ORIGINAL RESEARCH

Treatment Goals in Schizophrenia: A Real-World Survey of Patients, Psychiatrists, and Caregivers in the United States, with an Analysis of Current Treatment (Long-Acting Injectable vs Oral Antipsychotics) and Goal Selection

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¹Lundbeck LLC, Deerfield, IL, USA; ²Adelphi Real World, Bollington, UK; ³Otsuka Pharmaceutical & Commercial Development, Inc., Princeton, NJ, USA **Purpose:** To understand similarities and differences in patient treatment goals as selected by US psychiatrists, adult patients with schizophrenia, and their caregivers in a real-world setting in the United States, including stratification by current medication and age.

Patients and Methods: Data were drawn from the Adelphi Schizophrenia Disease Specific Programme[™], a point-in-time survey of psychiatrists and their consulting adult patients with schizophrenia, conducted from June to October 2019. Psychiatrists completed record forms for their next 8 consecutive outpatients and (where possible) 2 inpatients matching inclusion criteria. Participating psychiatrists, patients, and caregivers completed treatment goal questionnaires as part of the survey.

Results: Psychiatrists (n = 124) provided data on 1204 patients with schizophrenia, including 1135 on drug treatment (207 inpatients [18%] and 928 outpatients [82%]); questionnaires were completed by 555 patients and 135 caregivers. Decrease in disease symptoms was identified as the most important patient treatment goal by patients (64%), psychiatrists (selecting for 63% of patients), and caregivers (selecting for 68% of patients). Patients, psychiatrists, and caregivers similarly rated the least important goals (less sexual problems and less weight gain). Patients indicated their current medication helped to reach their most important goals: decrease in disease symptoms (68%) and thinking more clearly (39%). Findings based on analysis of treatment goals by treatment and age were similar to overall trends

Conclusion: These findings, including identification of a primary consensus goal of decrease in disease symptoms, may help with discussions between patients with schizophrenia, psychiatrists, and caregivers to inform effective management strategies and encourage shared decision-making.

Keywords: long-acting injectable antipsychotic, oral antipsychotic, real-world survey, schizophrenia, treatment goals, treatment preferences

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Introduction

Schizophrenia is a chronic and disabling severe mental illness with considerable impact, not only on patients' health, well-being, and functioning, but also on their surrounding environment.¹ Relapses are common among patients with schizophrenia;

thus, prevention of relapse and subsequent hospitalization is an important public health concern.^{2,3}

The introduction of long-acting injectable antipsychotics (LAIs) was an important step forward in addressing relapse and rehospitalization in patients with schizophrenia, but these agents remain underused in clinical practice. 4,5 Compared with oral antipsychotics (OAPs), LAIs may improve treatment adherence and continuity.² Based on the strong link between nonadherence and relapse, LAIs may thus potentially reduce risk of relapse, slow disease progression, and improve longterm outcomes.^{3,5–10} Currently, however, the American Psychiatric Association only recommends LAIs for patients who prefer such treatment or have a history of poor or uncertain adherence.^{4,5}

Survey data indicate that healthcare providers consider LAIs primarily for patients who have had repeated hospitalizations due to relapses, 11,12 although recent studies have demonstrated the advantage of LAIs over OAPs in first-episode or early-phase schizophrenia. 13-15 The recent US PRELAPSE study³ also showed that aripiprazole oncemonthly use in patients with early-phase schizophrenia can significantly delay first hospitalization compared with clinician's choice (treatment as usual, which could include LAIs, as well as OAPs).

Patient preferences related to treatment goals and outcomes in schizophrenia treatment, and the relative importance of certain treatment goals have been assessed in several studies, 16-18 including one focused on early-phase schizophrenia in patients from Germany, Italy, and the UK.¹⁹ Among the findings of these studies was that patients with schizophrenia judged improvement in positive symptoms as the most important potential treatment benefit. 17,18 and although psychiatrists and patients had general concordance on treatment goals, psychiatrists tended to emphasize traditional outcomes (for example, the absence of positive and negative symptoms), whereas patients placed greater value on global measures such as improved quality of life. 16 The early-phase European study by Bridges et al¹⁹ found that half of the 100 participants emphasized clinical goals, including reduced disease symptoms, while the remainder focused on functional goals; patients emphasizing functional goals were more likely to be receiving an LAI (44% vs 26%) and to state a preference for LAIs (46% vs 32%) compared with the overall group, which preferred OAPs to LAIs (61% vs 39%). In general, however, US data on patient attitudes and perceptions toward treatment goals and pharmacotherapy options in schizophrenia (particularly LAIs) are lacking. Additionally, greater understanding of the perspectives of

patients with schizophrenia, including the impact of disease severity on treatment goals, is needed in routine clinical practice.

Given that benefits of effective schizophrenia treatment may include improvements in both physical and psychosocial facets of life, a deeper understanding of how patients prioritize treatment goals may facilitate patient engagement and shared decision-making, inform effective clinical management strategies, and yield insight on patients' willingness to receive LAI treatment. 19 A US, real-world, point-in-time survey was undertaken to understand priorities in patient treatment goals as selected by patients with schizophrenia and their psychiatrists and caregivers, taking into consideration differences in patient treatment goals by treatment group (LAIs vs OAPs), age group, and disease severity.

Methods

Survey Study Design

Data were drawn from the Adelphi Schizophrenia Disease Specific ProgrammeTM, a point-in-time survey of psychiatrists and their consulting adult patients with schizophrenia conducted from June to October 2019 in the United States. Participating psychiatrists had to be actively involved in management and treatment of patients with schizophrenia; they completed questionnaires about themselves, their practice, and their workload, as well as patient record forms (PRFs) for their next 8 consecutively consulting outpatients and 2 consecutive inpatients who matched the inclusion criteria. If inpatients were not available, then 2 additional outpatients were allowed to be included in the survey by the responding psychiatrist. The PRFs included patient demographics, diagnosis information, disease severity, consultation and hospitalization history, and clinical and subjective assessments of the patients and their treatments. Participation of psychiatrists was financially incentivized, with reimbursement upon survey completion according to fair market research rates.

Patients were eligible if they were aged ≥18 years, had a physician-confirmed diagnosis of schizophrenia, and were not currently participating in a clinical study. Patients meeting inclusion criteria were invited by their psychiatrist to voluntarily complete a patient self-completion questionnaire (PSC). If a caregiver was present at the consultation for a patient for whom the psychiatrist had completed a PRF, the caregiver was invited to voluntarily complete a caregiver self-completion questionnaire (CSC).

A complete description of the methods of the survey has been previously published and validated.²⁰ Using a check box, participating patients, psychiatrists, and caregivers provided informed consent for use of their anonymized and aggregated data for research and publication in scientific journals. Psychiatrists provided informed consent during their self-completion of an electronic screening questionnaire; patients and caregivers provided their informed consent via a check box on the front of a selfcompleted pen-and-paper questionnaire, which was provided by the psychiatrists. Data were collected in such a way that respondents could not be identified directly; all data were aggregated and de-identified before receipt. The survey was conducted according to the EphMRA Code of Conduct and was approved by the Western Institutional Review Board (study protocol number AG8618).

Among other questions, psychiatrists, patients, and caregivers were each asked to select the 3 most important and 3 least important treatment goals for the patient from a list of 13 possible goals. Patients and caregivers also chose up to 3 treatment goals that the current medication helped to reach and up to 3 goals that were not met by the current medication. The full list of treatment goals included in the PRF, PSC, and CSC is provided in Table 1. Psychiatrist-reported data from the survey reflect the percentage of patients under the care of the responding psychiatrist for whom the

Table I Potential Treatment Goals Listed in Survey

- Decrease in disease symptoms (hallucinations and/or delusions)
- Think more clearly (able to concentrate)
- Fuller range of happy and sad emotions (feel more deeply)
- Less restlessness or urges to move (less unrest or feeling fidgety)
- Decrease in hospitalizations for relapse
- Less sexual problems (less sexual dysfunction or intimacy issues)
- Increased interest in work, studies, and/or hobbies (daily activities)
- Improved interaction and communication with others
- · Less weight gain
- Increased ability to take care of self (being independent, able to cook, clean, etc)
- Spend more time with family and friends (socialize more often)
- Less anxious and more relaxed (able to handle stressful situations)
- Less fatigue or feeling slowed down (missing energy)

psychiatrist had completed a PRF. For patients and caregivers, the percentages reported reflect the percentage of responding patients and caregivers.

In each PRF, psychiatrists rated disease severity using the 7-point Clinical Global Impression-Severity scale (CGI-S)²¹ to answer the question: "Considering your total clinical experience with this particular population, how mentally ill is the patient at this time?" The choices available were: 1 = normal. not at all ill; 2 = borderline mentally ill; 3 = mildly ill; 4 = moderately ill; 5 = markedly ill; 6 = severely ill; and 7 = markedly illamong the most extremely ill patients. As part of the PSC, patients completed the Work Productivity and Activity Impairment questionnaire (WPAI) about employment, absenteeism, impairment at work, productivity loss, and activity impairment.²² Caregivers also completed the WPAI in relation to how the patient's condition has impacted their employment and activities. Differences in patient treatment goals stratified by treatment group (LAIs vs OAPs), age group (18–35, 36–65, and >65 years), and disease severity based on CGI-S²³ and WPAI²⁴ were also assessed.

Statistical Methods

This was an exploratory survey and the statistical analyses were primarily descriptive in nature. Descriptive statistics (mean, standard deviation, frequency) are presented for demographic and clinical information to describe the survey sample. Categorical variables are presented as frequency and percentage distributions when appropriate. Categorical variables were compared using either Fisher's exact test for variables with two categories or chi-squared test for variables with more than two categories. Ordinal variables are reported as frequencies and percentages and/or medians and interquartile ranges, as appropriate for the individual variables. Continuous variables (for example, age, time since diagnosis, and questions with numeric rating scale responses) are presented as means and standard deviations, medians and interquartile ranges, and score ranges. Continuous variables from two groups were compared using the t-test or Mann–Whitney test depending on the distribution.

Data completeness are indicated by frequencies and percentages. The majority of variables for this analysis represent observed responses to individual closed-ended questions. These variables were analyzed as observed, with no imputation of missing data or aggregation across questions. Analyses that required statistical comparisons were conducted using STATA® 10 (StataCorp LP, College Station, Texas) and SPSS® 15.0 (IBM®, Armonk, New York).

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Results

Survey Participants

Overall, 124 psychiatrists provided data for 1204 patients with schizophrenia, including 1135 on drug treatment comprising 207 inpatients (18%) and 928 outpatients (82%). A total of 247 patients (34%) had a caregiver. In all, 555 patients completed a PSC and 135 caregivers completed a CSC. Of the 1135 patients on drug treatment, more were receiving an OAP (884 [78%]) than a LAI (251 [22%]). A significantly greater proportion of patients in the LAI-treatment group (68 [27%]) than in the OAPtreatment group (139 [16%]) were being treated as inpatients (P < 0.001).

Patient Demographics and Characteristics

Patient demographics and characteristics reported by psychiatrists are presented in Table 2. Compared with patients receiving OAPs, those on LAIs had a significantly longer mean time since their schizophrenia diagnosis (10.3 vs 7.8 years; P = 0.031) and a shorter mean time on their current medication regimen (1.7 vs 2.5 years; P = 0.009). More patients treated with LAIs were unable to finish their education due to schizophrenia (51% vs 38%; P = 0.002) and did not complete high school (25% vs 16%; P =0.003), and fewer were employed full-time (4% vs 11%; P < 0.001). Furthermore, more patients on LAIs were unemployed due to disability (56% vs 40%; P <0.001) and had a caregiver (37% vs 26%; P = 0.001) than those taking OAPs.

Differences in Patient Treatment Goals by Participant Group

Data for patient treatment goals most frequently selected as being most important and least important are presented in Figure 1 (top 5 goals selected based on average of patient, psychiatrist, and caregiver responses; data for all 13 goals are shown in Supplementary Table 1). "Decrease in disease symptoms" was most frequently selected as one of the top three choices for most important treatment goals for patients (64%), psychiatrists (selected for 63% of patients), and caregivers (selected for 68% of patients; Figure 1A). "Decrease in hospitalizations for relapse" was the second most important patient treatment goal for both psychiatrists (selected for 41% of patients) and caregivers (selected for 38% of patients), but was the seventh most important goal for patients (21%). The second most important treatment goal for patients (47%) was to "think more clearly," a goal that was the fourth most important for psychiatrists (selected for 29% of patients) and caregivers (selected for 29% of patients). For least important treatment goals, patients, psychiatrists, and caregivers all included "less sexual problems" (43-59%) and "less weight gain" (38-44%) among their top 3 choices (Figure 1B).

Patient treatment goals "met" and "not met" by the current medication as reported by patients and caregivers are presented in Figure 2 (top 5 goals selected based on average of patient and caregiver responses; data for all 13 goals are shown in Supplementary Table 2). Patients indicated that the treatment goals "decrease in disease symptoms" (68%) and "think more clearly" (39%) were met by their current medication (Figure 2A). Caregivers reported that the patient treatment goals "decrease in disease symptoms" (70%) and "decrease in hospitalizations for relapse" (41%) were met by patients' current medication. A greater proportion of caregivers (30%) than patients (11%) identified "think more clearly" as a treatment goal that was not being met by the patients' current medication (Supplementary Table 2).

Differences in Treatment Goals Stratified by Treatment

Focusing on patient-reported treatment goals by drug therapy, patients in each treatment group reported similar preferences for their most important treatment goals ("decrease in disease symptoms" [OAPs: 65%; LAIs: 62%] and "think more clearly" [OAPs: 46%; LAIs: 53%]) and least important treatment goals ("less sexual problems" [OAPs: 42%; LAIs: 48%] and "less weight gain" [OAPs: 43%; LAIs: 46%]; Table 3).

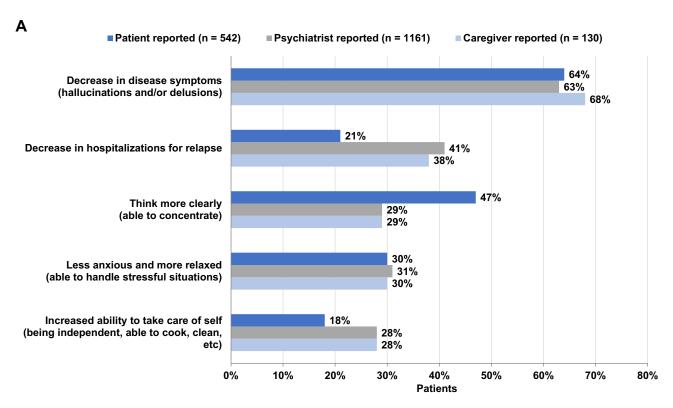
A higher proportion of patients receiving LAIs than those receiving OAPs rated "fuller range of happy and sad emotions" (36% vs 28%) and "increased ability to take care of self" (22% vs 17%) as the most important treatment goals and "improved interaction and communications" (24% vs 16%) as a least important treatment goal. "Improved interaction and communication" was also more frequently an unmet treatment goal in patients receiving LAIs vs OAPs (24% vs 16%). "Less restlessness or urges to move" was chosen as a least important treatment goal by a higher percentage of patients taking OAPs than LAIs (39% vs 30%) and was more frequently indicated as an unmet goal with the current treatment (39% vs 30%; Table 4). Furthermore, patients taking LAIs reported more frequently that their current medication helped to

Table 2 Patient Demographics and Characteristics

Parameter	Overall	OAPs	LAIs	P value
	(N = 1135)	(n = 884)	(n = 251)	(OAPs vs LAIs)
Mean age, y (SD)	40.0 (15.8)	40.4 (16.2)	38.8 (14.3)	0.15 (TT)
Sex, n (%)				0.83 (FE)
Female	470 (41)	368 (42)	102 (41)	
Male	665 (59)	516 (58)	149 (59)	
Mean BMI, kg/m² (SD)	27.4 (4.9)	27.2 (4.8)	27.8 (5.4)	0.15 (TT)
Race/ethnicity, n (%)				0.25 (CH)
White	697 (61)	559 (63)	138 (55)	
African-American	267 (24)	191 (22)	76 (30)	
Native-American	7 (<1)	6 (<1)	l (<l)< td=""><td></td></l)<>	
Asian	42 (4)	35 (4)	7 (3)	
Hispanic/Latino	73 (6)	54 (6)	19 (8)	
Middle Eastern	19 (2)	16 (2)	3 (1)	
Mixed race	27 (2)	21 (2)	6 (2)	
Other	3 (<1)	2 (<1)	l (<i)< td=""><td></td></i)<>	
Mean time since diagnosis, y (SD)	8.2 (10.8)	7.8 (10.5)	10.3 (12.0)	0.031 (TT)
Mean time on current regimen, y (SD)	2.3 (3.6)	2.5 (3.9)	1.7 (1.9)	0.009 (TT)
Education level, n (%)				0.003 (CH)
Did not complete high school	200 (18)	138 (16)	62 (25)	
High school diploma/GED	572 (50)	442 (50)	130 (52)	
College degree: 2 y	202 (18)	169 (19)	33 (13)	
College degree: 4 y	95 (8)	80 (9)	15 (6)	
Graduate degree	28 (2)	26 (3)	2 (<1)	
Trade school/certificate program	25 (2)	18 (2)	7 (3)	
Other	13 (1)	11 (1)	2 (<1)	
Unable to finish education due to schizophrenia	353 (41)	254 (38)	99 (51)	0.002 (FE)
Employment status				<0.001 (CH)
Full time, n (%)	109 (10)	99 (11)	10 (4)	
Part time, n (%)	253 (23)	201 (23)	52 (21)	
Homemaker	43 (4)	30 (4)	13 (5)	
Student	80 (7)	71 (8)	9 (4)	
Retired	64 (6)	57 (7)	7 (3)	
Unemployed due to disability, n (%) ^a	481 (43)	343 (40)	138 (56)	
Unemployed due to other reason, n (%)	59 (5)	50 (6)	9 (4)	
On long-term sick leave, n (%)	14 (1)	8 (I)	6 (2)	
Long-term sick leave/retired/ unemployed due to schizophrenia, n (%) ^b	499 (88)	358 (85)	141 (95)	<0.001 (FE)
Mean duration of sick leave/retirement/ unemployment, mo (SD)	11.9 (10.7)	11.7 (10.5)	12.7 (11.1)	0.43 (TT)
Patient has caregiver, n (%)	309 (29)	221 (26)	88 (37)	0.001 (FE)
Professional, n (%)	89 (29)	62 (28)	27 (31)	0.68 (FE)
Mean h/wk providing care (professional caregiver) (SD)	51.2 (48.5)	58.4 (54.2)	34.6 (25.5)	0.033 (TT)

Notes: ^aPercentages based on total n of 1114 (following removal of "don't know" responses). ^bPercentages based on total n of 568 (total number of patients on long-term sick leave/retired/unemployed due to disability, following removal of "don't know" responses).

Abbreviations: BMI, body mass index; CH, chi-square test; FE, Fisher's exact test; GED, general education development; LAIs, long-acting injectable antipsychotics; OAPs, oral antipsychotics; SD, standard deviation; TT, t-test.



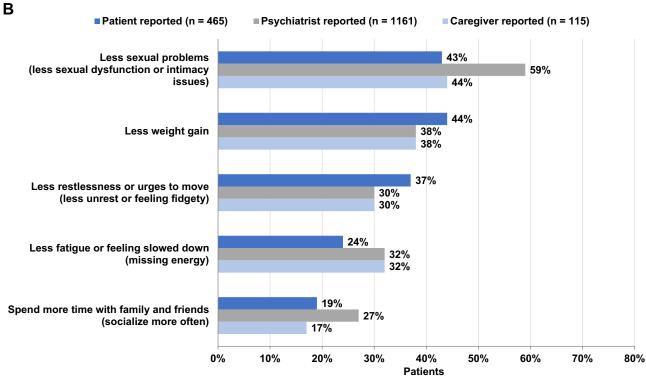


Figure I Most important (A) and least important (B) Patient treatment goals. Respondents were instructed to select the top 3 most important treatment goals and the top 3 least important treatment goals for patients; categories represent the 5 most frequently selected treatment goals (based on average of patient, psychiatrist, and caregiver responses).

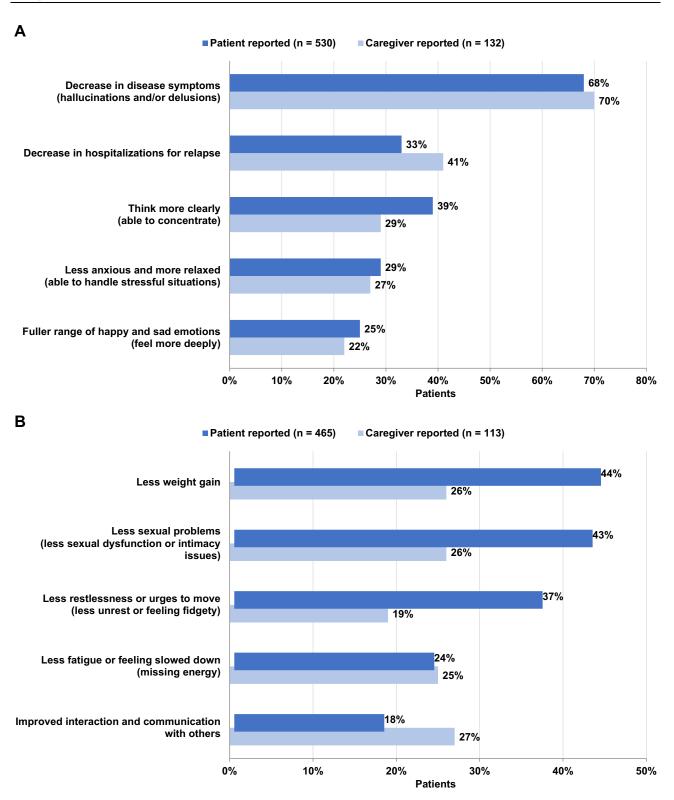


Figure 2 Patient treatment goals met (A) And not met (B) by current medication. Respondents were instructed to select up to 3 goals; categories represent the 5 most frequently selected goals (based on average of patient and caregiver responses).

Table 3 Most Important and Least Important Patient Treatment Goals by Responder Group and Treatment^a

Most Important Treatment Goals, n (%)	Patient Reported		Psychiatrist Reported		Caregiver Reported	
	OAPs (n = 418)	LAIs (n = 115)	OAPs (n = 891)	LAIs (n = 244)	OAPs (n = 90)	LAIs (n = 30)
Decrease in disease symptoms (hallucinations and/or delusions)	273 (65)	71 (62)	556 (62)	166 (68)	63 (70)	23 (77)
Decrease in hospitalizations for relapse	86 (21)	28 (24)	336 (38)	128 (52)	30 (33)	16 (53)
Increased ability to take care of self (being independent, able to cook, clean, etc)	72 (17)	25 (22)	248 (28)	74 (30)	22 (24)	7 (23)
Think more clearly (able to concentrate)	192 (46)	61 (53)	264 (30)	65 (27)	30 (33)	7 (23)
Less anxious and more relaxed (able to handle stressful situations)	121 (29)	37 (32)	303 (34)	56 (23)	29 (32)	7 (23)
Improved interaction and communication with others	95 (23)	24 (21)	213 (24)	60 (25)	23 (26)	8 (27)
Increased interest in work, studies, and/or hobbies (daily activities)	103 (25)	28 (24)	203 (23)	46 (19)	11 (12)	4 (13)
Spend more time with family and friends (socialize more often)	69 (17)	15 (13)	165 (19)	35 (14)	23 (26)	7 (23)
Less weight gain	68 (16)	18 (16)	96 (11)	35 (14)	11 (12)	5 (17)
Fuller range of happy and sad emotions (feel more deeply)	118 (28)	41 (36)	93 (10)	24 (10)	21 (23)	7 (23)
Less restlessness or urges to move (less unrest or feeling fidgety)	39 (9)	15 (13)	92 (10)	21 (9)	8 (9)	3 (10)
Less fatigue or feeling slowed down (missing energy)	42 (10)	12 (10)	75 (8)	13 (5)	7 (8)	2 (7)
Less sexual problems (less sexual dysfunction or intimacy issues)	48 (11)	10 (9)	29 (3)	9 (4)	2 (2)	2 (7)
Least Important Treatment Goals, n (%)	Important Treatment Goals, n (%) Patient Repor	Reported Psychiatrist Reported			Caregiver Reported	
	OAPs (n = 359)	LAIs (n = 97)	OAPs (n = 891)	LAIs (n = 244)	OAPs (n = 78)	LAIs (n = 27)
Less sexual problems (less sexual dysfunction or intimacy issues)	151 (42)	47 (48)	535 (60)	138 (57)	36 (46)	10 (37)
Less weight gain	153 (43)	45 (46)	338 (38)	96 (39)	31 (40)	11 (41)
Less fatigue or feeling slowed down (missing energy)	89 (25)	20 (21)	282 (32)	86 (35)	27 (35)	10 (37)
Less restlessness or urges to move (less unrest or feeling fidgety)	139 (39)	29 (30)	266 (30)	76 (31)	25 (32)	5 (19)
Spend more time with family and friends (socialize more often)	69 (19)	19 (20)	237 (27)	72 (30)	15 (19)	4 (15)
Fuller range of happy and sad emotions (feel more deeply)	59 (16)	18 (19)	195 (22)	58 (24)	7 (9)	4 (15)
Increased interest in work, studies, and/or hobbies (daily activities)	53 (15)	15 (15)	152 (17)	49 (20)	16 (21)	4 (15)
Less anxious and more relaxed (able to handle stressful situations)	49 (14)	10 (10)	148 (17)	38 (16)	12 (15)	9 (33)

(Continued)

Table 3 (Continued).

Improved interaction and communication with others	58 (16)	23 (24)	133 (15)	36 (15)	16 (21)	6 (22)
Increased ability to take care of self (being independent, able to cook, clean, etc)	54 (15)	15 (15)	153 (17)	32 (13)	17 (22)	5 (19)
Think more clearly (able to concentrate)	42 (12)	8 (8)	99 (11)	25 (10)	7 (9)	I (4)
Decrease in hospitalizations for relapse	92 (26)	23 (24)	87 (10)	17 (7)	14 (18)	4 (15)
Decrease in disease symptoms (hallucinations and/or delusions)	24 (7)	5 (5)	48 (5)	9 (4)	1 (1)	4 (15)

Notes: ^aRespondents were instructed to select the top 3 most important and top 3 least important goals for patients; goals are presented in decreasing order of response rate for the psychiatrist group (average of oral antipsychotic [OAP] and long-acting injectable antipsychotics [LAI] subgroups), ie, the goal most commonly selected by psychiatrists is presented in the top row; psychiatrist-reported data reflect the percentage of patients for whom the goal was identified.

achieve a "decrease in hospitalizations for relapse" than patients taking OAPs (39% vs 31%).

In both treatment groups, patients' most important treatment goals tended to overlap with the goals they selected as having been met by their current medication, and their least important treatment goals tended to coincide with those selected as not having been met by their current medication (Figures 1 and 2).

When treatment goals for the patient (most/least important; met/unmet by treatment) were examined by responder type (psychiatrist, patient, and caregiver), few differences were seen between goals for patients on LAIs and those on OAPs, similar to the trends observed for the overall (combined) responder population (Tables 3 and 4).

A "decrease in disease symptoms" was identified as an important patient treatment goal for a larger proportion of patients on LAIs vs OAPs based on responses from psychiatrists (68% vs 62%) and caregivers (77% vs 70%; Table 3). These responder groups also selected "less anxious and more relaxed" as an important treatment goal for a larger proportion of patients taking OAPs compared with LAIs (psychiatrists: 34% vs 23%; caregivers: 32% vs 23%).

Caregivers more often identified the most important patient treatment goals for patients receiving LAIs compared with OAPs as "decrease in disease symptoms" (77% vs 70%) and "decrease in hospitalizations for relapse" (53% vs 33%; Table 3), and reported larger proportions of patients receiving LAIs vs OAPs met these goals (80% vs 74% and 63% vs 35%, respectively; Table 4). For patients receiving OAPs vs LAIs, a larger proportion of caregivers chose "think more clearly" as an important treatment goal for the patient (33% vs 23%; Table 3).

Differences in Treatment Goals by Age Group

Overall, no significant differences in patient treatment goals by age group were identified and findings were generally similar to overall trends. Psychiatrists (selecting goals for 40% of patients) were more concerned about the ability of their patients aged >65 years to take care of themselves than either caregivers or patients (both 31%). Sexual problems were categorized as a lesser concern for patients aged >65 years by all 3 responder groups (patients, psychiatrists, and caregivers).

Differences in Treatment Goals by Psychiatrist-Reported Clinical Global Impression-Severity Scale Score and Treatment

Surveyed psychiatrists reported that 74% of all patients in the survey who were currently receiving LAIs (n = 239)had moderate-to-severe disease (CGI-S scores of 4-7) compared with 55% of patients receiving OAPs (n = 834). Similar levels of CGI improvement (much improved/very much improved) were reported for both treatment groups (68-69%). For moderately to severely ill patients receiving LAIs (grouping moderately, markedly, severely, and extremely ill patients: n = 178) compared with their counterparts receiving LAIs who had mild disease (grouping borderline and mildly ill patients: n = 61), psychiatrists more commonly chose "decrease in disease symptoms" (71% vs 52%) and "think more clearly" (31% vs 18%) as the most important treatment goals. For patients with mild disease severity compared with those with more severe disease, "increased interest in work,

Table 4 Patient Treatment Goals Met or Not Met by Current Medication by Responder Group and Treatment^a

Treatment Goals Helped to Reach by Current Medication, n (%)	Patient I	Reported	Caregiver Reported		
	OAPs (n = 410)	LAIs (n = 112)	OAPs (n = 92)	LAIs (n = 30)	
Decrease in disease symptoms (hallucinations and/or delusions)	282 (69)	75 (67)	68 (74)	24 (80)	
Think more clearly (able to concentrate)	162 (40)	41 (37)	29 (32)	7 (23)	
Decrease in hospitalizations for relapse	129 (31)	44 (39)	32 (35)	19 (63)	
Less anxious and more relaxed (able to handle stressful situations)	118 (29)	36 (32)	26 (28)	8 (27)	
Fuller range of happy and sad emotions (feel more deeply)	100 (24)	28 (25)	19 (21)	8 (27)	
Increased ability to take care of self (being independent, able to cook, clean, etc)	80 (20)	23 (21)	17 (18)	4 (13)	
Increased interest in work, studies, and/or hobbies (daily activities)	87 (21)	16 (14)	13 (14)	5 (17)	
Less restlessness or urges to move (less unrest or feeling fidgety)	51 (12)	19 (17)	4 (4)	2 (7)	
Improved interaction and communication with others	73 (18)	11 (10)	19 (21)	6 (20)	
Spend more time with family and friends (socialize more often)	59 (14)	16 (14)	22 (24)	6 (20)	
Less weight gain	38 (9)	14 (13)	6 (7)	4 (13)	
Less fatigue or feeling slowed down (missing energy)	38 (9)	11 (10)	8 (9)	2 (7)	
Less sexual problems (less sexual dysfunction or intimacy issues)	21 (5)	11 (10)	2 (2)	I (3)	
Treatment Goals Not Met by Current Medication, n (%)	Patient Reported		Caregiver Reported		
·					
	OAPs (n = 359)	LAIs (n = 97)	OAPs (n = 78)	LAIs (n = 25)	
Less sexual problems (less sexual dysfunction or intimacy issues)					
Less sexual problems (less sexual dysfunction or intimacy issues) Less weight gain	(n = 359)	(n = 97)	(n = 78)	(n = 25)	
. , , , , , , , , , , , , , , , , , , ,	(n = 359)	(n = 97) 47 (48)	(n = 78) 22 (28)	(n = 25) 3 (12)	
Less weight gain	(n = 359) 151 (42) 153 (43)	(n = 97) 47 (48) 45 (46)	(n = 78) 22 (28) 21 (27)	(n = 25) 3 (12) 7 (28)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety)	(n = 359) 151 (42) 153 (43) 139 (39)	(n = 97) 47 (48) 45 (46) 29 (30)	(n = 78) 22 (28) 21 (27) 15 (19)	(n = 25) 3 (12) 7 (28) 2 (8)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse Less fatigue or feeling slowed down (missing energy)	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26) 89 (25)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24) 20 (21)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8) 21 (27)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12) 7 (28)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse Less fatigue or feeling slowed down (missing energy) Improved interaction and communication with others	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26) 89 (25) 58 (16)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24) 20 (21) 23 (24)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8) 21 (27) 19 (24)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12) 7 (28) 7 (28)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse Less fatigue or feeling slowed down (missing energy) Improved interaction and communication with others Spend more time with family and friends (socialize more often)	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26) 89 (25) 58 (16) 69 (19)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24) 20 (21) 23 (24) 19 (20)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8) 21 (27) 19 (24) 16 (21)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12) 7 (28) 7 (28) 7 (28) 8 (32)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse Less fatigue or feeling slowed down (missing energy) Improved interaction and communication with others Spend more time with family and friends (socialize more often) Fuller range of happy and sad emotions (feel more deeply) Increased ability to take care of self (being independent, able to cook,	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26) 89 (25) 58 (16) 69 (19) 59 (16)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24) 20 (21) 23 (24) 19 (20) 18 (19)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8) 21 (27) 19 (24) 16 (21)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12) 7 (28) 7 (28) 7 (28) 8 (32) 5 (20)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse Less fatigue or feeling slowed down (missing energy) Improved interaction and communication with others Spend more time with family and friends (socialize more often) Fuller range of happy and sad emotions (feel more deeply) Increased ability to take care of self (being independent, able to cook, clean, etc)	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26) 89 (25) 58 (16) 69 (19) 59 (16) 54 (15)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24) 20 (21) 23 (24) 19 (20) 18 (19) 15 (15)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8) 21 (27) 19 (24) 16 (21) 17 (22)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12) 7 (28) 7 (28) 7 (28) 8 (32) 5 (20) 7 (28)	
Less weight gain Less restlessness or urges to move (less unrest or feeling fidgety) Decrease in hospitalizations for relapse Less fatigue or feeling slowed down (missing energy) Improved interaction and communication with others Spend more time with family and friends (socialize more often) Fuller range of happy and sad emotions (feel more deeply) Increased ability to take care of self (being independent, able to cook, clean, etc) Increased interest in work, studies, and/or hobbies (daily activities)	(n = 359) 151 (42) 153 (43) 139 (39) 92 (26) 89 (25) 58 (16) 69 (19) 59 (16) 54 (15) 53 (15)	(n = 97) 47 (48) 45 (46) 29 (30) 23 (24) 20 (21) 23 (24) 19 (20) 18 (19) 15 (15)	(n = 78) 22 (28) 21 (27) 15 (19) 6 (8) 21 (27) 19 (24) 16 (21) 17 (22) 21 (27)	(n = 25) 3 (12) 7 (28) 2 (8) 3 (12) 7 (28) 7 (28) 7 (28) 8 (32) 5 (20) 7 (28) 6 (24)	

Notes: a Respondents were instructed to select up to 3 goals; goals are presented in decreasing order of response rate for the patient group (average of oral antipsychotic [OAP] and long-acting injectable antipsychotic [LAI] subgroups), ie, the goal most commonly selected by patients is presented in the top row.

studies, and/or hobbies" (33% vs 16%), "spend more time with family and friends" (28% vs 11%), and "less anxious and more relaxed" (25% vs 18%) were more common treatment goals selected by psychiatrists.

Unemployment among patients was higher with moderate and severe disease than mild disease (mild disease: 63% [n = 35; normal/borderline/mildly ill]; moderate disease: 84% [n = 49]; severe disease: 82% [n = 28; markedly/severely/extremely ill]). Likewise, in patients who completed the WPAI questionnaire, the mean proportion of activity impairment due to schizophrenia increased with disease severity (mild: 32% [n = 32]; moderate: 49% [n = 47]; severe: 56% [n = 28]).

For patients with severe disease compared with those with mild disease, psychiatrists more often chose "decrease in disease symptoms" (71% vs 52%), "decrease in hospitalizations for relapse" (59% vs 54%), "increased ability to take care of self" (41% vs 31%), and "think more clearly" (34% vs 18%) as the most important treatment goals. For patients with mild disease compared with their counterparts with severe disease, psychiatrists more often chose social treatment goals: "increased interest in work, studies, and/or hobbies" (33% vs 12%), "spend more time with family and friends" (28% vs 7%), and "less anxious and more relaxed" (25% vs 16%).

Regardless of disease severity (mild [normal/border-line/mildly ill]: n = 61; moderate: n = 102; severe [mark-edly ill/severely ill/extremely ill]: n = 76), the top 2 patient treatment goals selected by psychiatrists for patients were "decrease in disease symptoms" (52%, 72%, and 71%, respectively) and "decrease in hospitalizations for relapse" (54%, 52%, and 59%, respectively).

Work Productivity and Activity Impairment Questionnaire: Caregiver Assessments

Results for the WPAI questionnaire showed that there were no significant differences among patients receiving LAIs vs OAPs with respect to work time missed, impairment while working, overall work impairment, or activity impairment as assessed by caregivers.

Discussion

Based on the results of this exploratory, real-world survey of US psychiatrists, patients, and their caregivers, patients with schizophrenia who were receiving LAIs appeared to have a longer duration of disease since their diagnosis and greater disease severity, and were significantly more likely to have a caregiver compared with patients taking OAPs. The survey results suggest that LAIs may represent an underused treatment option in clinical practice (in other words, these agents are mainly prescribed to patients who are later in the disease course with more severe illness), despite a growing body of evidence of utility in younger patients who are earlier in the course of their illness; 3,9,14,15,25-27 however, longitudinal studies of prescribing patterns over the course of disease would be required to determine if this were true. Similar proportions of patients in each treatment group chose "decrease in disease symptoms" and "think more clearly" as their most important treatment goals; "less sexual problems" and "less weight gain" were selected as the least important. Caregivers reported that the treatment goal of "decrease in hospitalizations for relapse" was met more frequently for patients treated with LAIs than with OAPs. There was consensus among patients, psychiatrists, and caregivers on the most important treatment goal—"decrease in disease symptom"-regardless of patients' current medication or age. This treatment goal was also the top choice of psychiatrists for patients with moderate and severe schizophrenia.

Identification of a primary consensus goal—"decrease in disease symptoms"—across the patient journey may help with discussions between patients, psychiatrists, and caregivers to inform effective management strategies early in the course of illness and encourage shared decisionmaking. The untapped potential of this collaborative process was highlighted by the findings of a study by Potkin et al.²⁸ which found that the majority (67%) of patients and caregivers were not involved in antipsychotic treatment decision-making, and that psychiatrists offered LAIs as a treatment option to their patients only 50% of the time, despite all of the patients being indicated for a change in therapy. The authors of the study stressed the importance of increasing patient engagement in the decision-making process, as well as providing both psychiatrists and patients with more information on the simplicity, safety, and tolerability of LAIs, including positive feedback based on experiences/outcomes, to reduce resistance to new therapies on the part of patients and their providers. More recently, the PRELAPSE study^{3,29} concluded that proper training of practitioners can facilitate effective communications with patients regarding the advantages of LAIs and foster engagement in shared decisionmaking, resulting in high LAI acceptance rates even in early-stage disease.

The analysis by disease severity in the present survey showed a clear shift in psychiatrists' perceptions regarding their patients' most important treatment goals moving from one end of the disease severity spectrum to the other. The top goals selected for their patients with mild disease were more focused on improving productivity and social functioning ("increased interest in work, studies, and/or hobbies." "spend more time with family and friends," and "less anxious and more relaxed"), whereas the goals selected for their patients with severe disease were more focused on symptoms and activities of daily living ("decrease in disease symptoms," "decrease in hospitalizations for relapse," and "increased ability to take care of self"). As expected, unemployment and disability were shown to increase in parallel with disease severity. Information regarding the correlation of disease severity and treatment goals, in conjunction with the broader analysis of treatment goals discussed above, can be crucial to psychiatrists as they engage with patients and caregivers in the shared decision-making process regarding the optimal individualized treatment strategy.

There were several limitations associated with this survey. First, the only patients included in the sample were consecutively consulting patients of the surveyed psychiatrists who met the patient inclusion criteria, and they may not fully represent the overall US population of patients with schizophrenia. In addition, patients who consult their psychiatrist frequently were more likely to be included in this sample. Second, although psychiatrists were requested to collect data on a series of consecutively consulting patients to avoid selection bias, in the absence of randomization, adherence was contingent on the integrity of the participating psychiatrists rather than via a formalized source-verification procedure. Moreover, diagnosis in the target patient group was based primarily on the judgement and diagnostic skills of the respondent psychiatrist rather than on a formalized diagnostic checklist, although patients were managed in accordance with the same routine diagnostic procedures representative of that clinical practice setting. Third, the quality of data was largely dependent on the accurate reporting of information by psychiatrists, patients, and caregivers, which may be subject to recall bias. Fourth, the inclusion criteria that governed the selection of psychiatrists (treatment of a minimum number of patients with schizophrenia and active involvement in patient management and treatment decisions) were minimal and, consequently, inclusion was likely influenced by willingness to take part in the survey. Selection bias was therefore possible because the participating psychiatrists were a pragmatic sample and may not represent the overall population of psychiatrists treating patients with schizophrenia. Finally, due to the point-in-time design of the survey, it cannot be used to demonstrate cause and effect.

Conclusions

This survey identified a primary consensus goal of a decrease in disease symptoms across the patient journey, as well as partial discordance among the responder groups with respect to their top 3 treatment goals. The findings of this survey highlight the importance of shared decision-making and may help facilitate and encourage discussions among patients, psychiatrists, and caregivers to inform effective management strategies early in the course of illness.

Abbreviations

CGI-S, Clinical Global Impression-Severity scale (CGI-S); CSC, caregiver self-completion questionnaire; LAI, long-acting injectable antipsychotic; OAP, oral antipsychotic; PRF, patient record form; PSC, patient self-completion questionnaire; WPAI, Work Productivity and Activity Impairment questionnaire.

Data Sharing Statement

All data that support the findings of this study are the intellectual property of Adelphi Real World. Legitimate research requests for supportive data will be considered. Requests for access should be addressed directly to Jason Shepherd at jason.shepherd@adelphigroup.com.

Ethics Approval and Informed Consent

Participating patients, psychiatrists, and caregivers provided informed consent for use of their anonymized and aggregated data for research and publication in scientific journals. Data were collected in such a way that respondents could not be identified directly; all data were aggregated and de-identified before receipt. The survey was conducted according to the EphMRA Code of Conduct and was approved by the Western Institutional Review Board (study protocol number AG8618).

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Author Contributions

All authors were involved in 1) conception or design, or analysis and interpretation of data; 2) drafting and revising the article; 3) providing intellectual content of critical importance to the work described; and 4) final approval of the version to be published, and therefore meet the criteria for authorship in accordance with the International Committee of Medical Journal Editors (ICMJE) guidelines (accessible at: http://www.icmje.org/). In addition, all named authors take responsibility for the integrity of the work as a whole, and have given their approval for this version to be published.

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Disclosure

Dr Fitzgerald is an employee of Lundbeck LLC. J. Shepherd, H. Bailey, M. Berry, and J. Wright are employees of Adelphi Real World. Dr Chen is an employee of Otsuka Pharmaceutical & Commercial Development, Inc. Abilify Maintena® (aripiprazole once-monthly 400 mg), a longacting injectable antipsychotic indicated in the US for the treatment of schizophrenia and maintenance monotherapy of bipolar I disorder in adult patients, is distributed by Otsuka America Pharmaceutical, Inc., Rockville, MD, USA and comarketed by Otsuka and Lundbeck, Deerfield, IL, USA. The authors report no other conflicts of interest in this work.

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