

## Supplementary Material

**Article Title:** Beneficial Effects of Concomitant Long-Acting Injectable Antipsychotics on Time to Rehospitalization in Patients With Treatment-Resistant Schizophrenia Receiving Clozapine: A Retrospective Cohort Study

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### **DISCLAIMER**

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Supplementary Table 1 Number of patients using each LAI.

CLO+LAI(n=55)	
<b>Clopenthixol</b>	5
<b>Flupentixol</b>	3
<b>Fluphenazine</b>	28
<b>Haloperidol</b>	16
<b>Risperidone</b>	3

Supplementary Table 2 Number of patients using different oral antipsychotic combinations, among the CLO+OAP and CLO+LAI groups

	CLO+OAP (n=156)	CLO+LAI (n=55)
<b>None</b>	0	46
<b>Amisulpride</b>	5	0
<b>Aripiprazole</b>	4	0
<b>Chlorpromazine</b>	6	0
<b>Chlorpromazine+Risperidone</b>	1	0
<b>Clotiapine</b>	3	0
<b>Clotiapine+Olanzapine</b>	1	0
<b>Flupentixol</b>	3	0
<b>Haloperidol</b>	32	3
<b>Loxapine</b>	1	0
<b>Olanzapine</b>	8	0
<b>Olanzapine+Risperidone</b>	1	0
<b>Paliperidone</b>	1	0
<b>Risperidone</b>	12	2
<b>Sulpiride</b>	70	3
<b>Sulpiride+Olanzapine</b>	1	0
<b>Sulpiride+Risperidone</b>	1	0
<b>Trifluoperazine</b>	2	0
<b>Ziprasidone</b>	1	0
<b>Zotepine</b>	2	1
<b>Zuclopenthixol</b>	1	0

Supplementary Table 3 Cox regression of hazard ratio for rehospitalization in a year  
(Full model)

	Full model		Final model	
	HR (CI)	p	HR (CI)	p
<b>CLO vs. CLO+LAI</b>	2.53 (1.57-4.07)	<b>&lt;.001</b>	2.50 (1.56-4.01)	<b>&lt;.001</b>
<b>CLO+OAP vs. CLO+LAI</b>	2.66 (1.60-4.42)	<b>&lt;.001</b>	2.69 (1.62-4.46)	<b>&lt;.001</b>
<b>CLO+OAP vs. CLO</b>	1.05 (0.82-1.36)	.697	1.08 (0.84-1.38)	.569
<b>Age</b>	1.02 (1.01-1.03)	<b>&lt;.001</b>	1.02 (1.01-1.03)	<b>&lt;.001</b>
<b>Clozapine daily dosage</b>	1.00 (1.00-1.00)	<b>.025</b>	1.00 (1.00-1.00)	<b>.002</b>
<b>Previous admissions</b>	1.10 (1.08-1.12)	<b>&lt;.001</b>	1.10 (1.08-1.12)	<b>&lt;.001</b>
<b>Sex</b>	1.24 (1.01-1.53)	<b>.044</b>	1.23 (0.99-1.51)	.051
<b>Involuntary admission</b>	0.91 (0.45-1.84)	.795	-	-
<b>Lithium</b>	1.22 (0.77-1.93)	.390	-	-
<b>Valproate</b>	1.08 (0.87-1.34)	.473	-	-
<b>Carbamazepine</b>	1.65(0.23-12.03)	.624	-	-
<b>Lamotrigine</b>	0.92 (0.12-6.75)	.931	-	-
<b>Antidepressant</b>	1.09 (0.80-1.47)	.601	-	-
<b>Benzodiazepine</b>	0.92 (0.75-1.15)	.470	-	-
<b>Anticholinergics</b>	1.14 (0.92-1.43)	.238	-	-
<b>Laxatives</b>	1.11 (0.90-1.37)	.341	-	-

HR (CI): hazard ratio (95% confidence interval).

Full model: prior to backward stepwise elimination.

Final model: after backward stepwise elimination.

Bold p-values indicate a statistically significant difference.

Continuous variables: age, clozapine daily dosage, number of previous admissions

Categorical variables: sex (reference set at female), compulsory hospitalization; and concomitant use of mood stabilizers (lithium, valproate, carbamazepine, or lamotrigine), antidepressants, benzodiazepines, anticholinergics, and laxatives.

Supplementary Table 4 Cox regression of hazard ratio for rehospitalization in a year of the first hospitalization (Full model)

Full model		
	HR (CI)	p
<b>CLO vs. CLO+LAI</b>	2.39 (1.57-3.64)	<b>&lt;.001</b>
<b>CLO+OAP vs. CLO+LAI</b>	2.51 (1.59-3.98)	<b>&lt;.001</b>
<b>Age</b>	1.01 (1.01-1.02)	.092
<b>Clozapine daily dosage</b>	1.00 (1.00-1.00)	<b>.001</b>
<b>Previous admissions</b>	1.11 (1.09-1.13)	<b>&lt;.001</b>
<b>Sex</b>	1.27 (1.05-1.53)	<b>.014</b>
<b>Involuntary admission</b>	0.74 (0.18-3.00)	.670
<b>Lithium</b>	1.34 (0.92-1.94)	.127
<b>Valproate</b>	1.21 (0.99-1.48)	.058
<b>Carbamazepine</b>	2.59(0.15-20.37)	.989
<b>Lamotrigine</b>	0.98 (0.31-3.09)	.973
<b>Antidepressant</b>	1.29 (0.99-1.67)	<b>.046</b>
<b>Benzodiazepine</b>	0.92 (0.76-1.11)	.394
<b>Anticholinergics</b>	1.07 (0.88-1.30)	.515
<b>Laxatives</b>	1.05 (0.87-1.28)	.592

Supplementary Table 5 Hazard ratio for rehospitalization in each selected subgroup

	CLO (n = 508)		CLO+OAP (n = 156)		CLO+LAI (n = 55)	
	N	HR (95% CI)	N	HR (95% CI)	N	HR (95% CI)
	508	<b>2.50 (1.56-4.01)</b>	156	<b>2.69 (1.62-4.46)</b>	55	1
<b>Grouped by daily clozapine dosage</b>						
≤200 mg	168	<b>2.98 (1.41-6.30)</b>	43	<b>2.86 (1.21-6.74)</b>	24	1
201-399 mg	198	<b>2.59 (1.00-6.72)</b>	34	<b>2.98 (1.07-8.28)</b>	14	1
≥400 mg	142	<b>2.94 (1.23-7.06)</b>	79	<b>3.04 (1.27-7.31)</b>	17	1
<b>Grouped by numbers of previous admission</b>						
0-2	341	<b>4.77 (1.18-19.28)</b>	98	<b>4.32 (1.04-17.95)</b>	17	1
3-5	108	<b>2.20 (1.10-4.41)</b>	37	<b>3.33 (1.57-7.06)</b>	19	1
≥6	59	<b>2.27 (1.06-4.87)</b>	21	2.25 (0.95-5.33)	19	1

The reference was set at the CLO + LAI group.

Bold values indicate a statistically significant difference.

Adjusted covariates: age; sex; previous hospitalizations (in daily clozapine dosage subgroups); daily clozapine dosage (in number of previous admission subgroups); compulsory hospitalization; and concomitant use of mood stabilizers (lithium, valproate, carbamazepine, or lamotrigine), antidepressants, benzodiazepines, anticholinergics, and laxatives.