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Introduction

- Patients with treatment-resistant depression (TRD) who are candidates for interventional treatments (IV ketamine, intranasal esketamine, transcranial magnetic stimulation) are often not identified or referred late for treatments from which they may benefit.
- This mixed-method study examined how a small group of US prescribers identify TRD patients and what delays referral or initiation of interventional treatments, including IV ketamine, esketamine, and transcranial magnetic stimulation (TMS).

Objective/Hypothesis

- To characterize current TRD patient identification practices, referral patterns, and barriers to interv-entional care among US prescribers.

Methods

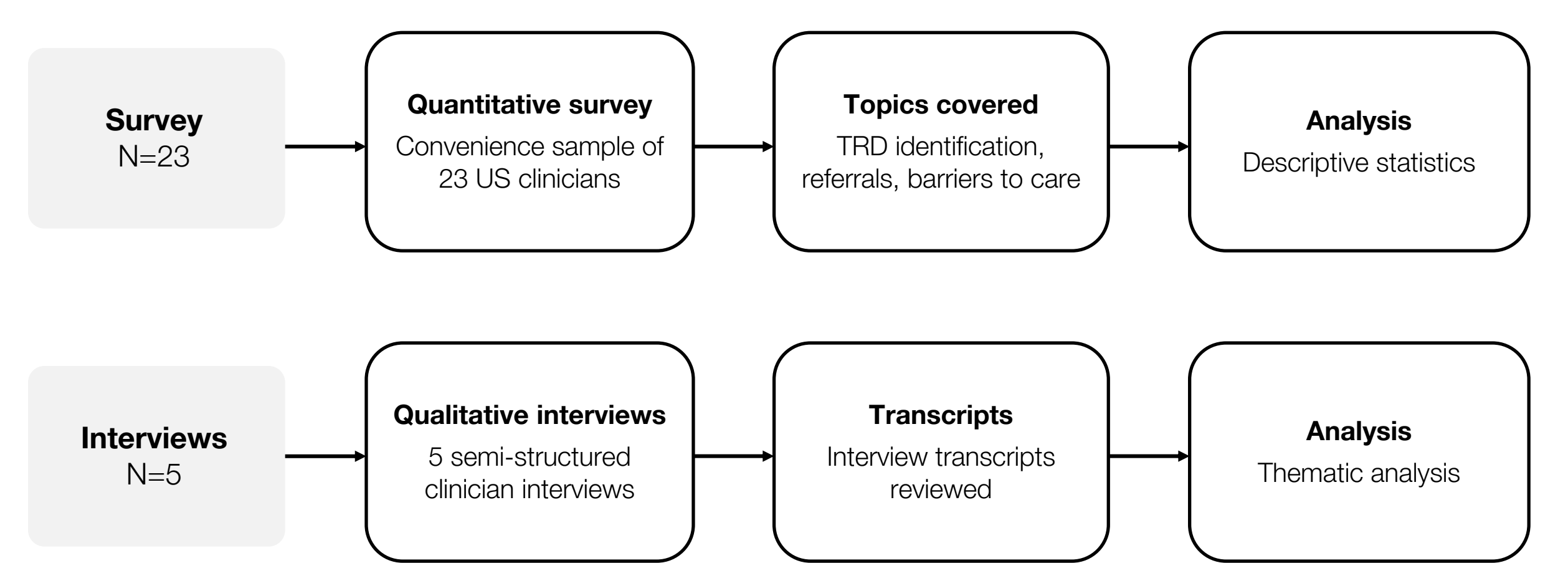
Study design

- Using a mixed-methods survey, a convenience sample of 23 clinicians was surveyed to understand identification of TRD patients, referrals, and barriers to interventional care.
- 5 qualitative clinician interviews were conducted with a semi-structured interview guide.

Analysis

- Responses were summarized using descriptive statistics and semi-structured interview transcripts were reviewed and thematically analyzed.

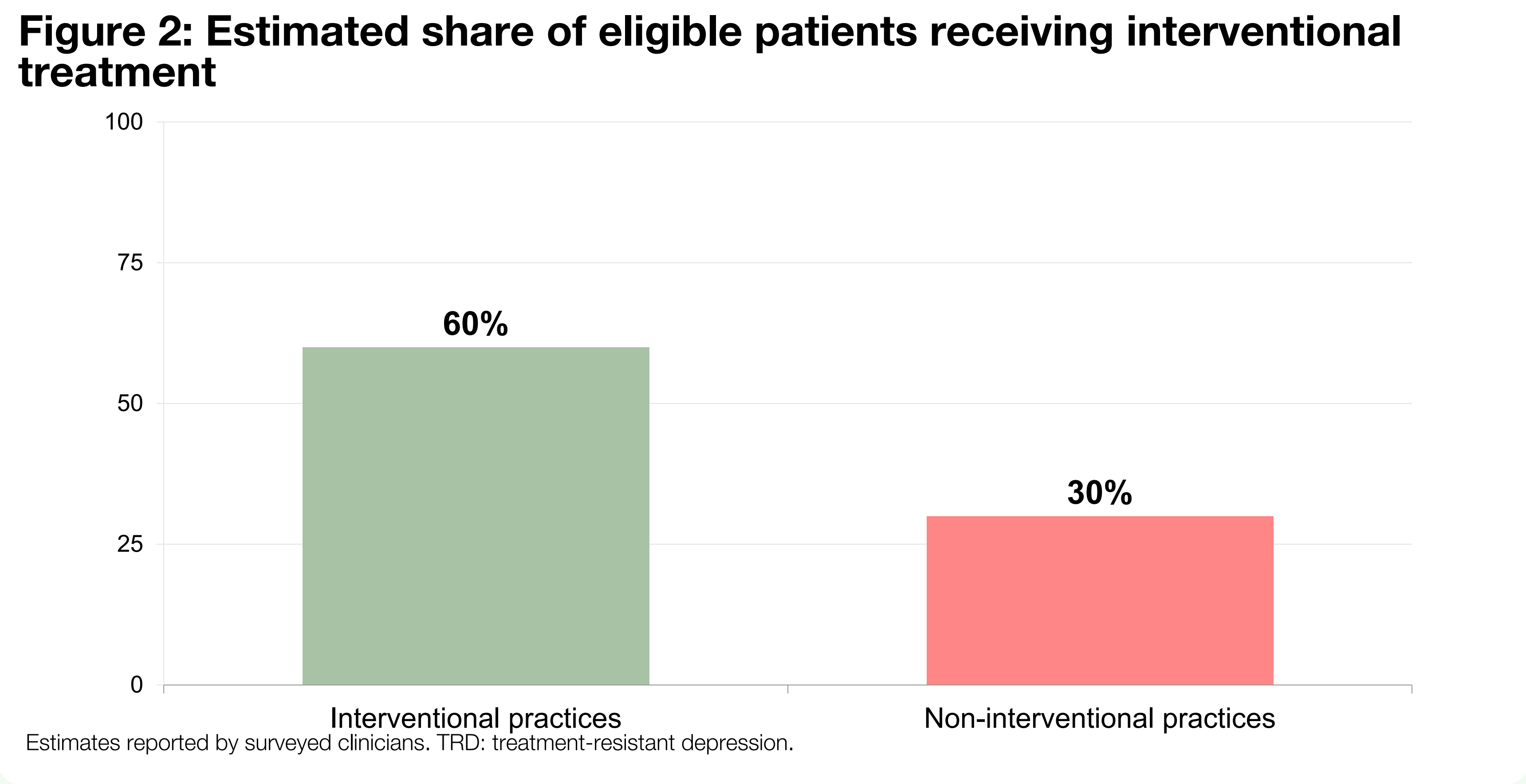
Figure 1: Mixed-methods study design



Abbreviations: TRD: treatment-resistant depression; TMS: transcranial magnetic stimulation; IV: intravenous; PHQ-9: Patient Health Questionnaire-9.

Results

- Most surveyed clinicians determined the TRD status of patients on an ad-hoc basis.
- Failure of a defined number of oral antidepressants was the primary criterion for initiating interventional treatment.
- The PHQ-9 was the most commonly used scale to assess disease severity and treatment response.



Barriers to interventional care

- Patient resistance (**100%**), and costs and insurance coverage (**93%**) were cited as barriers to interventional therapies.

Timing of interventional treatment

- Although more than **90%** of interventionalists were willing to begin interventional care after three or fewer failed oral antidepressants, nearly **60%** of patients arrived having failed four or more.

Table 1: Key survey findings

Finding	Estimate
Eligible patients receiving interventional treatment — interventional practices	60%
Eligible patients receiving interventional treatment — non-interventional practices	30%
Clinicians citing patient resistance as a barrier	100%
Clinicians citing costs and insurance coverage as a barrier	93%
Interventionalists willing to begin interventional care after ≤3 failed oral antidepressants	>90%
Patients arriving having already failed ≥4 oral antidepressants	~60%

Conclusions

- In a small convenience sample of clinicians, findings suggest that TRD identification in real-world practice is largely informal and inconsistent, and eligible patients are frequently referred for interventional care later than preferred.
- Patient resistance and insurance/cost barriers were the dominant obstacles.
- These findings characterize where delays and variability arise in the TRD treatment pathway.

COI Disclosure

Research was conducted by Osmind in collaboration with Compass Pathways. D.A., J.L., J.Q., and W.S. All authors are employees of and/or hold equity in Osmind. V.W., Y.H., L.O., and M.S. are employees of a subsidiary of Compass Pathways PLC and hold shares and/or share options in Compass Pathways PLC.

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TRD is identified **ad hoc**, and eligible patients reach interventional care **later** than clinicians would like. **Patient resistance** and **insurance costs** are the leading barriers.