## EXHIBIT 33

# **Diabetes Information**

Prevalence in the US population				
	General population	People with schizophrenia	People with bipolar disorder	
diabetes	6%-7%¹²²	13%-18%	10%-26%	
smoking	34%-39% 57	74%-90%°	35%*-55%*.10	
obesity (вмг≥зо)	20%²-31%¹¹	40%-60%12	30%³	

\*For mania

### **Testing and Monitoring Patients on Atypical Antipsychotics**

- Any patient treated with atypical antipsychotics should be monitored for symptoms of hyperglycemia including polydipsia, polyuria, polyphagia, and weakness¹³
- ADA recommends that patients' weight be reassessed at 4, 8, and 12 weeks after starting on an atypical antipsychotic, and then quarterly during routine visits¹⁴

Patients	Starting on an Atypical Antipsy	ichotic <sup>19</sup>
With an established diagnosis of diabetes	With risk ractors for diabetes	Any patient
Monitor regularly for worsening	■ Test fasting glucose at the start	疆'Monitor for symptoms
of glucose control	of treatment	of hyperglycemia
	Test fasting glucose periodically during treatment	
	curry accurrent	

Diabetes Risk Factors for Asymptomatic Adults¹⁵			
<b>S</b> Age:≥45 years	■ Hypertension (BP ≥149/90 mmHg)		
■BMI ≥25 kg/m²	■ HDL–C <35 mg/dL and/or triglycerides >250 mg/dL		
■ Habitual physical inactivity	■ Polycystic ovary disease		
■First-degree relative with diabetes	■ IGT or IFG on previous testing		
■ High-risk ethnic group (eg, African American, Latino, Native American, Asian American)	Other clinical conditions associated with insulin resistance (eg, acanthosis nigricans)		
■ Having delivered a baby weighing >9 lb or having been	■History of vascular disease		
diagnosed with gestational diabetes	■History of smoking		

See Important Safety Information on reverse side and accompanying full Prescribing Information.



### Diabetes Information

- Hyperglycemia, in some cases extreme and associated with ketoacidosis, hyperosmolar coma, or death, has been reported in patients treated with atypical antipsychotics, including SEROQUEL
- The relationship of atypical use and glucose abnormalities is complicated by the possibility of increased risk of diabetes in the schizophrenic population and the increasing incidence of diabetes in the general population<sup>13</sup>
- The results of retrospective studies of SEROQUEL and diabetes have been discrepant¹⁴
- Postmarketing reports of diabetes or diabetes-related events are very rare (<0.01%) with SEROQUEL.

  These reports were confounded by preexisting or coexisting risk factors and/or had limited information¹⁵
  </p>
- SEROQUEL is an atypical that has had over 16 million patient exposures worldwide since its approval in 1997. AstraZeneca believes that the available scientific and medical data do not establish that SEROQUEL causes diabetes

#### Important Safety Information

SEROQUEL is indicated for the treatment of acute manic episodes associated with bipolar I disorder, as either monotherapy or adjunct therapy with lithium or divalproex, and the treatment of schizophrenia. Patients should be periodically reassessed to determine the need for continued treatment.

Elderly patients with dementia-related psychosis treated with atypical antipsychotic drugs are at an increased risk (1.6 to 1.7 times) of death compared to placebo (4.5% vs 2.6%, respectively). SEROQUEL is not approved for the treatment of patients with dementia-related psychosis.

Prescribing should be consistent with the need to minimize the risk of tardive dyskinesia. A rare condition referred to as neuroleptic malignant syndrome has been reported with this class of medications, including SEROQUEL.

Precautions include the risk of seizures, orthostatic hypotension, and cataract development.

The most commonly observed adverse events associated with the use of SEROQUEL in clinical trials were somnolence, dry mouth, dizziness, constipation, asthenia, abdominal pain, postural hypotension, pharyngitis, SGPT increase, dyspepsia, and weight gain.

Patients starting treatment with atypical antipsychotics who have or are at risk for diabetes should undergo fasting blood glucose testing at the beginning of and during treatment. Patients who develop symptoms of hyperglycemia should also undergo fasting blood glucose testing.

#### See accompanying full Prescribing Information.

References: 1. CDC. National diabetes fact sheet. Available at: http://www.cdc.gov/diabetes/pubs/estlmates.htm. Accessed April 7, 2005. 2. Mokdad AH, Bowman BA, Ford ES, et al. JAMA. 2001;286: 1195-1200. 3. Regenold WT. Thapar FIK, Marano C, et al. J Affect Disord. 2002;70:19-26. 4. Bushe C, Holl R. Br J Psychiatry. 2004;184(suppl 47):s67-s71. 5. Cassidy F, et al. Elevated frequency of diabetes mellitus in hospitalized manic-depressive patients. Am J Psychiatry. 1999;155:1417-1420. 6. Gopalaswamy AD. Smoking in chronic schizophrenia. Br J Psychiatry. 1986;149:523. 7. Masterson E, O'Shea B. Smoking and mellignancy in schizophrenia and the motivation for smoking. Available on-line at: http://www.findarticles.com/particles/mi\_qas804/s\_2002/dal\_np030259/print. Accessed on April 22, 2005. 9. Grant BF, et al. Nicotine dependence and psychiatric disorders in the United States: results from the national epidemiologic survey on alcohol and related conditions. Arch Gen Psychiatry. 2004;15:1107-1115. 10. Ucok A, et al. Cigarette smoking among patients with can be paid and bipolar disorders. Psychiatry Cilin Neurosci. 2004;58:434-437. 11. AOA. AOA Fact Sheets: obesity in the U.S. Available on-line at: http://www.brasteristeristy.org/subs/fastacts/obesity\_U.S.shtml. Accessed May 2, 2005. Cigarette 12. Catapano L, Castle D. Obesity in schizophrenia with can be done about it? Australasian Psychiatry. 2004;12:23-25. 13. SERICOUEL!" (quetiapine furnerate) (package insert), Wilmington, DE: AstraZeneca Pharmaceuticals LP, 2004. 14. American Diabetes Association, American Association for the Study of Obesity. Consensus development conference on antipsychotic drugs and obesity and dlabetes. Diabetes Care. 2004;27:596-601. 15. American Diabetes Association, Standards of medical care in diabetes. Diabetes Care. 2005;28(suppl 1):S4-536. 16. Data on the, DA-SER-30.

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