Long-Acting Injectable Antipsychotic Adherence Among Texas Medicaid Enrollees with Schizophrenia

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Background & Objective

- It is estimated that 1.6% of the US population was diagnosed with schizophrenia in 2010.¹
- Total schizophrenia costs more than doubled between 2013 (~\$150 billion) and 2019 (\$340 billion).²
- Schizophrenia is associated with increased mortality compared to the general population primarily due to suicide and comorbidities such as cardiovascular diseases.³
- Long-acting injectable (LAI) antipsychotics (APs) have advantages over oral APs, including higher adherence rates and lower healthcare resource utilization.⁴
- However, comparisons among individual LAI second generation antipsychotics (SGAs) have resulted in inconclusive findings since many studies did not adjust for confounders.⁵⁻⁷
- Study Objectives: (1) Compare adherence among patients with schizophrenia initiating SGA LAIs and (2) Determine factors associated with adherence after controlling for covariates.

Methods

- Study Design: Retrospective cohort study
- **Database:** Texas Medicaid claims data (eligibility, pharmacy, medical)
- Study Population: Continuously enrolled adults (18-63 years) diagnosed with schizophrenia and with ≥ 1 SGA LAI prescription claim
- Index date: Date of the first SGA LAI prescription claim
- Outcome: Adherence Proportion of days covered (PDC):

number of days with medication available PDC = ---follow–up period (365 days)

Proportion Adherent: PDC \geq 0.80 \rightarrow Adherent

Mean PDC

| Study Period: | 6 pre-ir | -month 12-month ndex period post-index pe | ו eriod | | |
|-----------------------------------|----------------------------------|--|------------|---------------------|--------------------|
| | July 1, 2014 Jan 1, 2015 | Index Date | Jun 30, | , 2019 | Jun |
| | Earliest Pre-Index (6 months) | Identification Pariod |] | Latest Pos (12 m | st-Index onths) |
| | | Identification Penod | | | |

• **Data Analysis:** Inverse probability of treatment weighting (IPTW) was conducted to balance baseline characteristics among the six SGA LAI groups. Covariates included: clinical characteristics (comorbidities, psychotropic medications, baseline adherence, baseline healthcare utilization) and demographics (age, gender).

| Outcomes |
|--|
| Mean PDC |
| Proportion Adherent (PDC \geq 0.80) |

Multivariable analysis

Linear regression Logistic regression

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Results



Multivariable Logistic Regression of Factors Associated with Adherence (PDC≥0.80)

| | Odds Ratio | 95% CI | | P-value |
|--|------------|--------|-------|---------|
| SGA LAI (Reference: PP1M) | | | | |
| Aripiprazole lauroxil once monthly | 0.69 | 0.52 | 0.90 | 0.0064 |
| Aripiprazole lauroxil 2-month (1064 mg) | 1.93 | 1.14 | 3.29 | 0.0153 |
| Aripiprazole monohydrate | 1.15 | 0.97 | 1.37 | 0.0990 |
| Paliperidone palmitate 3-month | 11.30 | 6.59 | 19.39 | <.0001 |
| Risperidone 4-week | 1.11 | 0.90 | 1.36 | 0.3254 |
| Comorbid Psychiatric Condition at Baseline | | | | |
| Bipolar disorder | 0.97 | 0.83 | 1.13 | 0.7006 |
| Depression | 0.90 | 0.77 | 1.05 | 0.1796 |
| Anxiety | 1.07 | 0.92 | 1.25 | 0.3894 |
| Substance use disorder | 0.55 | 0.41 | 0.73 | <.0001 |
| Other mental health disorders | 0.89 | 0.66 | 1.19 | 0.4368 |
| Comorbid Psychiatric Condition in Follow-up Period | | | | |
| Bipolar disorder | 0.75 | 0.65 | 0.87 | 0.0002 |
| Depression | 0.69 | 0.59 | 0.80 | <.0001 |
| Anxiety | 0.78 | 0.67 | 0.91 | 0.0017 |
| Substance use disorder | 0.55 | 0.43 | 0.72 | <.0001 |
| Other mental health disorders | 1.06 | 0.65 | 1.74 | 0.8119 |
| Psychotropic Medications at Baseline | | | | |
| Anxiolytics | 1.12 | 0.95 | 1.31 | 0.1777 |
| Antidepressants | 1.16 | 0.98 | 1.37 | 0.0845 |
| Hypnotics/sedatives/sleep disorder agents | 0.996 | 0.81 | 1.23 | 0.9722 |
| Psychotropic Medications in the Follow-up Period | | | | |
| Anxiolytics | 0.90 | 0.77 | 1.06 | 0.2119 |
| Antidepressants | 0.85 | 0.71 | 1.01 | 0.0688 |
| Hypnotics/sedatives/sleep disorder agents | 0.87 | 0.71 | 1.06 | 0.1541 |
| Baseline Adherence and Comorbidity | | | | |
| Adherence to oral antipsychotics (PDC) | 1.02 | 1.01 | 1.02 | <.0001 |
| CCI | 0.99 | 0.93 | 1.06 | 0.3179 |
| Baseline Hospitalizations and Outpatient Visits | | | | |
| Schizophrenia-related hospitalizations | 1.44 | 1.00 | 1.96 | 0.0509 |
| All-cause outpatient visits | 0.99 | 0.96 | 1.02 | 0.3844 |
| Demographics | | | | |
| Age | 0.999 | 0.994 | 1.005 | 0.7850 |
| Male | 1.09 | 0.96 | 1.25 | 0.1982 |

| ard Error | P-value |
|-----------|---------|
| | |
| .97 | 0.0121 |
| .93 | 0.2196 |
| .28 | 0.1511 |
| .80 | <.0001 |
| .55 | 0.2896 |
| | |
| .15 | 0.1702 |
| .15 | 0.1947 |
| .15 | 0.7233 |
| .86 | <.0001 |
| .25 | 0.4602 |
| | |
| .13 | 0.0007 |
| .12 | <.0001 |
| .14 | 0.0015 |
| .72 | <.0001 |
| .78 | 0.0740 |
| | |
| .20 | 0.0546 |
| .25 | 0.1380 |
| .55 | 0.6905 |
| | |
| .18 | 0.0264 |
| .32 | 0.3925 |
| .45 | 0.2017 |
| | |
| 0.03 | <.0001 |
| .49 | 0.6213 |
| | |
| .49 | 0.0691 |
| .21 | 0.2752 |
| | |
| 0.04 | 0.2387 |
| .00 | 0.2047 |
| | |

Conclusions & Discussion

- This large observational real-world study showed that 3-month administration frequency was superior regarding SGA LAI adherence.
- Compared to PP1M, *aripiprazole lauroxil* once monthly was associated with suboptimal adherence regarding the expected mean PDC and the likelihood of being adherent.
- Aripiprazole lauroxil 2-month was associated with higher odds of being *adherent*, but not with mean PDC.
- *No difference* was observed between PP1M and risperidone 4-week or PP1M and aripiprazole monohydrate.
- Unique contribution: There is limited evidence comparing SGA LAIs; previous studies primarily focused on comparing PP1M with risperidone, as well as PP1M with PP3M.
- This is the first multiple group comparison among six different SGA LAIs.
- This comparative research will provide evidence for payers, policymakers, and practitioners when making decisions about treatment options.
- Limitations: (1) Diagnosis in medical claims may not represent the purpose for SGA LAI prescriptions; (2) Hospitalizations may have resulted in underestimating adherence; (3) PP3M patients transitioned from PP1M, so patients were exhibiting prior high adherence behaviors; (4) Results may not be generalizable outside of Texas Medicaid.
- Future studies could explore safety profile comparisons among SGA LAIs, and how patient experience influences treatment decisions and adherence in real-world settings.

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