

**To:** CN=Patrizia Cavazzoni/OU=AM/O=LLY@Lilly; CN=Jared G Kerr/OU=AM/O=LLY@Lilly  
**CC:** CN=Abelardo Altamirano/OU=AM/O=LLY@Lilly; CN=Steve Bandak/OU=AM/O=LLY@Lilly;  
CN=Florence Leroux/OU=AM/O=LLY@Lilly; CN=Paulo T Resende/OU=AM/O=LLY@Lilly;  
CN=Javier Vela/OU=AM/O=LLY@Lilly  
**Date:** 07/08/2002 06:02:52 PM  
**From:** CN=Elizabeth Brunner/OU=AM/O=LLY  
**Subject:** Regulatory request in Mexico concerning Hyperglycemia

Jared and Patrizia

As per my telephone communication with both of you regarding the Mexican MOH request to include an package insert regarding Hyperglycemia for Zyprexa, I am sending you an exact translation of the document:  
The letter was directed to our Medical Director, Abelardo Altamirano, we received this on July 1st.

"In virtue of the reports of hyperglycemia associated with the use of Olanzapine, we are requesting that in the Zyprexa presentation you include information in the way of "insert" , in a simple and clear text in order for it to be understood by the patients, the following:

- Route of administration, schedule and relationship with meals (if any)
- Most frequent and important side effects, pointing out which could be alarm symptoms for them to consult a physician
- Drug-drug interactions
- Precautions
- Contraindications

We suggest that the information be no longer than 1 page.  
This information should be submitted for it's approval at this office, without the required payments, no longer that 1 month after the date of reception of this document"

Here is what the Mexican PI states :

**ADVERSE EVENTS:**

- Diabetic Coma <0.01%
- Hyperglycemia <0.01%
- DKA <0.01

**Abnormalities in Laboratory Tests:**

"Very occasional random plasma glucose levels from clinical studies  $\geq$  200 mg/dl (suggestive of potential diabetes), and glucose levels  $>$  160 mg/dl but  $<$  200 mg/dl (suggestive of potential hyperglycemia) in patients with baseline plasma levels  $\geq$  140 mg/dl."

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We responded with the global response document (excluding the information on Japan) and a letter of disapproval to their request. This document was sent July 4th. We will meet with them on Friday 12 and will keep you updated.

Please let me know what else is needed.

regards, Eliabeth Brunner  
CRP Mexico

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