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## Understanding the Health System Conditions Affecting the Use of Long-Acting Injectable Antipsychotics in the Treatment of Schizophrenia in Clinical Practice: A US Healthcare Provider Survey

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Purpose: To describe factors that enable the routine use of long-acting injectable antipsychotics (LAIs) for appropriate patients in the current clinical practice, including changes in LAI prescribing due to the COVID-19 pandemic and expectations for prescribing in 2021 in the United States (US).

Methods: Frequent LAI prescribers recruited from a nationwide panel in 2020 completed an online survey regarding practice characteristics, perspectives on healthcare system conditions enabling routine use of LAIs, and prescribing patterns and changes in patterns during the COVID-19 pandemic.

**Results:** Of 408 prescribers who completed the survey, 77.7% were physicians and 59.1% had  $\geq$ 10 years of psychiatry practice. More than half of frequent prescribers (57.1%) reported treating >20% of their patients with schizophrenia with LAIs. The American Psychiatric Association (APA) guideline was followed by 64.0% of prescribers. Most prescribers identified poor adherence to antipsychotics as a circumstance when LAIs are recommended (94,9%) and patient/caregiver involvement in treatment decisions as a key factor impacting the decision to prescribe LAIs (97.3%). Most prescribers reported that LAI prescribing rates were unchanged in 2020 (59.8%). Similar proportions of prescribers expected no change (44.1%) or an increase (42.9%) in LAI prescribing rates in 2021. The number of patients followed, cost of treatment, and availability of staff to administer LAIs were the main driving factors identified by prescribers expecting an increase in LAI prescribing rates.

**Conclusion:** LAIs were commonly recommended to patients with poor adherence, and patient/caregiver involvement was an important factor affecting prescribers' treatment decisions. LAI prescribing rates remained unchanged during the COVID-19 pandemic in 2020. Keywords: COVID-19, healthcare providers, long-acting injectable antipsychotics, prescribing patterns, schizophrenia

### Introduction

Schizophrenia is a debilitating, chronic psychiatric disorder characterized by symptoms of delusions and hallucinations.<sup>1,2</sup> The annual prevalence of schizophrenia is estimated to range between 0.25% and 0.64% in the United States (US).<sup>3-5</sup>

Oral antipsychotics (OAPs) are commonly used in routine care for schizophrenia.<sup>2,6</sup> However, adherence to medication that requires daily oral administration may be difficult to maintain. Thus, long-acting injectable antipsychotics (LAIs), which are administered biweekly, monthly, or once every few months, have the potential to improve adherence and symptom control and reduce the rate of hospitalizations in patients with schizophrenia compared to OAPs.<sup>7-9</sup> Indeed, the 2021

American Psychiatric Association (APA) schizophrenia practice guideline recommends the use of LAIs in patients with a history of nonadherence and by patient choice,<sup>2</sup> while the Florida Medicaid guidelines recommend LAIs as the initial treatment after a trial of an OAP monotherapy or among patients who are nonadherent or refractory to initial treatment.<sup>10</sup>

Despite guideline recommendations and evidence demonstrating improved antipsychotic adherence and symptom control with LAIs vs OAPs,<sup>2,10–13</sup> LAIs are infrequently prescribed. Reports indicate that fewer than 20% of patients with schizophrenia are prescribed LAIs in Western countries.<sup>14,15</sup> Multiple factors may influence the decision to treat with LAIs vs OAPs, including patient characteristics, preferences, and costs.<sup>15,16</sup> For instance, LAI prescription was found to be associated with female sex and younger age in two separate studies of real-world treatment patterns.<sup>16,17</sup> Physician and patient perspectives on LAIs may also create barriers to their use.<sup>15,18</sup> In a survey-based study of physicians discussed the option of LAIs with only 50% of their patients, notably because they were concerned about potentially damaging the relationship established with their patients.<sup>18</sup> Furthermore, patient attitudes toward the option of using LAIs were largely neutral or unfavorable. Additional factors that may influence the utilization of LAIs vs OAPs in clinical practice include costs related to storage, time and effort related to their administration, and the availability of trained healthcare professionals to administer doses.<sup>15</sup> Given the high risk of relapse and hospitalization when patients with schizophrenia are nonadherent or inadequately treated,<sup>19</sup> removing these barriers to the use of LAIs may reduce the clinical and public health burden associated with the condition.

More recently, the COVID-19 pandemic has challenged care provision for patients with schizophrenia and put certain in-person encounters needed to ensure patients' stability, such as the administration of LAIs, at risk.<sup>20,21</sup> Disruption of continuity of care in schizophrenia and subsequent decompensation may place, in the context of a global pandemic, an additional strain on healthcare systems worldwide, further increasing emergency department visits and hospitalizations and contributing to COVID-19 outbreaks.<sup>21</sup> However, the effect of COVID-19 on LAI use may as well have been minimal based on the relatively infrequent administration of LAIs.

Given the evolving guidelines related to LAI use, the role of LAIs in reducing hospitalizations, and the challenges associated with provision of in-person care during the COVID-19 pandemic, there is a need to better evaluate characteristics of healthcare providers and clinical practices enabling the routine use of LAIs for appropriate patients, as well as changes in LAI prescribing due to the COVID-19 pandemic and expectations for prescribing in 2021.

### Methods

### Data Source

For this descriptive, hypothesis-generating study, de-identified, individual-level data were collected nationwide between November 17, 2020, and December 12, 2020, through an anonymous cross-sectional double-blinded online survey of US-based frequent LAI prescribers (ie, those who treat  $\geq$ 1 patient with LAIs monthly) via an online panel, which comprised US-based healthcare professionals with a psychiatry specialty. Study participants received a link to the online questionnaire containing 41 questions and a description of the study and were required to provide their informed consent prior to responding to the survey questions.

The survey collected information about the characteristics of participating prescribers, the prescribers' perspectives on healthcare system conditions enabling routine use of LAIs, and the prescribers' perspectives on the evolution of LAI prescribing patterns and the conditions that influenced them both during the COVID-19 pandemic and in 2021.

All study materials, including the study protocol and the questionnaire, were approved by the New England Independent Review Board (now WIRB-Copernicus Group Independent Review Board) and the study was granted an institutional review board exemption (#17-1364883-1).

### **Study Participants**

Study participants recruited from a nationwide online panel were screened for the following criteria: (1) had a primary specialty in psychiatry, (2) were prescribed LAIs for the treatment of schizophrenia in the past 12 months, and (3) had  $\geq 1$  patient with schizophrenia treated with LAIs in their caseload in an average month (ie, were considered a frequent LAI

prescriber). The threshold for identification as a frequent LAI prescriber was determined based on the 50th percentile or higher in the distribution of annual patient days on LAIs per prescriber obtained across all US LAI prescribers in 2019 (data on file).

### Statistical Analysis

The responses from the survey were summarized using means, medians, and standard deviations (SDs) for continuous variables and frequencies and proportions for categorical variables. Completed surveys did not contain missing data because answers to all questions were required. All analyses were conducted using SAS Enterprise Guide 7.1 (SAS Institute, Cary, NC).

### Subgroup Analysis

Of all the frequent prescribers included in the study, two subgroups were identified. Specifically, the frequent prescribers were stratified based on the proportion of patients with schizophrenia in a monthly caseload who were treated with LAIs: (1) moderately frequent LAI prescribers (ie, those who treated  $\leq 20\%$  of patients with schizophrenia with LAIs) and (2) very frequent LAI prescribers (ie, those who treated  $\geq 20\%$  of patients with schizophrenia with LAIs). Prescribers who answered "don't know or unsure" when queried for the percentage of their patients with schizophrenia who were treated with LAIs were excluded from either subgroup.

## Results

### Characteristics of Frequent LAI Prescribers and Their Clinical Practices

Between November 17, 2020, and December 12, 2020, a total of 408 prescribers (of 648 who viewed the survey) completed the online survey (Table 1). Most of the frequent LAI prescribers were physicians (77.7%), 17.2% were nurse practitioners, and 5.1% were physician assistants. Most frequent LAI prescribers had >10 years of practice in psychiatry (59.1%). Almost all practiced in urbanized areas or urban clusters (92.7%), with the most common clinical settings being community health centers (27.9%), private practices (22.8%), and inpatient units of psychiatric hospitals (15.2%).

The availability of LAI administration services on site was reported by 71.6% of frequent LAI prescribers; other commonly reported on-site services included counseling (82.1%) and social work/referral services (71.8%). The average time spent with a patient during an encounter (excluding the first encounters) was in the range of 11–30 minutes for 87.8% of frequent LAI prescribers. The prescribers surveyed identified medical staff involved in the care of patients with schizophrenia as including psychiatrists (81.6%), registered nurses (64.0%), social workers (61.8%), psychologists/ therapists (55.4%), and nurse practitioners (49.3%).

### Psychiatric Caseload of Frequent LAI Prescribers

More than two-thirds of frequent LAI prescribers had total monthly psychiatric caseloads of  $\leq 200$  patients (71.6%) (Figure 1A) and a subset of patients with schizophrenia in their psychiatric caseload  $\leq 40\%$  (71.5%) (Figure 1B). More than half (57.1%) of frequent LAI prescribers reported treating over 20% of their patients with schizophrenia with LAIs (Figure 1C). In absolute terms, just over half (53.2%) of frequent LAI prescribers treated >10 patients with schizophrenia with LAIs in a given month (Figure 2).

### Education on LAI Use

When reporting receiving routine training and/or education on LAI use, 75.7% of prescribers (including physicians, nurse practitioners, and physician assistants) and 60.3% of registered nurses reported receiving frequent training and/or education, while the number of pharmacists, patients or caregivers, social workers, and psychologists or therapists who reported receiving routine LAI training and/or education was lower (Table 2). The prescribers noted their main resources for education on LAI use as online continuing medical education (CME; 79.9%), professional literature (78.9%), and participation in conferences or congresses (58.6%) (Table 2). Frequent LAI prescribers reported using

 Table I Characteristics of Frequent LAI Prescribers

	Frequent LAI Prescribers, n (%) N = 408
Job title	
Physician	317 (77.7)
Nurse practitioner	70 (17.2)
Physician assistant	21 (5.1)
Years of practice in psychiatry	
<5 years	67 (16.4)
5–10 years	100 (24.5)
II-I5 years	68 (16.7)
>15 years	173 (42.4)
Geographic region of clinical practice	
South	148 (36.3)
Northeast	105 (25.7)
West	86 (21.1)
Midwest	69 (16.9)
Type of geographic area of clinical practice	
Urbanized area (≥50,000 people)	266 (65.2)
Urban cluster (2500–49,999 people)	112 (27.5)
Rural (<2500 people)	29 (7.1)
None of the above	I (0.2)
Clinical setting in which healthcare providers	(
prescribed the most LAIs	
Community health center	114 (27.9)
Private practice	93 (22.8)
Inpatient unit of psychiatric hospital	62 (15.2)
Outpatient clinic of hospital	61 (15.0)
Inpatient psychiatry unit of general hospital	34 (8.3)
Academic practice	12 (2.9)
PACT	11 (2.7)
Long-term care	5 (1.2)
Academic practice	12 (2.9)
None of the above	15 (3.7)
Do not know/unsure	I (0.2)
Services provided in clinical practice	. ()
Counseling (individual, family, group)	335 (82.1)
Social work/referral services	293 (71.8)
LAI administration services	292 (71.6)
Pharmacy	186 (45.6)
Peer specialists	139 (34.1)
Crisis center	133 (32.6)
Partial hospitalization program	94 (23.0)
Day programs	93 (22.8)
None of the above	12 (2.9)
Do not know/unsure	2 (0.5)
Average time spent with a patient during an encounter	
≤10 minutes	9 (2.2)
I I–20 minutes	163 (40.0)
21–30 minutes	195 (47.8)
>30 minutes	39 (9.6)
Do not know/unsure	2 (0.5)
bo not know/unsure	

Abbreviations: LAI, long-acting injectable; OAP, oral antipsychotics; PACT, Program of Assertive Community Treatment.

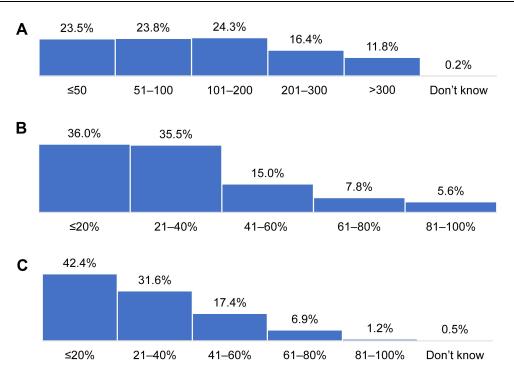


Figure I Monthly caseload of frequent LAI prescribers per respondents (N = 408). (A) Number of patients in monthly caseload; (B) share of patients with schizophrenia in monthly caseload; (C) share of patients treated with LAIs among those with schizophrenia. Abbreviation: LAI, long-acting injectable.

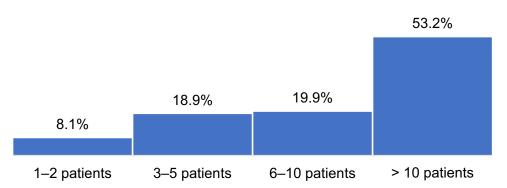


Figure 2 Number of patients with schizophrenia on LAIs in monthly caseload of frequent LAI prescribers per respondents (N = 408). Abbreviation: LAI, long-acting injectable.

2.9 resources (SD = 1.3; median = 3.0) on average for ongoing education on LAI use. Of note, 7.4% of prescribers reported that no LAI education was provided at their clinical practice.

Specialists identified by prescribers as providing education on LAI use to patients and caregivers were psychiatrists (81.9%), registered nurses (61.3%), and nurse practitioners (45.3%) (Table 2). The main resources and approaches to patient and caregiver education were in-person conversations (78.2%), paper information sheets (59.8%), and website referrals (eg, to advocacy groups; 43.9%) (Table 2).

# Schizophrenia Treatment Guidelines Followed by Frequent LAI Prescribers and Their Clinical Practices

Most frequent LAI prescribers reported following the APA practice guideline for the treatment of patients with schizophrenia (64.0%); 53.9% reported that their clinical practice followed this guideline (Figure 3). More than

	Frequent LAI Prescribers, n (%) N = 408
Individuals routinely trained or educated on LAI use	
Prescribers (physicians, nurse practitioners, physician assistants)	309 (75.7)
Registered nurses	246 (60.3)
Pharmacists	122 (29.9)
Patients or caregivers	77 (18.9)
No education provided at clinical practice	30 (7.4)
Social workers	25 (6.1)
Psychologists/therapists	24 (5.9)
Do not know/unsure	16 (3.9)
Professionals educating patients on LAI use	
Psychiatrist	334 (81.9)
Registered nurse	250 (61.3)
Nurse practitioner	185 (45.3)
Pharmacist	133 (32.6)
Social worker	90 (22.1)
Psychologist/therapist	69 (16.9)
Physician assistant	67 (16.4)
Other physician (not specialized in psychiatry)	20 (4.9)
Other	8 (2.0)
No education provided to patients and caregivers	2 (0.5)
Do not know/unsure	I (0.2)
Resources available for LAI education	
For prescribers	
Online CME	326 (79.9)
Reading professional literature	322 (78.9)
Participating in conferences/congresses	239 (58.6)
Professional association	166 (40.7)
Direct communication from pharmacies	81 (19.9)
Conducting research	37 (9.1)
None of the above	3 (0.7)
Do not know/unsure	3 (0.7)
For patients/caregivers	
In-person conversations	319 (78.2)
Paper information sheets	244 (59.8)
Website referrals (eg, to advocacy groups and other websites)	179 (43.9)
Phone calls	106 (26.0)
Videos	73 (17.9)
Emails	60 (14.7)
Apps	28 (6.9)
Resources from pharmaceutical companies	5 (1.2)
No resources available at clinical practice	12 (2.9)
Do not know/unsure	15 (3.7)

#### Table 2 Training and Education on LAI Use, According to Prescribers

Abbreviations: CME, continuing medical education; LAI, long-acting injectable.

half (52.9%) of prescribers reported that both they and their clinical practice follow the APA schizophrenia practice guideline. About one-quarter of responding prescribers (24.5%) and clinical practices (24.0%) did not follow any formal clinical guidelines when treating patients with schizophrenia (Figure 3).

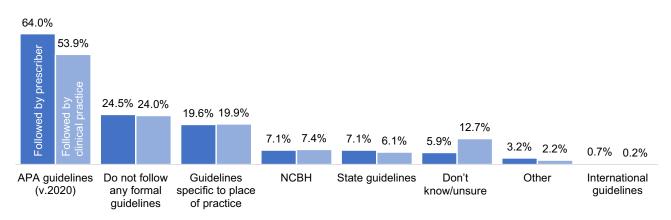


Figure 3 Schizophrenia treatment guidelines followed by respondents (N = 408).

Abbreviations: APA, American Psychiatric Association; NCBH, National Council for Behavioral Health.

## Circumstances in Which Frequent LAI Prescribers Recommended LAIs and Factors Impacting Prescribing of LAI in Eligible Patients

Most frequent LAI prescribers reported poor adherence to antipsychotics as a circumstance when LAIs are recommended in patients with schizophrenia (94.9%) (Table 3). Of note, 87.0% of frequent LAI prescribers reported that adherence to antipsychotics at their clinical practice was measured based on self-reported information from patients and/or caregivers. Other frequently reported circumstances for LAI prescribing included multiple relapses (86.8%), failure of treatment with oral medications (78.2%), and chronic schizophrenia (72.8%). Over one-third of frequent LAI prescribers (36.0%) reported that they recommend use of LAIs in the early stage of schizophrenia (Table 3).

Among patients clinically eligible for LAIs, the key factors impacting the decision to prescribe LAIs were patient and caregiver involvement in treatment decisions (97.3%), prescriber experience with managing adverse events (AEs) related to LAI use (95.8%), and cost of treatment for patients (90.2%) (Figure 4). Of note, 89.2% of the responding prescribers reported inviting patients and caregivers to participate in decisions regarding the best treatment plan.

# Collaboration of Frequent LAI Prescribers with Nonprescribing Members of Clinical Teams

More than three-quarters (80.6%) of LAI prescribers reported collaborating with nonprescribing members of the treatment team on schizophrenia treatment plans, including LAI selection (Table 3). Responding prescribers most frequently collaborated with registered nurses (51.7%), social workers (51.2%), psychologists/therapists (46.6%), and pharmacists (34.3%).

### Factors Affecting LAI Prescribing During the COVID-19 Pandemic

Most frequent LAI prescribers reported that LAI prescribing rates remained unchanged during the COVID-19 pandemic (59.8%), while 17.6% reported a decrease and 13.7% reported an increase in prescribing rates (Figure 5). Among those who reported that prescribing remained unchanged (n = 244), most identified patient concerns with COVID-19 exposure (56.6%) as a factor impacting the dynamics of LAI prescribing; patient concerns with transportation (45.5%) and staff concerns with COVID-19 exposure (42.2%) were also common responses (Figure 6A). Among those reporting that prescribing decreased (n = 72), most named patient and staff concerns with COVID-19 exposure (55.6% and 45.8%, respectively) as key factors impacting the dynamics of prescribing; facility closure or restricted visits (43.1%) and patient concerns with transportation (43.1%) were also common responses (Figure 6B). Among those reporting that prescribing increased (n = 56), common factors contributing to LAI prescribing dynamics were increased confidence in adherence with LAIs (44.6%) and patient concerns with transportation (44.6%) (Figure 6C).

	Frequent LAI Prescribers, n (%) N = 408
Circumstances where frequent prescribers recommended	
prescribing LAIs	
Poor adherence to antipsychotic medication	387 (94.9)
Multiple relapses	354 (86.8)
Failure of oral medications	319 (78.2)
Chronic schizophrenia	297 (72.8)
At patient's request	283 (69.4)
Early stage of schizophrenia	147 (36.0)
None of the above	2 (0.5)
Do not know/unsure	5 (1.2)
Antipsychotic adherence determination methods	
Self-reported information from patients and/or caregivers	355 (87.0)
Information from other healthcare professionals on the team	279 (68.4)
Refill information from pharmacy	255 (62.5)
Other <sup>a</sup>	27 (6.6)
Do not know/unsure	9 (2.2)
Nonprescribers collaborating with prescribers on	
schizophrenia treatment plans	
Registered nurses	211 (51.7)
Social workers	209 (51.2)
Psychologists/therapists	190 (46.6)
Pharmacists	140 (34.3)
Other <sup>b</sup>	13 (3.2)
None	73 (17.9)
Do not know/unsure	6 (1.5)

 Table 3 LAI Prescribing Patterns and Clinical Practice Characteristics of Frequent LAI

 Prescribers

**Notes**: <sup>au</sup>Other" responses included MAR (n = 14), clinical evaluation (n = 7), blood level measurement (n = 4), and PSYCKES/EMR (n = 2). <sup>bu</sup>Other" responses included case manager (n = 5), occupational therapist or recreational therapist (n = 4), licensed practical nurse (n = 3), support staff (n = 1), and correctional staff (n = 1).

Abbreviations: EMR, electronic medical records; LAI, long-acting injectable; MAR, medical administration records; OAP, oral antipsychotics; PSYCKES, Psychiatric Services and Clinical Knowledge Enhancement System.

## Factors Affecting Expected LAI Prescribing in 2021

Similar proportions of frequent LAI prescribers expected no change (44.1%) or an increase (42.9%) in LAI prescribing rates in 2021, while only 3.4% expected a decrease (Figure 7). Among those who expected that prescribing would remain unchanged (n = 180), the cost of treatment for patients (45.0%), followed by clinical staff availability to administer LAIs on site (38.9%), and the number of patients with schizophrenia followed at their clinical practice (37.8%) were cited as common factors impacting the expected dynamics of prescribing (Figure 8A). Among those who reported that prescribing would increase (n = 175), the number of patients with schizophrenia followed at their clinical practice (36.0%), the cost of treatment for patients (36.0%), and clinical staff availability to administer LAIs on site (33.7%) were common factors identified as impacting the expected dynamics of prescribing (Figure 8B). Among the responding prescribers who reported that prescribers who reported that prescribers who reported that prescribers who reported that prescribers who administer LAIs on site (37.1%) as a factor impacting the expected dynamics of prescribing (Figure 8C).

### Subgroup Analysis

The subgroups of moderately frequent LAI prescribers included 173 respondents, and the subgroup of very frequent prescribers included 233 respondents. A higher proportion of very frequent (80.7%) relative to moderately frequent (42.2%) LAI prescribers had >20% of patients with schizophrenia in their monthly psychiatric caseload. Higher

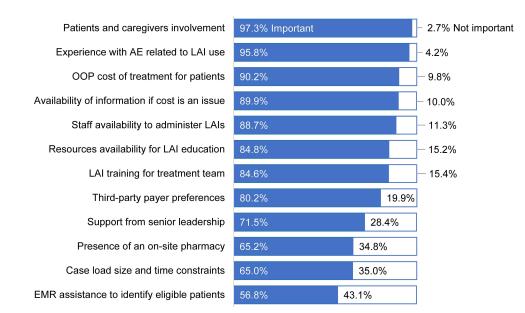
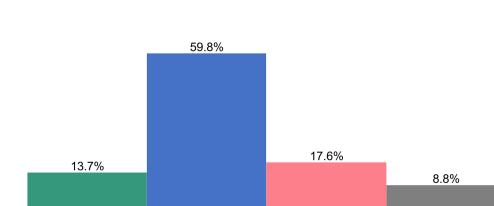
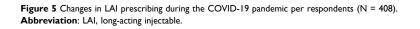


Figure 4 Factors impacting LAI prescribing in eligible patients for respondents (N = 408). Respondents rated the level of importance of each factor on a scale of I-10, with I being not at all important and 10 being very important. Values of 4 and above were classified as important.



Remained unchanged

Abbreviations: AE, adverse events; EMR, electronic medical records; LAI, long-acting injectable; OOP, out-of-pocket.



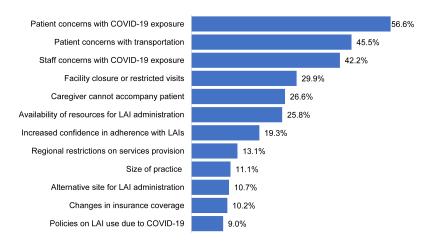
Increased

proportions of very frequent relative to moderately frequent LAI prescribers appeared to work in a community health center (31.3% vs 23.1%) or inpatient unit of a psychiatric hospital (18.5% vs 11.0%) and have LAI administration services (82.8% vs 56.1%) and staff available to inject LAIs (94.8% vs 81.5%) at their clinical practice. Higher proportions of very frequent relative to moderately frequent LAI prescribers appeared to collaborate on schizophrenia treatment plans with registered nurses (60.5% vs 40.5%) and social workers (56.2% vs 45.1%), as well as when prescribing LAIs at a patient's request (73.8% vs 63.0%) or for early-stage schizophrenia (49.4% vs 17.9%).

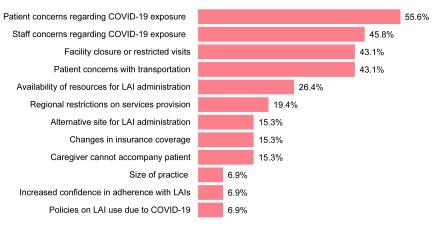
Decreased

Don't know

In addition, the two subgroups differed both with respect to the perception of change in LAI prescribing during the COVID-19 pandemic and the expected change in 2021. A higher proportion of the very frequent relative to moderately frequent LAI prescribers reported increased LAI prescribing during the COVID-19 pandemic (19.3% vs 6.4%). Very frequent LAI prescribers were also more likely to expect LAI prescribing to increase in 2021 than moderately frequent LAI prescribers (51.1% vs 31.8%).



Α





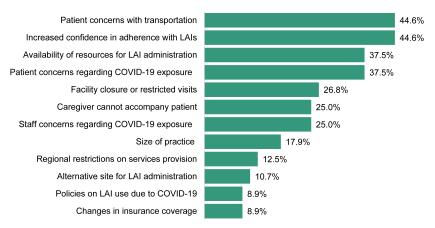


Figure 6 Factors contributing to changes observed in LAI prescribing during the COVID-19 pandemic. (A) Among those who reported that LAI prescribing remained unchanged (N = 244); (B) among those who reported that LAI prescribing decreased (N = 72); (C) among those who reported that LAI prescribing increased (N = 56). Abbreviation: LAI, long-acting injectable.

In terms of education on LAI use, higher proportions of very frequent relative to moderately frequent LAI prescribers reported that registered nurses (69.1% vs 48.0%) and patients or caregivers (24.5% vs 11.6%) received routine training on LAI use at their clinical practices and that registered nurses were involved in educating patients and caregivers on LAI use (69.5% vs 49.7%). Higher proportions of very frequent relative to moderately frequent LAI prescribers reported availability of various resources to educate patients and caregivers on LAIs, specifically, in-person conversations (84.1%)

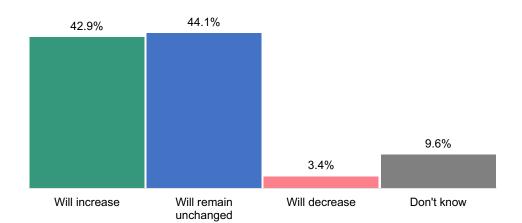


Figure 7 Expected changes in LAI prescribing during the next year (2021) per respondents (N = 408). Abbreviation: LAI, long-acting injectable.

vs 69.9%), paper information sheets (68.2% vs 48.6%), website referrals (48.9% vs 37.0%), phone calls (29.6% vs 20.8%), videos (22.3% vs 12.1%), and apps (9.4% vs 3.5%).

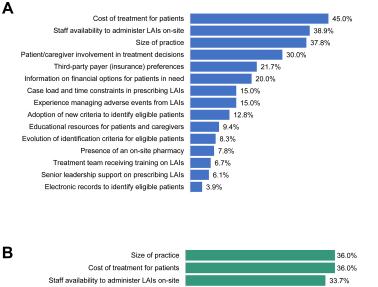
### Discussion

The existing literature on LAI prescribing in the US has focused on the clinical benefits (eg, lower discontinuation rates and higher number of prescription fills) or reduced healthcare resource utilization (eg, IP admissions, hospital days, and ER visits) associated with LAIs vs OAPs.<sup>7,22–24</sup> This study provides additional and timely insights into the characteristics of frequent LAI prescribers and the factors impacting their decision to prescribe LAIs, as well as historic and anticipated changes in LAI prescribing over time due to the impacts of the COVID-19 pandemic. Using the data collected in late 2020 across all US regions, this study reflects a recent treatment landscape for schizophrenia in the US, which evolves as therapeutic guidelines are updated.

In this study, frequent LAI prescribers were predominantly physicians with at least a decade of experience in psychiatry who were routinely trained on LAI use and had LAI administration services available on site at their clinical practice. Over half of these prescribers were treating more than 20% of their patients with schizophrenia with LAIs. To put this finding in context, only 10%–20% of antipsychotic treatment in the US involves LAIs (based on anecdotal reports).<sup>14,15</sup>

Poor adherence to antipsychotics was reported as a key circumstance for recommending LAIs. This finding is consistent with the current recommendations in the APA guideline,<sup>2</sup> which was the most followed guideline among the responding prescribers and their clinical practices. Interestingly, over one-third of frequent LAI prescribers and close to half of very frequent LAI prescribers reported that they also recommend the use of LAIs in the early stage of schizophrenia. While LAIs are typically used in patients with multiple relapses, literature and guidelines are increasingly shifting toward consideration of LAI use earlier in the course of illness to prevent future nonadherence and relapse.<sup>2,10</sup>

Most frequent prescribers reported recommending LAIs at the patients' request (69.4%; 73.8% in the very frequent prescriber subgroup). Patient/caregiver involvement in treatment decisions was consistently identified as a top consideration when initiating LAI prescribing in eligible patients, and almost 90% of frequent LAI prescribers reported inviting patients and caregivers to participate in treatment plan decisions in their clinical practice. These findings all suggest that joint decision-making with patients/caregivers has been adopted by the surveyed frequent LAI prescribers. This is in contrast to a prior study by Potkin et al that showed patient/caregiver involvement in 25% of psychiatrist-patient conversations about LAI prescription based on a smaller sample (N = 60) that predominately consisted of prescribers at community mental health centers in 2011–2012.<sup>18</sup> Thus, frequent LAI prescribers may place a higher value on patient/caregiver involvement compared to other prescribers, and the importance of this factor in prescribing decisions may have also increased over the past decade. Indeed, because LAI-naïve patients typically express indifferent or negative attitudes



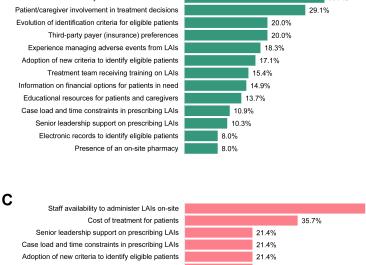




Figure 8 Factors contributing to expected changes in LAI prescribing over the next year (2021). (A) Among those who expected LAI prescribing to remain unchanged (N = 180); (B) among those who expected LAI prescribing to increase (N = 175); (C) among those who expected LAI prescribing to decrease (N = 14). Abbreviation: LAI, long-acting injectable.

towards LAIs,<sup>18</sup> patients may become more cooperative and accepting of LAIs when the treatment option is discussed in the context of shared decision-making.<sup>2</sup> Eventually, as demonstrated by Pietrini et al, switching adults with stabilized schizophrenia from OAPs to the equivalent LAIs could be associated with significant improvement in patient-reported attitude toward treatment, subjective experience of treatment, and health-related quality of life.<sup>25</sup> Taken together, patient education and shared decision-making are of utmost importance for the improvement of patient acceptance of LAIs and, subsequently, clinical outcomes during treatment.

With reduced access to healthcare due to the COVID-19 pandemic starting in 2020, there was concern that patients with schizophrenia would not receive in-person LAI treatment in a timely manner, which could lead to decompensation and relapse.<sup>20,21</sup> The majority of the frequent LAI prescribers surveyed reported that their LAI prescribing remained unchanged during the COVID-19 pandemic. This is consistent with a report by Gannon et al, who found that LAI prescribing largely remained stable before and after the implementation of social distancing and COVID-19 restrictions in March 2020 at a Pittsburgh ambulatory clinic.<sup>20</sup> Gannon et al cited established rapport between staff and patients and emphasis on the importance of ongoing adherence as key reasons for this lack of workflow and treatment disturbance, further emphasizing the significance of communication between healthcare professionals and patients for LAI prescribing behaviors, the current findings are encouraging and highlight the perceived value of continuing LAI treatment for patient stability, even during such emergency scenarios. Due to the complexity and unpredictable nature of the ongoing COVID-19 pandemic, further research is warranted to evaluate the changes it introduced to the treatment of patients with schizophrenia, as well as to understand its potential effect on health-related outcomes of patients with schizophrenia.

### Limitations

The findings of this study should be interpreted in light of some limitations. First, the analysis of prescriber characteristics, treatment patterns, and historical and anticipated future trends in this study was based on self-reporting by the participating prescribers, which may be subject to recall bias. Second, although attempts were made to minimize selection bias (eg, enrolling a relatively large number of participants from a nationwide panel), there remained a potential for bias given that a healthcare provider's decision to be on a panel and to contribute time to the survey was not random. Although frequent LAI prescribers from all US geographic regions were included, regions may not have been proportionally captured, so no statements regarding the national representativity of the study findings, or their generalizability to countries outside of the US, can be made. Lastly, due to the timeframe of this study, the results related to the COVID-19 pandemic only covered part of the first year of the pandemic.

## Conclusions

Over half of respondents who were frequent LAI prescribers in the US were treating  $\geq 20\%$  of their patients with schizophrenia with LAIs. Poor adherence to antipsychotics was a key circumstance for recommending LAIs, while patient/caregiver involvement in treatment decisions was a top consideration for prescribing LAIs in eligible patients. The majority of the frequent LAI prescribers surveyed reported that their LAI prescribing remained unchanged during the COVID-19 pandemic and expected LAI prescribing to remain unchanged or to increase in 2021. The insights from this study may influence strategies for ensuring routine use of LAIs in eligible patients.

## **Data Sharing Statement**

Due to the nature of this research, participants of this study did not agree for their data to be shared publicly, so supporting data are not available. Therefore, restrictions apply to the availability of these data, which are not publicly available.

## **Ethics Approval and Informed Consent**

Study participants provided their informed consent prior to responding to the survey questions. All study materials were approved by the New England Independent Review Board (now WIRB-Copernicus Group Independent Review Board).

## Acknowledgments

The authors thank the prescribers who participated in this survey. Medical writing assistance was provided by Christine Tam, an employee of Analysis Group, Inc., which provided consulting services to Janssen Scientific Affairs, LLC for the conduct of this study. Sophia Monaghan and Camilo Obando contributed to the study design, data collection, and results interpretation. Pamela Baker contributed to the study design and data collection. Sophia Monaghan was an employee of Janssen Scientific Affairs, LLC. Camilo Obando and Pamela Baker are employees of Janssen Scientific Affairs, LLC and

hold stock in Johnson & Johnson. Some data in this manuscript were presented virtually at the Psych Congress 2021, October 29 – November 1, 2021, in San Antonio, Texas, USA.

### Funding

This work was supported by Janssen Scientific Affairs, LLC.

### Disclosure

Maryia Zhdanava, Patrick Lefebvre, Todor I. Totev, Aditi Shah, Kristy Sheng, and Dominic Pilon are employees of Analysis Group, Inc., a company that provided consulting services to Janssen Scientific Affairs, LLC for the conduct of this study. H. Lynn Starr is an employee of Janssen Scientific Affairs, LLC and holds stock in Johnson & Johnson, Inc. The authors report no other conflicts of interest in this work.

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