

# Predictors of second-generation antipsychotic-induced weight gain: a longitudinal study with antipsychotic-naïve patients

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## Introduction

- Weight gain (WG) is a common adverse event of second-generation antipsychotics (SGAs) associated with a significant increase in morbidity, social stigma, and treatment non-adherence [1, 2].
- Retrospective studies report a higher risk of WG in the first 3 months of treatment. Predisposing factors include premorbid BMI, female sex, younger age, and being diagnosed with a psychotic disorder [3].

## Objectives

- To assess short-term (i.e. baseline to 6 weeks) and medium-term (6 weeks to 6 months) WG in a cohort of paediatric and adult patients naïve to SGAs.
- To study the differential predictive value of sociodemographic variables and of anthropometric and metabolic variables at baseline associated with the short- and medium-term WG.

## Methods

- Six-month longitudinal, observational, multicenter study.
- N=208 antipsychotic-naïve pediatric and adult patients (lifetime exposure to SGAs fewer than 10 days)
- Assessments:** anthropometric changes within two time periods:  
Basal - 6-weeks (short-term): weight and BMI changes  
6 weeks - 6months (medium-term): weight and BMI changes
- Metabolic markers:**

Total Cholesterol	Glucose	Leptin
LDL-cholesterol	Insulin	Adiponectin
HDL-cholesterol	HbA1c	Triglycerides
- Two multivariate regression analyses were performed to estimate the effect of sociodemographic, anthropometric and metabolic factors at baseline on short- and medium-term WG.

<sup>1</sup>HbA1c: Glycated Hemoglobin  
BMI: Body-mass Index

## Results

### Sociodemographic Variables

208	
N	
Age	Mean 37,97±20,48 years
	>18 years 74,6%
	Range 13-90
Males	56%
Psychotic disorder	56,8 %



Figure 2: Mean Daily BMI Change



Figure 3: Mean Daily Weight Change (gr)

### BASAL-6 WEEKS REGRESSION MODEL VARIABLES INCLUDED

	Confidence Interval (95%)		Correlations		Explained Variance (p)
	Inferior Limit	Superior Limit	Parcial	Semipartial	
Gender (females)	-0,934	-0,408	-0,263	-0,256	<b>6,55% *&lt;0,001</b>
BMI_Basal	-0,108	-0,05	-0,278	-0,272	<b>7,40% *&lt;0,001</b>
Leptina_Bas	0,003	0,026	0,138	0,131	<b>1,72% * 0,011</b>
Whole model adjusted*				0,342	<b>11,7% * 0,011</b>
Dependent Variable: Change BMI Baseline to 6 weeks					

\*After adjusting the model for age, sex, baseline BMI, leptin, glucose, insulin, HbA1c, adiponectin, cholesterol, and triglycerides).

Table 1: Change BMI baseline-6weeks regression model

### 6 WEEKS-6 MONTHS REGRESSION MODEL: VARIABLES INCLUDED

	Confidence Interval (95%)		Correlations		Significatio n
	Inferior Limit	Superior Limit	Parcial	Semiparci al	
Adiponectin Basal	0,026	0,105	0,195	0,195	<b>1,88% * 0,022</b>
Leptin Basal	-0,042	-0,003	-0,139	-0,137	<b>3,80% * 0,001</b>
Whole model adjusted*				0,210	<b>4,4% * 0,022</b>
Dependent Variable: Change BMI 6 weeks-6 months					

\*After adjusting the model for age, sex, baseline BMI, leptin, glucose, insulin, HbA1c, adiponectin, cholesterol, and triglycerides).

Table 2: Change BMI 6 weeks-6 months regression model

## Conclusions

- Most of the weight gain during the six month follow-up takes place in the first 6 weeks of treatment..
- In the short-term period male gender, basal-BMI, and leptin levels predict WG.
- In medium-term basal leptin and adiponectin levels predict WG.
- Most WG can not be explained by the variables studied.

## References

- [1] Newcomer JW., 2005. Second-generation (atypical) antipsychotics and metabolic effects: a comprehensive literature review. *Drugs, 19Suppl 1: 93-98*
- [2] Kane JM, Ishiwata T, Cornell CU., 2013. Non-adherence to medication in patients with psychotic disorders: epidemiology, contributing factors and management strategies. *World Psychiatry, Oct;12(3):216-26*
- [3] Gebhardt S, Heberhausen M, Heinzl-Gutenbrunner M, Gebhardt N, Remschmidt H, Krieg JC, Hebebrand J, Theisen FM., 2009. Antipsychotic-induced body weight gain: predictors and a systematic categorization of the long-term weight course. *J Psychiatr Res, Mar;43(6):620-6*

## DISCLOSURES:

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