Most Common Treatments in Patients with Treatment Resistant Depression Based on European Cohort Study Real-World Evidence

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OBJECTIVES

• To report the most commonly received treatments at baseline in the Treatment Resistant Depression Cohort in Europe (EOTC).

BACKGROUND

- Treatment resistant depression (TRD) is generally defined as failure to respond to at least two different treatments with antidepressants, received at an adequate dose for adequate duration, in the same major depressive episode.¹
- TRD affects 10–30% of patients with major depressive disorder,^{2–4} and carries with it a significant unmet need.^{5, 6}
- The EOTC study aimed to examine treatment patterns and outcomes among TRD patients in Europe, providing real-world data on the burden of disease.

METHODS

- A prospective, multicentre, observational cohort study of adults with TRD in Europe was conducted (NCT03373253; **Figure 1**).⁷
- At baseline, patients had a Montgomery-Åsberg
 Depression Rating Scale score ≥20, had failed at least
 two different oral antidepressant treatments and were
 initiating a new treatment for depression.
- Medical records, clinician-rated scales and patient-reported questionnaires were used to collect patient characteristics and medical history.
- For patients receiving combination or augmentation therapies (**Figure 2**), all ongoing pharmacological treatments were recorded at baseline.

RESULTS

- In total, 411 patients were included, reporting 54 different pharmacological treatments used at baseline either as monotherapy, or as part of combination or augmentation therapy.
- At baseline, the mean duration of the current major depressive episode was 2.6 years (136.3 weeks), and the mean number of previous episodes was 3.4; as per the inclusion criteria, all patients had moderate or severe depression at baseline (**Table 1**).
- Nine different medications were received by ≥10% of patients: venlafaxine (24.8%), mirtazapine (21.2%), quetiapine (14.1%), bupropion (14.1%), duloxetine (14.1%), vortioxetine (13.1%), sertraline (10.9%), escitalopram (10.9%) and trazodone (10.2%; **Figure 3**).
- Of these, all were oral antidepressants except quetiapine, an antipsychotic.
- Of the eight oral antidepressants received by ≥10% of patients, two were selective serotonin reuptake inhibitors (sertraline and escitalopram), two were serotonin norepinephrine reuptake inhibitors (venlafaxine and duloxetine), and four were recorded as 'other' (trazodone, bupropion, mirtazapine and vortioxetine).

CONCLUSIONS



In this study, the pharmacological treatments used at baseline were heterogeneous, reflecting the heterogeneity of TRD and its treatment.



The most commonly received treatments were predominantly oral antidepressants.



However, the third most received treatment was quetiapine, mainly used in clinical practice as an augmentation therapy, suggesting it is a routine treatment for patients with TRD.

NEUROPSYCHIATRY

SUMMARY of the 9 drugs used in SSRIs and SNRIs 1 was the antipsychotic, quetiapine, mainly used in clinical practice as an augmentation therapy FIGURE 1. EOTC study design Baseline data Observational period

Baseline data collection^a

Event-driven^b MADRS and CGI-S measurement

New antidepressant treatment (dose/frequency and treatment in line with local clinical practice)

Baseline

Month 6

Month 12

Scheduled data collection

Scheduled data collection

Scheduled data collection

^aBaseline data were documented ±14 days of baseline date (on which new treatment was started). ^bAny clinically relevant worsening/improvement in the current major depressive episode. CGI-S: Clinical Global Impression-Severity; EOTC: European Observational TRD Cohort; MADRS: Montgomery-Åsberg Depression Rating Scale; TRD: treatment resistant depression.

FIGURE 2. TRD treatment strategies

Monotherapy



Involves one therapy of a single class approved for the treatment of MDD such as an oral antidepressant³

Combination Therapy



Involves the prescription of at least two antidepressants from multiple classes, such as an SSRI plus an alpha-2 receptor antagonist³

Augmentation



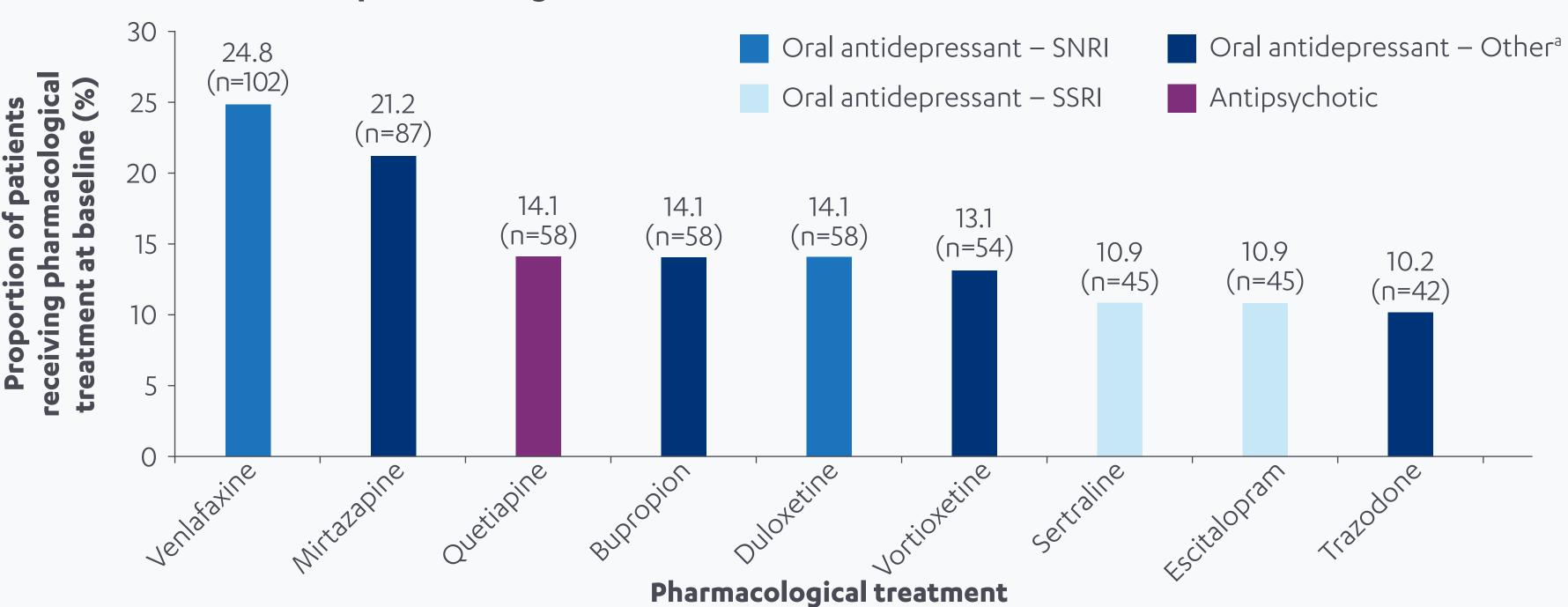
Involves the prescription of an antidepressant plus another agent that, while not primarily licensed as an antidepressant, may enhance the clinical response of the antidepressant therapy, such as an antidepressant plus an antipsychotic³

MDD: major depressive disorder; SSRI: selective serotonin reuptake inhibitor; TRD: treatment resistant depression.

TABLE 1. Baseline characteristics

Mean (SD), unless otherwise stated	All patients (N=411)
Socio-demographics	
Age, years	51.0 (10.8)
Female patients, % (n)	62.3 (256)
Psychiatric and medical history	
Age at diagnosis of MDD, years	37.2 (13.1)
Years since MDD diagnosis	13.7 (11.2)
Number of previous episodes	3.4 (5.6)
Duration of current MDE, weeks	
Mean	136.3 (203.8)
Median (min, max)	69.6 (10-2,242)
Clinical characteristics	
MADRS total score	31.8 (6.0)
Depression severity: MADRS score	e category
Severe, % (n)	32.6 (134)
Moderate, % (n)	67.4 (277)

FIGURE 3. Most common pharmacological treatments



Some drugs were received in combination; therefore, the total is greater than 100%. Figure includes all pharmacological treatments received by ≥10% of patients at baseline. a'Other' includes or all antidepressants not classed as an SNRI or SSRI, such as tetracyclic antidepressants or serotonin modulators. SNRI: serotonin norepinephrine reuptake inhibitor; SSRI: selective serotonin reuptake inhibitor.

ABBREVIATIONS: **CGI-S**: Clinical Global Impression-Severity; **EOTC**: European Observational TRD Cohort; **MADRS**: Montgomery-Åsberg Depression Rating Scale; **MDD**: major depressive disorder; **MDE**: major depressive episode; **SD**: standard deviation; **SNRI**: serotonin norepinephrine reuptake inhibitor; **SSRI**: selective serotonin reuptake inhibitor; **TRD**: treatment resistant depression; **UK**: United Kingdom.

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²Jaffe DH et al. BMC Psychiatry 2019;19:247; ³Al-Harbi KS et al. Patient Prefer Adherence 2012;6:369–88; ⁴Voineskos D et al. Neuropsychiatr Dis Treat 2020;16:221–34; ⁵Popova V et al. Am J Psychiatry 2019;176:428–38; ⁶Daly EJ et al. JAMA Psychiatry 2019;76:893–903; ⁷Heerlein K et al. J Affect Disord 2021;283:115–22. DISCLOSURES: KH: Employee of Janssen Global Services. YG, YK, SMH, TI, CVH: Employees of Janssen EMEA. ACKNOWLEGEMENTS: Substantial contributions to study conception/design, or acquisition/analysis/interpretation of data: KH, YG, KY, SMH, TI, CVH; Drafting of the publication, or revising it critically for important intellectual content: KH, YG, KY, SMH, TI, CVH; Final approval of the publication: KH, YG, KY, SMH, TI, CVH. We would like to thank the patients and their caregivers in addition to all the investigators and their teams who contributed to this study. The authors acknowledge Emma Francis Gregory, BA, Costello Medical, Cambridge, UK and Carolyn Walsh, PhD, Costello Medical, London, UK for medical writing and editorial assistance, and the Costello Medical Design Team for design support. This study was funded by Janssen EMEA. All costs associated with development of this presentation were funded by Janssen EMEA.