Unraveling the Treatment Preferences of Patients and Healthcare Providers in Treatment-Resistant Depression: A Targeted Literature Review

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Introduction

- Pharmacologic and somatic therapies used for treatment-resistant depression (TRD) differ in modality and associated monitoring requirements, potential benefits, and tolerability
- Some treatments are used off-label, without the rigorous reassurance concerning product safety and efficacy that is provided by regulatory approval
- As a result, individuals with TRD and their healthcare providers (HCPs) have to make difficult trade-offs when selecting a treatment

Objectives

- The targeted literature review (TLR) aimed to identify the treatmentrelated concepts that matter most to patients and their HCPs when they select a treatment for TRD. The specific objectives were to:
- Describe and summarize treatment-related concepts that have been included in previously published quantitative preference studies and themes arising in qualitative literature exploring treatment preferences
- Provide an overview of relevant concepts that might affect preference for TRD treatments

TABLE 1: Inclusion and exclusion criteria

Category	Inclusion criteria	Exclusion criteria
Population	Adult patients diagnosed with MDD or TRD or physicians treating MDD or TRD	 Child/adolescent studies General population studies Patients not diagnosed with depression Physicians not treating depression Nonhuman studies
Treatment	Disease burden, QoL, treatment satisfaction, treatment preference, treatment burden	Not applicable
Study design	 Qualitative studies Interviews Focus groups Nominal group technique Quantitative studies Quantitative stated preference studies (e.g., DCE, BWS, SW, MCDA, thresholding, BRA, conjoint analysis) Observational preference studies (e.g., revealed preferences) 	 Publications of studies with the following designs: Animal studies In vitro/ex vivo studies Gene expression/protein expression studies Case studies/case series Publications that are not of empirical studies or studies without primary data collection (e.g., reviews, editorials) Clinical trials
Language	English-language articles	Not applicable

BRA, benefit-risk analysis; BWS, best-worst scaling; DCE, discrete choice experiment; MCDA, multicriteria decision analysis; MDD, major depressive disorder; QoL, quality of life; SW, swing weighting; TRD, treatment-resistant depression.

Methods

- A TLR¹ was undertaken to identify qualitative and quantitative studies describing patient or HCP treatment preferences in TRD published since 2013, via the Ovid database (Embase, PubMed)
- Double-screening of titles/abstracts and full-text screening were conducted in May/June 2024. Data extraction for the studies that met all inclusion and no exclusion criteria (Table 1) was completed in June 2024
- Relevant clinical study reports and published clinical studies on the efficacy and safety of investigational and/or marketed treatments were included to identify relevant clinical trial endpoints in TRD

FIGURE 1: PRISMA diagram of literature search



PRISMA, Preferred Reporting Items for Systematic Reviews and Meta-Analyses. *Double screening of titles and abstracts.

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Results

Characteristics of included preference studies

- Twelve studies (n = 9 qualitative²⁻¹⁰; n = 3 quantitative¹¹⁻¹³) were included in the review (**Figure 1**)
- The majority of studies related to preferences for pharmaceuticals (n = 10) and half were undertaken in the US (n = 6). Qualitative studies most commonly used interview methods (n = 6), whereas all quantitative studies used discrete choice methods (n = 3) (**Figure 2**)





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Identified concepts that might affect preference for TRD treatments

• Fifty-two treatment-related concepts were identified as potentially relevant for patients or HCPs when selecting a treatment, including concepts associated with potential benefits, risks, and other treatment characteristics, such as those related to administration. Common concepts are included in **Figure 3**



DCE, discrete choice experiment; ECT, electroconvulsive therapy; HCP, healthcare provider; MDD, major depressive disorder; TRD, treatment-resistant depression. The numbers in the chart refer to the number of studies included. *Some studies cover multiple categories.

FIGURE 4: Evidence review summary and preference research gaps identified



- Treatment efficacies were often important in influencing treatment selection and were described by a range of different characteristics
- Key treatment benefits considered important by patients and HCPs included the chance of symptom response or remission, the chance of maintaining a response, and, to a lesser extent, the time to respond

Risks

Relevant risks were treatment-dependent and included potential metabolic effects (e.g., weight gain), cognitive or psychiatric effects (e.g., sleepiness, memory loss, dissociation), gastrointestinal side effects, and sexual dysfunction

HCP, healthcare provider; TRD, treatment-resistant depression



include the mode and frequency of administration, the need for monitoring, treatment duration, and the out-of-pocket cost of treatment

Research Gaps

- associated risks
- Previous studies have not explored the perspectives and preferences of patients or HCPs regarding aspects related to the availability of long-term safety data and FDA approval, which differentiate some TRD treatments and may be important considerations when choosing between different treatments

MDD



Γ	Frequency
	Treatment Location
	Monitoring
	Administration
	Treatment Duration
	Monotherapy or Combination Therapies

Other treatment characteristics that may be relevant for patients

There is a scarcity of preference literature that evaluates patient or HCP preferences related to novel treatments, their benefits, and

Limitations



This study is based on a targeted literature review conducted in English. While the approach follows systematic review principles, it may not capture all relevant studies, especially those in other languages, potentially introducing selection bias

Conclusions



Individuals with TRD and their HCPs place significant value on symptom improvement/response, remission, longlasting efficacy, and minimization of side effects.



Treatment modality, frequency, and setting are also relevant considerations that may influence patient and HCP treatment preferences



Understanding the treatment preferences of patients with TRD and their HCPs can facilitate shared decision-making and enable personalized treatment plans



The review will inform the selection of candidate attributes that are suitable for inclusion in a quantitative preference elicitation instrument

Acknowledgements

Hui Lu, Rosanne Janssens, and Jennifer A. Whitty drafted the poster. All authors (Josh Hamilton, Yuxian Du, Hui Lu, Kruti Joshi, Rosanne Janssens, Jennifer A. Whitty, and Chinwe Adebiyi) provided input into the poster and critically reviewed and approved the final version.

Novel Pathways in Depression (③)





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This study was funded by Janssen Scientific Affairs, LLC, a Johnson & Johnson company and conducted by Evidera.