 (0.9)                    | 0          |
| Other              | 5 (1.5)    | 3 (1.4)    | 4 (1.5)             | 2 (1.1)    | 4 (1.7)                    | 1 (0.6)    |

Supplementary Table 2. Study Completion/Withdrawal Information for Day 2 and Days 2

AD, antidepressant; ESK, esketamine nasal spray; PBO, placebo nasal spray.