

prospectively (2 years). Results presented in this report were based on data from patients with at least 12 months of available data in the Netherlands.

Results: There are 190 patients currently enrolled in the Netherlands and 118 patients have at least 12 months of available data. Of the 118 patients, the majority were male (62.7%) with a mean age of 37.7 ± 11.5 years and a mean time since schizophrenia diagnosis of 11.1 ± 21.5 years. The main reasons for switching to RLAI were lack of compliance (42.4%), adverse events (25.4%) and lack of efficacy (24.6%) with previous therapy. At 12 months, 66.9% of patients were still on RLAI treatment. Of the patients who discontinued RLAI, the mean time to discontinuation was 157.8 ± 76.5 days. Mean CGI-S score significantly improved from 4.05 ± 1.14 at baseline to 3.15 ± 1.38 at 12 months ($p < 0.001$). Additionally, the mean GAF score significantly improved from 43.8 ± 12.0 at baseline to 55.2 ± 14.7 at 12 months ($p < 0.001$).

Conclusion: These interim results showed that treatment with RLAI in patients with schizophrenia was associated with significant improvements in disease severity and functioning.

P0278

Patient and physician satisfaction with risperidone long-acting injection: 18-month interim results from the electronic schizophrenia treatment adherence registry in Belgium

J. Peuskens¹, M. Povey², J. Van der Veken³, A. Jacobs⁴, A. Lam⁵.
¹ *Universitair Psychiatrisch Centrum, KU Leuven Campus UC St.Jozef Kortenberg, Kortenberg, Belgium* ² *SGS Life Science Services, Wavre, Belgium* ³ *Janssen-Cilag, Berchem, Belgium* ⁴ *Johnson & Johnson Pharmaceutical Services, Beerse, Belgium* ⁵ *Johnson and Johnson Pharmaceutical Services, Toronto, ON, Canada*

Objectives: To evaluate patient and physician satisfaction with risperidone long-acting injection (RLAI) in patients with schizophrenia enrolled in the electronic Schizophrenia Treatment Adherence Registry (e-STAR) in Belgium.

Methods: e-STAR is an ongoing, international, prospective, observational study of patients with schizophrenia who start RLAI during their routine clinical management. Treatment satisfaction was assessed by the patient and physician on a 5-point scale from 'very good' to 'very bad'.

Results: 135 patients with mean age 40.9 ± 14 years and duration of illness 9.5 ± 9.2 years initiated treatment with RLAI, followed-up for at least 18 months were included in this analysis. At baseline, only 29.2% of patients expressed "good" or "very good" satisfaction while 21.1% of them expressed "bad" or "very bad" with their previous treatment. Similarly at baseline, 38.2% of physicians reported "good" or "very good" level of satisfaction and 14.6% rated their satisfaction as "bad" or "very bad" at that time. After initiation of RLAI, both patient and physician satisfaction with treatment improved dramatically. At 18 months, 76.5% of patients were satisfied ('good' or 'very good') with RLAI treatment and only 2.4% felt 'bad' and none reported 'very bad'. Physicians also expressed satisfaction with RLAI with 82.1% of them rated it as 'good' or 'very good'. Only one physician reported satisfaction below 'moderate'.

Conclusions: The low levels of patient and physician satisfaction with treatment prior to RLAI are likely to be a key decision driver to change therapy. After starting treatment with RLAI, both patient and physician satisfaction with the treatment substantially improved.

P0279

RGH-188, a d3/d2 dopamine receptor antagonist/partial agonist atypical antipsychotic candidate

I. Laszlovszky¹, B. Kiss², I. Gyertyan², G. Pasztor Meszaros², N. Seneca³, E. Schmidt², Z.S. Nemethy², G.Y. Bugovics², K. Saghy², J. Laszy², M. Kapas², G.Y. Nemeth¹, Z.S. Szombathelyi². ¹ *Gedeon Richter Plc., Medical Division, Budapest, Hungary* ² *Gedeon Richter Plc., Pharmacological and Drug Safety Research, Budapest, Hungary* ³ *Karolinska Institute, Department of Clinical Neuroscience, Psychiatry Section, Stockholm, Sweden*

Objectives: RGH-188 is an orally active, potent dopamine D3/D2 receptor antagonist/partial agonist atypical antipsychotic for the treatment of schizophrenia and bipolar mania.

Results: RGH-188 displayed high affinity to human D3 receptors (Ki: 0.085 nM) and approximately six- and thirty-times less affinity to human D2, and 5-HT1A receptors. In various in vitro and in vivo assays RGH-188 behaved either as an antagonist or as a partial agonist on dopamine D3 and D2 receptors.

RGH-188 displayed potent antipsychotic activity (0.1-0.8 mg/kg) in rodent models such as apomorphine-induced climbing, amphetamine- and phencyclidine-induced hypermotility, conditioned avoidance response. It significantly improved the learning performance of rats (0.02-0.2 mg/kg) impaired by scopolamine in a water-labyrinth learning paradigm. RGH-188 showed no EPS liability as it produced no catalepsy up to 100-fold therapeutic range.

In a nonhuman primate positron emission tomography (PET) study using ¹¹C-raclopride RGH-188 occupied striatal D2/D3 receptors in a dose dependent and saturable manner with an ED50 of 7 µg/kg iv. In healthy male subjects multiple administration of 1 mg RGH-188 resulted in over 70% D2/D3 receptor occupancy and the displacement showed correlation with RGH-188 and metabolites plasma levels.

After single administration to healthy volunteers, Tmax for RGH-188 was 3-4 hours and the terminal disposition half-life was 5-6 days. Over the dose range of 0.5-2.5 mg AUC of the parent drug was approximately dose-proportional. Systemic exposure to the pharmacologically active metabolites, desmethyl- and didesmethyl-RGH-188 was 20-30% and 50-200% of that to the parent, respectively.

P0280

Effect of clozapine and its metabolites on the intracellular calcium concentration in cells of isolated rat islets

L. Hao, W. Gaohua, X. Weidong, W. Xiaoping, W. Huiling.
Department of Psychiatry, Remin Hospital of Wuhan University, Wuhan, China

Objective: To study the different effects of clozapine and its metabolites on the intracellular Ca²⁺ concentration ([Ca²⁺]_i) in cells of isolated rat islets.

Methods: Under low or high glucose (3.3 mmol/L or 16.7 mmol/L), the cells of isolated rat islets was treated with 1 mmol/L clozapine, desmethyl-clozapine and clozapine N-oxide respectively, blank control group was also set, [Ca²⁺]_i represented by fluorescence intensity was measured by laser scanning confocal microscope after cells were loaded with calcium sensitive fluorescent indicator Fluo-4/AM.