

Canadian Consortium for Early Intervention in Psychosis

# Pharmacotherapy Approach in First Episode Psychosis



#### Howard C. Margolese, MDCM, MSc, FRCPC

Associate Professor, Dept. of Psychiatry, McGill University Director, PEPP-MUHC, First Episode Psychosis Program VP Research, CCEIP *Montreal, QC* 



- Research Support: Acadia, Amgen, SyneuRX International
- Paid Speaker: AbbVie, HLS Therapeutics, Janssen, Otsuka, Lundbeck, Sunovion
- Consultant: Abbvie, HLS Therapeutics, Janssen, Otsuka, Lundbeck, Teva



# **Objectives**

- Discuss medication treatment approach in first episode psychosis
- Highlight clinical order sets to help with this approach
- Explore patient decision aids to encourage patients to accept proposed treatments



# First Episode Psychosis Treatment Considerations

# **Goals of Pharmacotherapy in FEP**

- Reduce DUP
- Achieve and maintain remission
- Improve quality of life
- Prevent progression of the disease



- 1. Birchwood M. Aust N Z J Psychiatry 2000; 34:S181-4
- 2. Hogarty GE, Ulrich RF. J Psychiatr Res 1998; 32:243-50
- 3. Robinson D, et al. Arch Gen Psychiatry 1999; 56:241-7.

# Pharmacotherapy Discussion Points with FEP Patients

- Often useful to include the family in this discussion
- Discuss all treatment options with patients including LAIs and clozapine
- Most patients will refuse LAIs, but I discussed early, not seen as a punishment and can be discussed again later
- When discussing PO options, focus on side effects, tolerability, etc.
- Discuss adherence
- Discuss use of medications with ETOH and Cannabis



# **Pharmacotherapy Approach**

- Use metabolically favorable second-generation antipsychotic (SGA) medications first
- Offer LAI early in the course of treatment discussed at very first or second visit.
- If complete or partial non-adherence is suspected, LAI re-discussed
- If 2 failed adequate trials we strongly suggest clozapine
- We prefer a trial of an LAI before clozapine to differentiate between lack of efficacy and partial/non-adherence



# Pharmacotherapy: Safety First Protocol

- 1. Metabolically favourable SGA
  - Aripiprazole, Brexpiprazole, Cariprazine, Lurasidone, Asenapine (rarely used), Ziprasidone (rarely used);
  - Or Arip, Risp or Palip if LAI is being considered
  - Or straight to SGA-LAI Aripiprazole or Paliperidone
- 2. Choose
  - Different MF SGA from above if intolerability, lack of efficacy
  - If Partial/non-adherence suspected choose SGA-LAI
  - If severe EPS, consider quetiapine
- 3. SGA LAI or Clozapine
- 4. Clozapine



# Pharmacotherapy: Safety First Protocol

- Whatever medication is chosen first will usually work in 70-75% of patients for psychosis (1)
- Therefore, tolerability drives choice
- SGA LAIs are preferred when accepted given high rates of non-adherence
- FGA are not used given negative effects on brain volumes (2,3)



# Pharmacotherapy: Safety First Protocol

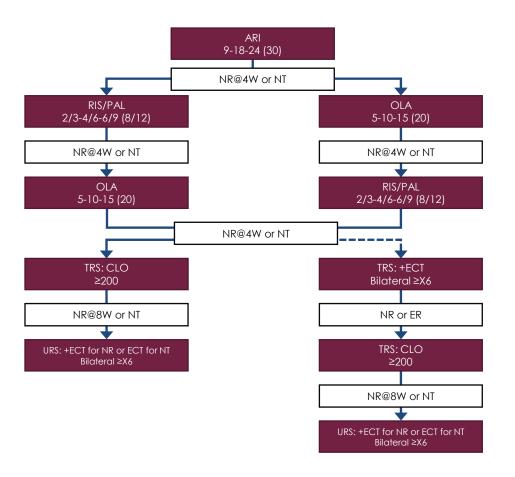
- Whatever medication is chosen, Optimize dose based on efficacy and tolerability
- Goal is to treat to symptom remission
- We do not wait to increase dose; if limited or partial response after 2 weeks, we increase the dose (1)



## Some Experts Recommend 3<sup>rd</sup>-Generation Antipsychotics as First-line Treatment for Antipsychotic-naïve Patients

Japanese Society of Clinical Neuropsychopharmacology Treatment Algorithm

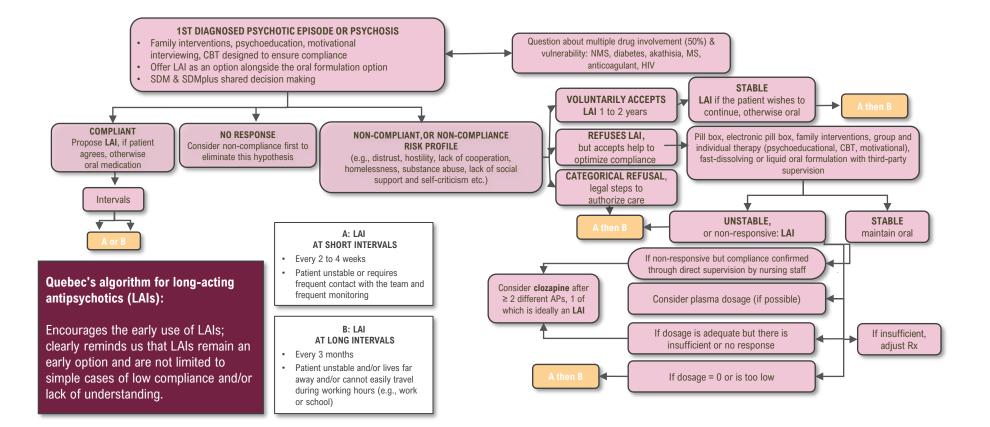
- 1<sup>st</sup>-line: Aripiprazole
- 2<sup>nd</sup>-, 3<sup>rd</sup>-line: Risperidone / paliperidone or olanzapine
- 4<sup>th</sup>-line: Clozapine



ARI, aripiprazole; BE-PSD, Brief Evaluation of Psychosis Symptom Domains; CLO, clozapine; ECT, modified electrocon vulsive therapy; EPS, extrapyramidal symptoms; LAI, long-acting injection; OLA, olanzapine; PAL, paliperidone; QUE, quetiapine; RIS, risperidone; TRS, treatment-resistant schizophrenia; URS, ultra-resistant schizophrenia

1. Huhn M, et al. Lancet 2019; 394:939-51.

### **QAAPAPLE Algorithm 2019**



1. Stip, Emmanuel, et al. "Antipsychotiques à Action Prolongée: Révision de l'algorithme QAAPAPLE." The Canadian Journal of Psychiatry. 2019; 0706743719847193.

#### **Order Sets**

https://www.epicanada.org/clinicalordersets

### What is a Clinical Order Set?

- A pre-defined template that provides support in making clinical decisions for a specific condition or medical procedure
- A grouping of orders that standardizes and expedites the ordering process for a common clinical scenario
- Clinical order sets guide clinicians while treating patients to ensure that they do not miss any critical components of care
- Order sets can be used to incorporate the latest evidence-based best practice to clinical workflow

# **Advantages of Clinical Order Sets**

- Aligned to provincial mandates (HQO Schizophrenia Care Standard)
- Reduces variability
- Integrates best practices (standardized approach)
- Identifies practice patterns
- Facilitates outcome evaluations
- Can be customized based on existing policies and procedures and to reflect variances in practice
- Can be integrated into Electronic Medical Records

### **CCEIP Order Sets**

In partnership with Think Research, CCEIP has developed four clinical order sets:

- **1. Initiation of Treatment for Early Phase Psychotic Disorders**
- 2. Optimization of Treatment for Early Phase Psychotic Disorders
- 3. Clozapine initiation
- 4. Cannabis and early psychosis

https://www.epicanada.org/clinicalordersets

# **Order Set Themes**

#### Administration

- Document Purpose
- Working Diagnosis

#### Substance Use Screening

- Alcohol Use Disorders Identification Test (AUDIT)
- Drug Abuse Screening Test, DAST-10
- Additional Information

#### Psychiatric Symptoms Assessment Tools

- Clinical Global Impression-Severity (CGI-S) Scale
- Brief Psychiatric Rating Scale (BPRS) 4-Item Positive Symptom Rating Scale

# **Order Set Themes (con't)**

#### Physical Assessment

- Movement Disorder Assessment Tools
  - Tools for Monitoring Antipsychotic Side Effects (TMAS)
  - Abnormal Involuntary Movement Scale (AIMS)
  - Extrapyramidal Symptom Rating Scale (ESRS)
- Vitals/Monitoring
- Lab Investigations (if not previously obtained)
- Diagnostics
- Allergies and Medication Review
- Antipsychotic Treatment Capacity Assessment

# **Order Set Themes (con't)**

#### Management of Psychosis

- It is recommended that preference be given to atypical antipsychotics in the treatment of early psychosis patients
- It is recommended that LAI (Long-Acting Injectable) antipsychotic therapy is offered during all phases of psychotic disorders, including the early phase
- To address high rates of partial/non-adherence in early psychosis patients, preference is given to medications available in a long-acting formulation
- Refer to Antipsychotic Treatment Selection Tool
- Atypical Antipsychotics
  - Oral Medication with LAI Formulations
  - LAI Antipsychotic Medication

#### OR

- Alternate Atypical Antipsychotic Medication
- Patient choice; refer to OPTIMA: Offering Patients Therapeutic Information on Medication Alternatives

# **Order Set Themes (con't)**

#### Adjunctive Management

- Anticholinergic Agents
- Benzodiazepines
- Other
- Cognitive Behavioural Therapy
- Smoking Cessation
- Psychoeducation and Health Lifestyle Information
  - Provide psychoeducation to patient, refer to iHope tool
- Referrals
- Additional Orders

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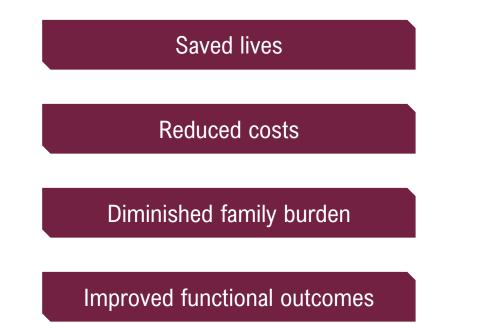
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Movement Disorder Assess:           □ tools for Monitoring Antipsych available at: http://epicanada.y           △ Abnormal Involuntary Moveme disorders/clinical-scales-moveme           □ Extrapyramidal Symptom Radius Extrapyramidal Symptom Radius (Taits/Monitoring)           □ Weigh patient, measure heigh Usits circumferences?           □ T°, HR°, RR°, BP° as per polici           □ CBC°           Chemistry           □ CBC°           Chemistry           □ Urine β HGG           □ Vinical Lab Investigations           □ Fasting glucose5           □ Prolactin	hotic Side Effects (TMAS) pro/project/bod-for-monitorin and Scale (AlMS) available a mt-disorders/aims-abnormal ng Scale (ESRS)  at: Weight <sup>6</sup> : kg, H cm cm cm eviously obtained)  Days Creatinine <sup>6</sup> Urine drug scre A1C5	t http://www.psychiatrictimes.com/clinical-scales-movem involuntary-movement-scale leight:m  BMI <sup>6</sup> :kg/m <sup>2</sup>	ent- ent- ent-

	ies on approved f s been reviewed a						
	nitiation of T	reatment i	or Early F	Phase Psych	notic Disorde	rs Order Set	АСТК
Physical Ass	essment Con	tinued					
Diagnostics							
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MRI Recei							
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	d Medication I lergen and react						
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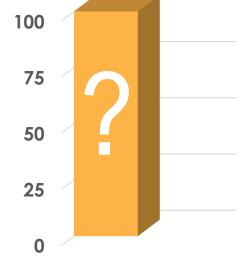
Document allergies of reconciliation has be						
Initi	ation of Treat	ment for Ear	rly Phase Psychoti	c Disorders	s Order Set	АСП
Management of	Psychosis					
***It is recommend	led that preference	-Acting Injectab		is offered durir	arly psychosis patients***9 ng all phases of psychotic	
		adherence in e	arly psychosis patients, p acting formulation*** <sup>0</sup>	vreference is giv	ven to medications available	
		tion Tool availab	ble at: https://vivomap.ca	lib/surveyStand	lalone/psychosis.php	
Atypical Antipsyc						
Oral Medication wit						
			(frequency) (frequency)			
□ palipendone	mg PO		(frequency)	[caution	-genamo,renalj v genamo,renalj	
LAI Antipsychotic M			(	[vaulioi	. Server et reherret e roll	
		nsvebotic should	t be established prior to i	nitiating treatm	ent with LAI formulation***1	>
			(frequency)			5
					ate) [caution-geriatric,renal]	C
Trisperidone micro:		ma IM	(frequency)			ţ
risperidone micro: [caution-geriatric,he ] OR	spheres	mg IM	(frequency)			e Document
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reconciliation has been	reviewed as per organizatio	mai process			
Initiat	ion of Treatment for	Early Phase P	sychotic Disorder	s Order Set	ACT
	ement Continued				
Other Anticonvulsants:					_
					_
Other:					_
Cognitive Behavio	oural Therapy e for cognitive behavioural t	horopy (CBT) <sup>13</sup> for a	asvehocie?		
Yes I			usychosis r		
If patient appropriate	e for CBT and CBT not offer	ed, please provide e	explanation (e.g. patient	refusal):	_
**	*Prescriber to initiate referra	al for CBT in the Ref	ferrals section, as approp	priate***	-
Smoking Cessatio	m				
	treatment combined with co		ffective than pharmacolo	gical treatment alone***12	1
Is patient a smoker?					
<ul> <li>Is patient a smoker?</li> <li>If patient is a smoke (e.g. patient refusal)</li> </ul>	Yes Not r and not offered smoking c	essation education/i	interventions, <sup>2,10</sup> please	provide explanation	
<ul> <li>Is patient a smoker?</li> <li>If patient is a smoke (e.g. patient refusal)</li> </ul>	Yes No r and not offered smoking c : o initiate referral to Smoking	essation education/i	interventions, <sup>2,10</sup> please	provide explanation	
Is patient a smoker? If patient is a smoke (e.g. patient refusal) ***Prescriber to Phamacological Ma nicotine patch	Yes No r and not offered smoking c : o initiate referral to Smoking	essation education/i	interventions, <sup>2,10</sup> please	provide explanation	
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Is patient a smoker? If patient is a smoke (e.g. patient refusal) ""Prescribert t Pharmacological Mi nicotine patch Psychoeducation	Yes Nit     rand not offered smoking c     i     i     initiate referral to Smoking     anagement     mg Topically daily for     and Health Lifestyle I	Cessation Education/ Cessation Counsel	interventions, <sup>2,10</sup> please lior in the Referrals secti- then notify MD/NP to rea	provide explanation on, as appropriate*** ssess (14 – 21 mg)	
Is patient a smoker?     If patient is a smoker     (e.g. patient refusal)     ""Prescribert (     Phamacological Ma     Inicotine patch     Provide education to     Diagnosis and co	Yes Nit     r and not offered smoking c     i     initiate referral to Smoking     anagement     mg Topically daily for     anad Health Lifestyle     o patient on the following top     uruse of illness/prognosi/re	Cessation education// Cessation Counsel	interventions, <sup>2,16</sup> please llor in the Referrals secti then notify MD/NP to rea ng, and electronically, as	provide explanation on, as appropriate*** ssess (14 – 21 mg)	
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Is patient a smoker?  If patient is a smoker?  If patient is a smoker?  Pharmacological M:  nicotine patch  Provide education tt  Diagnosis and cc  Treatment option  Risk of substa  Risk of substa  Importance of ad  **Prescriber  Offer family interven  Provide education tt  Comparison  Provide education terven	Yes Net and not offered smoking of initiate referral to Smoking anagement and Health Lifestyle I patient on the following tog iniciduality daily daily for and Health Lifestyle I patient on the following tog nurse of illness/prognosis/re is, including their potential e and options, including clozep nd recognition of warning si ad monitoring for warning	Cessation education// Cessation Counsel Cessation Counsel Information Covery Weeks, I Information Covery Information Grass and relapse pre grass and rela	Interventions, <sup>2,10</sup> please lifer in the Referrals section then notify MD/NP to react and electronically, as exists and electronically as exists and electronically as exists and electronically as and electronically as a sector and supports <sup>2,10</sup> (specify)	provide explanation on, as appropriate*** seess (14 – 21 mg) : applicable <sup>2</sup> : ions as well as illness nt strategies wws/ihope-tool/_***	
Is patient a smoker?  If patient is a smoker?  If patient is a smoker?  Pharmacological M:  nicotine patch  Provide education tt  Diagnosis and cc  Treatment option  Risk of substa  Risk of substa  Importance of ad  **Prescriber  Offer family interven  Provide education tt  Comparison  Provide education terven	Yes Net ar and not offered smoking of initiate referral to Smoking anagement mg Topically daily for and Health Lifestyle patient on the following top purse of illness/prognosis/re s, including their potential e and recognition of warning si ind monitoring for warning s	Cessation education// Cessation Counsel Cessation Counsel Information Covery Weeks, I Information Covery Information Grass and relapse pre grass and rela	Interventions, <sup>2,10</sup> please lifer in the Referrals section then notify MD/NP to react and electronically, as exists and electronically as exists and electronically as exists and electronically as and electronically as a sector and supports <sup>2,10</sup> (specify)	provide explanation on, as appropriate*** seess (14 – 21 mg) : applicable <sup>2</sup> : ions as well as illness nt strategies wws/ihope-tool/_***	
Is patient a smoker?     If patient is a smoker?     If patient is a smoker?     If patient refusal?     ""Prescribert?     Plancacological Mic     Inicotine patch     Provide education to     Diagnosis and co     Treatment option     Alternate treatme     Risk of relapse a     Risk of relapse a     Importance of ad     """Prescribert?     Offer family intervene     Provide education of	Yes Net r and not offered smoking c i initiate referral to Smoking anagement mg Topically daily for and Health Lifestyle I patient on the following top purse of illness/prognosi/re is, including their potential e and options, including clozap nd recognition of warning si noce use (patricularly cannat herence with treatment and pro consider use of the iHd ton to provide family-focus in healthy eating, physical ar family with contact informati Prescriber to initiate relevan	Cessation education// Cessation Counsel Cessation Counsel Information Covery Weeks, I Information Covery Information Grass and relapse pre grass and rela	Interventions, <sup>2,10</sup> please Ilor in the Referrals secti- then notify MD/NP to rea- ng, and electronically, as exis exis exis exis solution strategies ctions with treatment opt s adherence enhanceme inttp://epicanada.org/ne and support <sup>2,10</sup> upports <sup>2</sup> : (specify) eferrals section, as appr	provide explanation on, as appropriate*** seess (14 – 21 mg) : applicable <sup>2</sup> : ions as well as illness nt strategies wws/ihope-tool/_***	
Is patient a smoker?     If patient is a smoker?     If patient is a smoker?     If patient refusably     ""Prescriber for     Provide education to     Tratment option     Alternate treatment     Risk of relapse a     Risk of suicida a     Importance of ad     """Prescriber     Offer family intervene     Provide education o     Provide education o     "rotide education o     "Provide education o     "Provide education o     "Provide education o     "Provide education o     "rotide education o     "Provide education o     "rotide education o     "Provide education o     "Provide education o     "rotide education o	Yes Net ar and not offered smoking of initiate referral to Smoking anagement mg Topically daily for and Health Lifestyle patient on the following top purse of illness/prognosis/re s, including their potential e and recognition of warning si ind monitoring for warning s	Cessation education// Cessation Counsel Cessation Counsel Information Covery Weeks, I Information Covery Information Grass and relapse pre grass and rela	Interventions, <sup>2,10</sup> please lifer in the Referrals section then notify MD/NP to react and electronically, as exists and electronically as exists and electronically as exists and electronically as and electronically as a sector and supports <sup>2,10</sup> (specify)	provide explanation on, as appropriate*** seess (14 – 21 mg) : applicable <sup>2</sup> : ions as well as illness nt strategies wws/ihope-tool/ ***	

# If You Had A Treatment Strategy That:









Dr. John Kane, personal communication, with permission

#### **OPTIMA and OPTIC Shared Decision-Making Tools**

https://www.epicanada.org/tools-shareddecisionmaking

#### **OPTIMA: A Tool For Patient Engagement**



#### SHOULD YOU TAKE MONTHLY MEDICATION FOR YOUR SYMPTOMS?

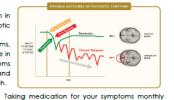
A consult decision aid for people with psychotic symptoms and healthcare professionals to discuss options.

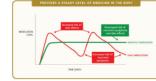
#### Why are you being offered monthly medication for your symptoms?

Your symptoms affect your ability to function well at home, school or work, and in your social life. Medication is a vital part of freating your symptoms and restoring your ability to take part in everyday activities. The main goal of taking medication is to help you stay well so you can stay out of hospital. Starting medication when symptoms first happen is the best way to do this.

#### Uninterrupted medication helps you stay well.<sup>1</sup>

- Being well is known as remission. This is shown in the graph titled "Possible Outcomes of Psychotic Symptoms".
- If you do not take medication for your symptoms, you cannot function well. You may need to be in hospital often, as you will have symptoms (relapses). Having relapses affects your brain and prevents full recovery. This is shown in the graph.





provides steady levels of medication in the body, as shown in the graph titled "Monthly Medication Compared with Daily Medication Provides a Steady Level of Medication in the Body".<sup>2</sup>

- This way of giving medication may decrease side effects, such as sleepiness and weight gain, compared with daily oral medication.<sup>3</sup> Daily medication is shown in the graph.
- Monthly medication may also increase effectiveness, as it decreases the risk of recurrent symptoms, compared with oral medication.
- · With monthly medication, you receive less medication than with daily medication.

#### OFFERING PATIENTS THERAPEUTIC INFORMATION MEDICATION ALTERNATIVES (OPTIMA)

Person's Name

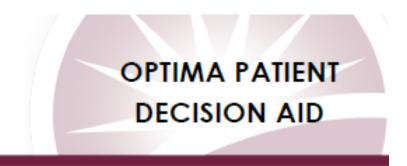
BENEFITS AND REASONS TO CHOOSE MONTHLY MI	EDICATION	YES	NO
When you take monthly medication rather than daily	Is it important to you to get along better with		
medication, you may get along at home better, be able to	family and friends?	_	
go to school or work, have a better social life and be more	Is it important to you to be able to go to		
satisfied with treatment. <sup>1</sup>	school or work?		
	Is it important to you to have a better social		
People who switch to monthly medication say their	life?		
symptoms, ability to function in their daily life, and	Is it important to you to be satisfied with		
satisfaction with treatment are much better with monthly	your treatment?		
medication. <sup>2,3</sup>		_	
Over two years, more people stay on treatment with	Is it important to you to stay on medication?		
monthly medication than daily medication. <sup>4</sup>			
If you need to be hospitalized with symptoms, you may	Is it important to you to spend less time in		
spend much less time in hospital with monthly medication	hospital?		
than with daily medication, about two-thirds less time. <sup>5</sup>			
People who forget to take their medication daily are more	Is it important to you to avoid being		
likely to have symptoms and more likely to be hospitalized	hospitalized with symptoms?		
than people taking monthly medication-6			
Over a one-year period, people taking daily medication			
are three times more likely to be hospitalized with			
symptoms than people taking monthly medication. <sup>1</sup>			
Relapses make it harder for you to recover. Usually, fewer	Is it important to you to have better long term		
relapses happen with monthly than with daily medication.	treatment result and fewer relapses?	_	_
Long-term treatment results are better with monthly			
medication. <sup>7</sup>			
Some monthly medication begins to work fast <sup>®</sup> This means	Is it important to you to recover faster and		
you may be able to recover faster and leave hospital	leave hospital sooner?		
sooner.			
When you take monthly medication, you have more steady	Is it important to you to have fewer side		
and even levels of medication in the body and may have	effects?		
fewer side effects, such as sleepiness and weight gain,			
than with daily medication. <sup>9</sup>			
You receive less medication with monthly than with daily	Is it important to you to receive less		
medication.	medication?		
When you take medication monthly, you do not need to	Is it important to you not to worry about		
worry about remembering to take medication daily.	remembering to take medication?		
Taking monthly treatment may be simpler for you than	Is it important to you to simplify your		
taking daily.	medication plan?		
If you forget to go for your monthly medication, we will	Is it important to you to stay connected to your		
reschedule your appointment with your treatment team to	treatment team?		
maintain your medication's effectiveness.			
maintain your medication's ettectiveness. People taking monthly medication better protect their	Is it important to you to protect your brain		

1. Margolese HC, Steiner W, Lalla F, Cattan C, Perillo A, Arshoff L. Development of a patient decision aid to educate patients with acute psychoses about long-acting injectable antipsychotic therapy. Institute on Psychiatric Services (IPS) 2016: The Mental Health Services Conference, from October 6 to 9, 2016, Washington, DC.



#### https://www.epicanada.org/optima





# SHOULD YOU TAKE MONTHLY MEDICATION FOR YOUR SYMPTOMS?

A consult decision aid for people with psychotic symptoms and healthcare professionals to discuss options.

*What does the acronym OPTIMA stand for?* **O**ffering **P**atients **T**herapeutic **I**nformation on **M**edication **A**Iternatives

#### The OPTIMA Tool Has 2 Parts:

#### 1) Education



#### SHOULD YOU TAKE MONTHLY MEDICATION FOR YOUR

#### SYMPTOMS?

A consult decision aid for people with psychotic symptoms and healthcare professionals to discuss options.

#### Why are you being offered monthly medication for your symptoms?

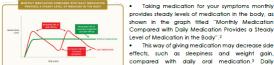
Your symptoms affect your ability to function well at home, school or work, and in your social life. Medication is a vital part of treating your symptoms and restoring your ability to take part in everyday activities. The main goal of taking medication is to help you stay well so you can stay out of hospital. Starting medication when symptoms first happen is the best way to do this.



 Being well is known as remission. This is shown in the graph titled "Possible Outcomes of Psychotic Symptoms". If you do not take medication for your symptoms. you cannot function well. You may need to be in hospital often, as you will have symptoms

(relapses). Having relapses affects your brain and prevents full recovery. This is shown in the graph.





shown in the graph titled "Monthly Medication Compared with Daily Medication Provides a Steady Level of Medication in the Body".2 This way of giving medication may decrease side effects, such as sleepiness and weight gain, compared with daily oral medication.<sup>3</sup> Daily medication is shown in the graph.

 Monthly medication may also increase effectiveness, as it decreases the risk of recurrent symptoms, compared with oral medication.

· With monthly medication, you receive less medication than with daily medication.

#### 2) Motivational Interview

#### OFFERING PATIENTS THERAPEUTIC INFORMATION MEDICATION ALTERNATIVES (OPTIMA)

Person's Name

BENEFITS AND REASONS TO CHOOSE MONTHLY ME	EDICATION	YES	NO
When you take monthly medication rather than daily	Is it important to you to get along better with		
medication, you may get along at home better, be able to	family and friends?	_	_
go to school or work, have a better social life and be more	Is it important to you to be able to go to		
satisfied with treatment. <sup>1</sup>	school or work?		
	Is it important to you to have a better social		
People who switch to monthly medication say their	life?	_	
symptoms, ability to function in their daily life, and	Is it important to you to be satisfied with		
satisfaction with treatment are much better with monthly	your treatment?		
medication. <sup>2,3</sup>		_	_
Over two years, more people stay on treatment with	Is it important to you to stay on medication?		
monthly medication than daily medication.4		_	_
If you need to be hospitalized with symptoms, you may	Is it important to you to spend less time in		
spend much less time in hospital with monthly medication	hospital?		
than with daily medication, about two-thirds less time. <sup>5</sup>			
People who forget to take their medication daily are more	Is it important to you to avoid being		
likely to have symptoms and more likely to be hospitalized	hospitalized with symptoms?		
than people taking monthly medication. <sup>6</sup>			
Over a one-year period, people taking daily medication are three times more likely to be hospitalized with			
symptoms than people taking monthly medication. <sup>1</sup>			
Relapses make it harder for you to recover. Usually, fewer	Is it important to you to have better long term		
relapses happen with monthly than with daily medication.	treatment result and fewer relapses?	•	-
Long-term treatment results are better with monthly	realment resolt and rewer relapses:		
medication. <sup>7</sup>			
Some monthly medication begins to work fast <sup>®</sup> This means	Is it important to you to recover faster and		
you may be able to recover faster and leave hospital	leave hospital sooner?	•	
sooner.	leave hospital society		
When you take monthly medication, you have more steady	Is it important to you to have fewer side		
and even levels of medication in the body and may have	effects2	-	-
fewer side effects, such as sleepiness and weight gain,	checks.		
than with daily medication.?			
You receive less medication with monthly than with daily	Is it important to you to receive less		
medication.	medication?	-	-
When you take medication monthly, you do not need to	Is it important to you not to worry about		
worry about remembering to take medication daily.	remembering to take medication?	-	_
Taking monthly treatment may be simpler for you than	Is it important to you to simplify your		
taking daily.	medication plan?	-	
If you forget to go for your monthly medication, we will	Is it important to you to stay connected to your		
reschedule your appointment with your treatment team to	treatment team?	_	_
maintain your medication's effectiveness.			
People taking monthly medication better protect their	Is it important to you to protect your brain		
brain from additional illness than people taking daily	from the illness?	-	_

1. Margolese HC, Steiner W, Lalla F, Cattan C, Perillo A, Arshoff L. Development of a patient decision aid to educate patients with acute psychoses about long-acting injectable antipsychotic therapy. Institute on Psychiatric Services (IPS) 2016: The Mental Health Services Conference, from October 6 to 9, 2016, Washington, DC.

### **Patient Handout**

#### **3 Important Sections**

- a) Should you...
- b) Why?
- c) Education on continuous treatment



# SHOULD YOU TAKE MONTHLY MEDICATION FOR YOUR SYMPTOMS?

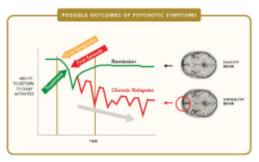
A consult decision aid for people with psychotic symptoms and healthcare professionals to discuss options.

#### Why are you being offered monthly medication for your symptoms?

Your symptoms affect your ability to function well at home, school or work, and in your social life. Medication is a vital part of treating your symptoms and restoring your ability to take part in everyday activities. The main goal of taking medication is to help you stay well so you can stay out of hospital. Starting medication when symptoms first happen is the best way to do this.

#### Uninterrupted medication helps you stay well.<sup>1</sup>

- Being well is known as remission. This is shown in the graph titled "Possible Outcomes of Psychotic Symptoms".
- If you do not take medication for your symptoms, you cannot function well. You may need to be in hospital often, as you will have symptoms (relapses). Having relapses affects your brain and prevents full recovery. This is shown in the graph.



### **OPTIMA Questionnaire**

#### Instructions

- Questionnaire lists benefits and risk (discomfort) of monthly medication and is the core of decision aid
- Series of questions about importance to patient of specific benefits of monthly medication
- Each question preceded by evidence statement
- Review each question to ensure patient understands
- Have patient respond **Yes or No** to each question
- Record response on decision aid

BENEFITS AND REASONS TO CHOOSE MONTHLY ME	DICATION	YES	NO
When you take monthly medication rather than daily medication, you may get along at home better, be able to	Is it important to you to get along better with family and friends?		
go to school or work, have a better social life and be more satisfied with treatment. <sup>1</sup>	Is it important to you to be able to go to school or work?		
People who switch to monthly medication say their	Is it important to you to have a better social life?		
symptoms, ability to function in their daily life, and satisfaction with treatment are much better with monthly medication. <sup>2,3</sup>	Is it important to you to be satisfied with your treatment?		
Over two years, more people stay on treatment with monthly medication than daily medication. <sup>4</sup>	Is it important to you to stay on medication?		
If you need to be hospitalized with symptoms, you may spend much less time in hospital with monthly medication than with daily medication, about two-thirds less time. <sup>5</sup>	Is it important to you to spend less time in hospital?		
People who forget to take their medication daily are more likely to have symptoms and more likely to be hospitalized than people taking monthly medication <sup>4</sup>	Is it important to you to avoid being hospitalized with symptoms?		
Over a one-year period, people taking daily medication are three times more likely to be hospitalized with symptoms than people taking monthly medication. <sup>1</sup>			
Relapses make it harder for you to recover. Usually, fewer relapses happen with monthly than with daily medication. Long-term treatment results are better with monthly medication. <sup>7</sup>	Is it important to you to have better long term treatment result and fewer relapses?		
Some monthly medication begins to work fast <sup>a</sup> This means you may be able to recover faster and leave hospital sooner.	Is it important to you to recover faster and leave hospital sooner?		
When you take monthly medication, you have more steady and even levels of medication in the body and may have fewer side effects, such as sleepiness and weight gain, than with daily medication. <sup>9</sup>	Is it important to you to have fewer side effects?		
You receive less medication with monthly than with daily medication.	Is it important to you to receive less medication?		
When you take medication monthly, you do not need to worry about remembering to take medication daily.	Is it important to you not to worry about remembering to take medication?		
Taking monthly treatment may be simpler for you than taking daily.	Is it important to you to simplify your medication plan?		
If you forget to go for your monthly medication, we will reschedule your appointment with your treatment team to maintain your medication's effectiveness.	Is it important to you to stay connected to your treatment team?		
People taking monthly medication better protect their brain from additional illness than people taking daily medication. <sup>10</sup>	Is it important to you to protect your brain from the illness?		

ENTS THERAPEUTIC INFORMATION MEDICATION ALTERNATIVES (OPTIMA

#### **OPTIC: Offering Patients Therapeutic Information about Clozapine, A Shared Decision-Making Tool**

by Lauren Said, Pharm.D and Howard C. Margolese, MD, CM, MSc, FRCPC

Poster presented at SIRS virtual

🐯 McGill

meeting on April 20th, 2021 Submission #3007067

#### Abstract

Patients suffering from treatment resistance schizophrenia (TRS) often have an inadequate clinical response and a less favorable functional outcome. Therefore, it is important to use the most effective treatment available. Although clozapine has proven benefits in treating TRS compared to other antipsychotics, it is only prescribed in less than 50% of eligible patients. Clozapine's list of side effects and blood monitoring requirements give it a negative reputation amongst patients and some physicians.

The primary focus of the OPTIC tool is to facilitate a balanced discussion between clinicians and patients that emphasizes the benefits of clozapine as the gold standard treatment for TRS. When clinicians are confident that clozapine is the best treatment for their patients, their genuine approach to offering clozapine will be better received. The OPTIC tool is composed of three parts, a clinician information handout, a patient handout and patient questionnaire.

The clinician handout discusses the benefits of clozapine in TRS, how to initiate a positive offer of clozapine to patients as well as how to best manage the side effects with lifestyle changes and medications. There is also a guide to initiating clozapine.



physicians on what information to discuss with patients when offering clozapine as a treatment option. Clozapine should be offered systematically to people with TRS as soon as they become eligible. This tool can facilitate a positive offer and increase acceptance of clozapine by your patient.

#### CLINICAL RECOMMENDATIONS ON THE USE OF CLOZAPINE

4

Clozapine is a second-generation antipsychotic medication that has proven to be more effective than other antipsycholics for patients with TRS, Clozapine has been shown to improve remission rates, and reduce psychotic symptoms, risk of relapse, length of hospitalization, suicidal risk, and mortality. Despite its effectiveness, it remains underused in Canada, and is often prescribed as a last resort.<sup>1</sup> Physicians and patients often have negative perceptions towards the medication due to the side effects and specific laboratory monitoring requirements. However, according to Canadian guidelines, clozapine should be offered as a 3rd line therapy, i.e., after two unsuccessful trials of adequate dose and duration using other antipsychotic medications.<sup>2</sup> This is also referred to as TRS. It is important to discuss clozapine's effectiveness in treating TRS with patients to allow them to make an informed decision about their treatment. Clozapine should be offered to both inpatients and outpatients who have poor treatment response or lingering symptoms due to partial response with their current treatment. Patients who are highly suicidal, who have substance use disorders and who present with polydipsia may also benefit from clozapine

· People with schizophrenia and a co-morbid substance abuse disorder (possibly) Contraindications: pre-existing myeloproliferative disorder, neutropenia, knows

 "We have taked about your goals and the importance of medication to manage your symptoms. We have tried different antipsychotics and have not succeeded in treating your symptoms. Now, I would like to talk to you about clozapine, which is an effective antipsychotic that is recommended for people like you who have not responded well to

Question 1: Why are you being offered clozapine for your symptoms? Review and help the patient understand the benefits of clozapine after two failed

antipsychotic treatments (remission). Every other trial after the second antipsychotic has a response rate of roughly 10-20%. except for clozapine which has a response rate of 60-77%?

- KEY POINTS TO REMEMBER ABOUT CLOZAPINE 1. SIDE EFFECTS
- When reviewing clozapine's side effects with patients, it is important to adequately describe

them so that patients can have an honest depiction of the medication. However, it should be mentioned that even though they are "common" side effects, not all patients will experience these symptoms. Furthermore, most of them can be adequately managed with a healthy lifestyle and/or pharmaceutical treatments. As for the severe side effects, given that clozapine is an older medication, a lot of information is available about its potential side effects and their associated risks are well documented. Also, many experts have recommended strategies to reduce the incidence of severe side effects. Therefore, they are now considered well-managed

#### 2. BENEFITS

Clozapine was shown to be more effective than other antipsychotics in many studies. In fact, one particular study showed that after 3 months of treatment with clozapine, patients showed significant improvement in symptoms, improved quality of life and greater improvement in their overall mental health compared to other antipsychotics.<sup>24</sup>

The two po parts were assist pati clozapine The cloza clearly high benefits o compare treatment discusses side effec managed

The clinici questionn motivatio concepts discussion the patier their treat determine suited to I end of the the patier make a d taking clo ready.

re developed to			OFFERING PATIENTS THERAPEUTIC INF	ORMATION ON CLOZAPINE (O	PTIC)	) ©
tients in deciding if	Canadian Consortium for Early Intervention	CLOZAPINE DECISION AID	Person's Name:		YES	NO
e is right for them.	in Psychosis	DECISION AID	When you take clozapine rather than other medication,	Is it important to you to get along better with family and friends?		
apine decision aid			you may get along at home better, be able to go to school or work, have a better social life and be more satisfied	Is it important to you to be able to go to school or work?		
ighlights the	SHOULD YOU TAKE CLOZAPINE FOR Y		with treatment. People who switch to clozapine say their symptoms, ability to function in their daily life, and	Is it important to you to have a better social life?		
of clozapine	A decision aid for people with psychotic symptoms and th antipsychotic medication options.	eir healthcare professionals to discuss	satisfaction with treatment is better. <sup>1</sup>	Is it important to you to be satisfied with your treatment?		
ed to other	Why are you being offered clozapine for your symp	otoms?	Over two years, more people stay on clozapine than other medication. <sup>8</sup>	Is it important to you to stay on medication?		
nts for TRS and	Your psycholic symptoms have not responded to the medications y your ability to function well at home, school or work, and in your soo your symptoms and restoring your ability to take part in everyday	cial life. Medication is a vital part of treating	People not optimally treated are more likely to have symptoms and more likely to be hospitalized than people taking clozapine. <sup>III</sup>	Is it important to you to avoid being hospitalized with symptoms?		
s how clozapine's cts can be	medication for people who have not responded well to two differe goal of taking medication is to help you stay well so that you can fu course, stay out of hospital. Taking medication that can adequately to best way to accomplish this.	nction well, reach your full potential, and, of	Relapses make it harder for you to recover. Usually, fewer relapses happen with clozapine since it is the most effective treatment for your condition. Long-term treatment results are better with clozapine. <sup>16</sup>	Is it important to you to have a better long- term treatment results and fewer relapses?		
d.	POSSIBLE QUT		You are being offered clozapine because you have symptoms that are not responding to other medications	Is it important for you to take the medication with the best chance of response?		
	What are the benefits of taking effective medications?		and clozapine offers the best chance of response. It is the gold standard of care for your condition. *	Is it important for you to take the gold standard treatment for your condition?		
cian administered naire uses	Being well is also called "remission". If the medication you are taking does not fully		Some patients who are now taking multiple medications might be able to take fewer medications when taking clozapine. <sup>4</sup>	Is it important to you to simplify your medication plan?		
	treat your symptoms, you cannot function well. You may need to be in the	Orusic Relayaer	The scheduled laboratory monitoring allows you to be more connected to your treatment team.	Is it important to you to stay connected to your treatment team?		
onal interviewing	hospital more often, as you will have entries worsening symptoms (relapses). Having relapses aftects your brain health and	mul - 000 ==-	Since clazapine causes fewer relapses, people taking clazapine can better protect their brain compared to people taking less effective medication.	Is it important to you to protect your brain from the illness?		
s to guide the	prevents you from recovering fully, as shown in the graph to the side.					NO
on thereby allowing			RISKS AND REASONS TO AVOID CLOZAPINE You may experience minimal discomfort when getting your	Are you able to tolerate some discomfort	YES	
ent to reflect on	LESPONSE TO CLOZAFINE COMPARED TO ANY OTHER ANTIPSTICHORIC 105	ine reduces psycholic symptoms, risk of	blood tests, but most patients tolerate this well. Optional question depending on availability of capillary blood	caused by regular blood testing? If you are uncomfortable with regular blood		
Itment goals and	2 40 neonle	length of hospitalization, and mortality for that have not shown a significant	testing: If you responded NO to the previous question:	tests, would this device be an acceptable alternative for you?		
ne if clozapine is	antipsyc		For people uncomfortable with blood tests, there is a device that allows for regular monitoring with a simple prick of a finger. A small drop of blood is then collected from the tip of the finger and is used for testing.	If yes, ask your doctor if this device is available in your area.		
help them. At the	any me	u have tried two different medications, trying dication other than clozapine has a very low	From the tip of the tinger and is used for testing. Blood tests are initially administered weekly then every two weeks and after 1 year, every 4 weeks (monthly). <sup>44</sup>	Are you able to tolerate the inconvenience of the blood monitoring schedule?		
ne questionnaire,	This is with	e of making you feel better, roughly 10-20%. hy we are suggesting that you try clozapine, ine has a high chance of improving your	Clozapine side effects may include sedation (tiredness), weight gain, hypersalivation (drooling or excess saliva),	Are you willing to exercise, eat healthy foods and take additional medication to manage		
ent is invited to		ms, with a response rate of 60-77%.2	constipation and palpitations. Most of these side effects can be adequately managed with diet, exercise, or other medication.**	side effects if they occur?		
decision about ozapine if he/she is			Like all of the medications offered to you, clozapine does have potential side effects. To gain a better understanding of clozapine's side effects, we should discuss them in comparison with any alternative medication.	Do you think you would be able to tolerate some of the potential side effects of clozapine?		

#### Discussion

OPTIC provides the necessary information to initiate a balanced discussion between clinician and clozapine eligible patient. It changes the conversation from a list of clozapine's many side effects to a discussion about how its benefits on reducing TRS symptoms significantly outweigh its side effects. The main objective is for clozapine to be adequately offered systematically to people with TRS as soon as they become eligible. This tool can facilitate a positive offer and increase acceptance of clozapine. The OPTIC tool is based on the OPTIMA tool which discusses the place of LAI in treatment of psychosis.

#### Conclusion

We hope that the use of the OPTIC tool will facilitate positive offers of clozapine thus increasing its acceptance amongst patients with TRS as soon as they become eligible. This will hopefully be translated by an increase in the number of TRS patients effectively treated with clozapine. The OPTIC tool has not yet been distributed as it was just recently finalized, therefore its impact on clozapine prescription rates cannot be measured at this time. It will be available for free on the Canadian Consortium for Early Intervention in Psychosis website (epicanada.org) under clinical tools.

Howes, Oliver D et al. "Treatment-Resistant Schizophrenia: Treatment Response and Resistance in Psychosis (TRRIP) Working Group Consensus Guidelines on Diagnosis and Terminology." The American journal of psychiatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychosis." Journal of clinical psychoatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychosis." Journal of clinical psychoatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychosis." Journal of clinical psychoatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychoatry." The American journal of psychiatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychoatry." The American journal of psychiatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychoatry." The American journal of psychiatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychoatry." The American journal of psychiatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly responding first-episode psychoatry." The American journal of psychiatry vol. 174,3 (2017): 216-229. Aqid, Ofer et al. "Early use of clozapine for poorly respondence psychiatry." 27,4 (2007): 369-73.

Williams, Richard et al. "What Is the Place of Clozapine in the Treatment of Early Psychosis in Canada?." Canadian journal of psychiatry. Revue canadienne de psychiatrie vol. 62,2 (2017): 109-114.

3. Canadian Psychiatric Association. Clinical practice guidelines: treatment of schizophrenia. The Canadian Journal of Psychiatry. 2005;50(13):1s-57s.

Margolese HC, Steiner W, Lalla F, Cattan C, Perillo A, Arshoff L. Development of a patient decision aid to educate patients with acute psychoses about long-acting injectable antipsychotic therapy. Institute on Psychiatric Services (IPS) 2016: The mental Health Services Conference, October 6-9 2016, Washington DC

# **OPTIC Consists of 3 Parts:**

https://www.epicanada.org/optic

- 1. Healthcare Provider Information Brochure
  - Provides information about clozapine
- 2. Patient Handout
  - Gives basic information about why clozapine is a good choice for them
- 3. Decision Aid Questionnaire
  - Uses questions to facilitate discussion of benefits and risks of clozapine vs. other treatments



#### **OPTIC - Patient Handout**

- A one-page double sided handout
- Provides information to guide the discussion:
  - Why are you being offered clozapine for your symptoms?
  - Effective medication helps you stay well
- What you need to know about clozapine:
  - Common and rare but serious side effects
  - Laboratory monitoring



## **OPTIC - Decision Aid Questionnaire**

- Questions are designed to highlight the potential benefits and discuss the potential side effects in a balanced manner
- At the end you ask for a decision and leave room of course for further discussion at another visit.

#### What are your options?

- A) Take clozapine
- B) Do not take clozapine
  - Talk to your doctor about other antipsychotic medication.



#### **TMAS: Tool for Monitoring Antipsychotic Side Effects**

https://www.epicanada.org/tmas

## Tool for Monitoring Antipsychotic Side Effects (TMAS)

#### Tool for Monitoring Antipsychotic Side Effects (TMAS)

#### Person's Name:

WHY MONITOR? Schizophrenia<sup>8, 34</sup> and use of antipsychotics<sup>13, 19, 24, 32, 33, 36</sup> are independently associated with increased motor and metabolic abnormalities, which can contribute to non-adherence to medication, and increased motoidity and mortality<sup>6, 18, 18, 22, 27, 38, 39, 41, 42</sup>. MINIMUM MONITORING FREQUENCY: For newly initiated medication: baseline, 1 (motor side effects only), 3 and 12 months. For persons on the same medication > 1 year: q 12 months.

Medical History	No	relevant moto	r/neurological histo	ry 🗖 Mo	tor/neurological c	isorders
Details:			/	,		
Family History in First Degree Relativ	ve 🛛 No	relevant moto	r/neurological histo	ry 🗖 Mo	tor/neurological c	lisorders
Details:			/	.,		
Assessment Date (dd/mm/yy)						
Assessment Completed By:						+
Automatical Completion Dy:						
Current Medication(s)						
SUBJECTIVE EXPERIENCE	SCORE:	0 = NONE	1 = QUESTIONABLE	2 = MILD	3 = MODERATE	4 = SEVERE
(≤ 1 week)	Score	Score	Score	Score	Score	Score
Parkinsonism						
Dyskinesia						
Akathisia						
Dystonia						
PARKINSONISM	SCORE:	0 = NONE	1 = QUESTIONABLE	2 = MILD	3 = MODERATE	4 = SEVERE
Score right/left sides as indicated	Score	Score	Score	Score	Score	Score
Facial expression – reduced						
Hands – tremor (resting)	R	R	R	R	R	R
rialias – rielior (resilig)	L	L	L	L	L	L
Hands – tremor	R	R	R	R	R	R
(with arms extended, fingers apart)	L	L	1	L	L	L
Hands – bradykinesia	R	R	R	R	R	R
Hanas – bradykinesia	1	L	1	L	L	1
Clip and a similarity	R	R	R	R	R	R
Elbow – rigidity	L	L	L	L	L	L
Gait – abnormality						
DYSKINESIA	SCORE:	0 = NONE	1 = QUESTIONABLE	2 = MILD	3 = MODERATE	4 = SEVERE
Score right/left sides as indicated	Score	Score	Score	Score	Score	Score
Face and mouth – with activation						
Tongue – with activation						
-	R	R	R	R	R	R
Upper extremities – arms, hands	L	L	1	L	L	L
Trunk – neck, shoulders, hips						
	R	R	R	R	R	R
Lower extremities – ankles/toes	L	L	1	L	L	L
	SCORE:	0 = NONE	1 = QUESTIONABLE	2 = MILD	3 = MODERATE	4 = SEVERE
AKATHISIA	Score	Score	Score	Score	Score	Score
Observed motor restlessness						
	SCORE:	0 = NONE	1 = QUESTIONABLE	2 = MILD	3 = MODERATE	4 = SEVERE
DYSTONIA	Score	Score	Score	Score	Score	Score
Observed dystonia						
Details – name affected body part	1					+
(e.g. head, extremities, trunk):						

#### B. ISSUE/ACTION/OUTCOME

DATE	ISSUE	ACTION	OUTCOME
		1	

#### **Tool for Monitoring Antipsychotic Side Effects (TMAS)**

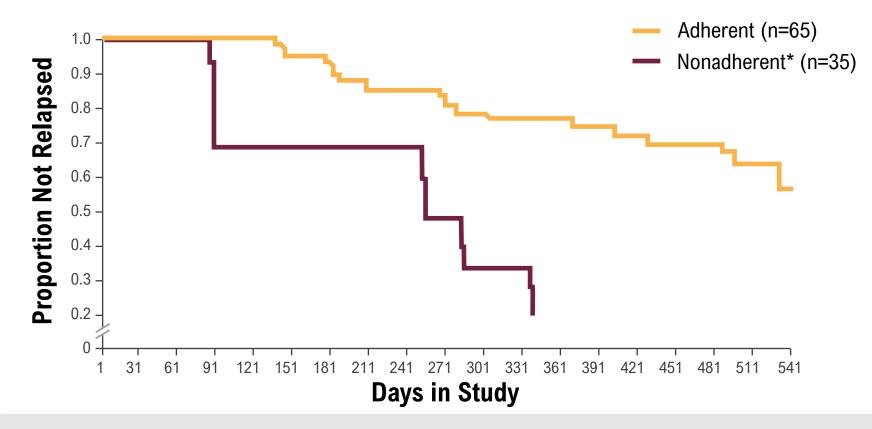
Person's Nam	e:						
C. METABOLIC	SIDE EFFECTS						
Baseline Medical	History Date:	V	√eight:	Height:	8	$3M\left(\frac{Weight [kg]}{Height [m]^{1}}\right) =$	
No relevant m	etabolic history	Cardiovasa	ular 🗖 Dysl	ipidemia	Diabetes		ertension
Obesity (BMI>)	>30)7	Smoker	D Sed	entary lifestyle (·	< 30 min exercis	e at least 4 days	
Details:							
	First Degree Relative				Distantes		· · · ·
Cardiovascula Details:	r (< age ov)	Hypertensic	in 🖬 Dysi	ipidemia	Diabetes	D Obe:	ыту
Jeiulis.							
Assessment Date	(dd/mm/yy)						
Assessment Com	pleted By:						
Date Requisition (*if applicable)	Provided						
Date Blood Work (*if applicable)	Completed						
Current Medicatio	on(s)						
Risk Factor	Abnormal Level	Test Results	Test Results	Test Results	Test Results	Test Results	Test Results
Weight	≥ 5% increase						
kgs/lbs	from baseline		_		_	_	
Waist	(i.e.=kgs/lbs)	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
vv aist circumference <sup>7†</sup>	M >102 (40)						
cm (inches)	F > 88 (35)	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
Blood pressure <sup>35</sup>	> 140/90 or						
mmHg	> 130/80						
	if diabetic	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
Triglycerides <sup>3#</sup>	> 1.7						
nmol/L	- 1.7	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
asting glucose <sup>3*</sup>							
nmol/L	> 5.6						
,		Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
HDL cholesterol <sup>3*</sup>	M ≤ 1.03						
nmol/L	F ≤ 1.30	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
LDL cholesterol4*	≥ 5.0						
mmol/L	2 3.0	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
Hemoglobin A1c	< 6.0 normal						
as indicated)*	< 7.0 for most						
%	diabetic persons	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result	Abnormal result
	an Diabetes Association	n provides gende	r and ethnicity ba	sed guidelines fo	r waist circumfer	ence	
D. OTHER SIDE EFFECTS DATE SIDE EFFECT NOTE				DATE	SIDE EFFECT NOTED		
	(e.g. GI, sedation, sexu	al, etc.)		24.1	(e.g. Gl, sedat	ion, sexual, etc.)	
					+		

#### E. ISSUE/ACTION/OUTCOME

E. ISSERVENEN/OFFCOME					
DATE	ISSUE	ACTION	OUTCOME		
2					

#### **Treatment Adherence and Relapse**

# Nonadherence Predicts Relapse in Patients with Recent-Onset Schizophrenia



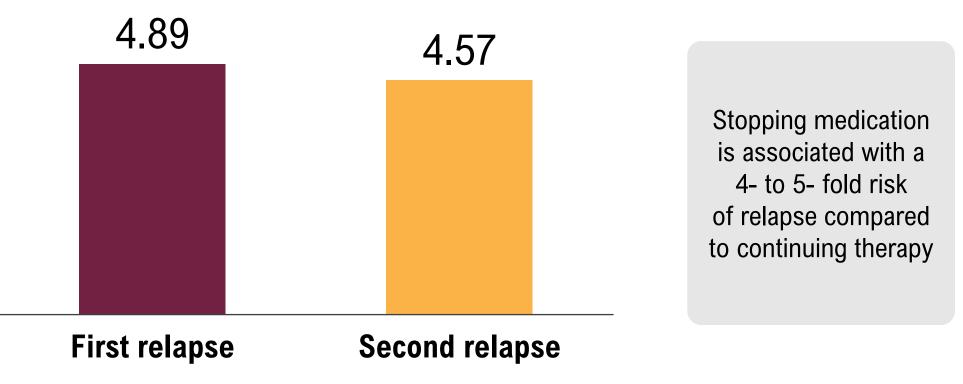
Missing as little as 25% of the prescribed dosage over a period of ≥2 weeks significantly raised the risk of returning psychotic symptoms.

\*For this study, nonadherence was defined as patients with <50% adherence of the prescribed medication dose for at least 2 weeks.

1. Subotnik KL et al. Am J Psychiatry 2011; 168(3):286-92.

## **Stopping Medication is the Most Powerful Predictor of Relapse**

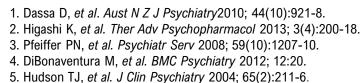
## Hazard Ratio compared to continuing therapy (n=104)



1. Adapted from Robinson D, et al. Arch Gen Psychiatry 1999; 56:241-7.

## **Key Contributors to Nonadherence**

- Lack of insight<sup>1,2</sup>
- Medication beliefs<sup>2</sup>
- Substance abuse<sup>2</sup>
- Complex dosing regimens<sup>3</sup>
- Adverse events<sup>4</sup>
- Stigma of antipsychotic medication<sup>5</sup>





## **Key Contributors to Relapse**

- Nonadherence 4X
- Substance use 3X
- Highly critical family / Significant others 2X
- Family history / Biology 2x



#### How Long Should We Continue Treatment After An Initial Episode?

Key question with no clear answers.



## Q & A

# To submit a question please use the "Ask A Question" button on the top right of your screen.





#### To attend the next presentation, please click your preferred Concurrent Session topic from the agenda below your video player.