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Artwork from Hospital Audiences, Inc. (HAI), a nonprofit organization that inspires healing, growth and learning through access to the arts for the culturally underserved.

Janssen Global Public Health:
HIV Medicines Access & Partnerships Program

APRIL 2014

Janssen Pharmaceutica, N.V.
www.janssen.com
We believe this requires two important components: ensuring quality and affordable medicines are available sustainably, and that these medicines are used in the most appropriate way in people living with HIV/AIDS (PLWHA). As a key element of our new Janssen Global Public Health group, our global HIV Medicines Access & Partnerships Program (the Program) is working to fulfill our responsibility to PLWHA in resource-limited settings.

Janssen’s HIV Medicines in Resource-Limited Countries

More PLWHA have access to HIV medicines than ever before. Access to HIV medicines for PLWHA who are new to treatment (“first-line”) remains the top public health priority for countries with high HIV burdens. Yet, over time, many patients will need to switch to a new “second-line” regimen.

The Program has a portfolio of three HIV treatments. Currently, the protease inhibitor PREZISTA® (darunavir) and non-nucleoside reverse transcriptase inhibitor (NNRTI) INTELENCE® (etravirine) are indicated in resource-limited countries for use in treatment-experienced patients only after previous drug combinations have failed. They are often the last treatment option for patients in these countries. The Company’s third HIV medicine, the NNRTI EDURANT™ (rilpivirine), is currently indicated in resource-limited countries for use in treatment-naïve adults.

One of our key responsibilities is to facilitate access to these medicines once they are approved in a way that safeguards their clinical effectiveness today and for the future.

Key Program Elements

The Program aims to improve the health of people worldwide through sustainable availability of our HIV medicines, medical education on appropriate and safe use of these medicines, and innovative collaborations.

1 As actual approved uses are determined by local regulatory authorities, please see package insert for precise indication.
Branded Availability, Enabled Generics & Generic Licensing

Our approach to providing effective and sustainable access to our HIV medicines in resource-limited settings continues to evolve based on patient needs. Agreements with Aspen Pharmacare for our branded HIV medicines at Special Effort Access Prices in SSA provided rapid access for patients in need during the Program’s formative years. These royalty-free agreements helped to establish distribution networks, lay the groundwork for the appropriate clinical use of our HIV medicines, and support appropriate, associated pharmacovigilance activities. They also accelerated the registration process of our branded HIV medicines, which is an important first step in facilitating the introduction of quality generic versions. These agreements were designed to ensure timely and sustainable access to all our HIV medicines in SSA, which is the region with the highest HIV burden and economic vulnerability.

For darunavir, our HIV medicine for treatment-experienced patients, Janssen announced in 2012 that it would not enforce its patents on the medicine provided the darunavir product is medically acceptable and is used only in SSA and LDCs. This policy assures generic manufacturers that they may manufacture high quality darunavir that is used in SSA and LDCs without a concern that we will accuse them of infringing on darunavir patents.

For rilpivirine, our HIV medicine indicated for treatment-naïve adults as part of combination HIV treatment for which we anticipate greater volumes, we have additional licenses for the

In 2012, Johnson & Johnson was ranked second among 20 global companies in the biannual Access to Medicine Index based on significant progress and improvements made to our access to medicines strategies, R&D portfolio, and philanthropy initiatives.
manufacture of generic versions. These agreements, covering as many as 112 countries, offer licensees the technical information and knowledge ("tech transfer") to manufacture the API and finished product. They also enable development of appropriate FDCs of rilpivirine with other HIV medicines within the licensed territories. Licensing to globally-recognized generic manufacturers supports a sustainable supply of affordable and quality generic versions of our HIV medicines.

All of our agreements go beyond simply licensing our patents; they strive to ensure that critical components of HIV medicine access are achieved, including timely in-country registration, supply chain development, and drug safety monitoring. We believe our agreements and efforts provide the best route to expanding access to our medicines in resource-limited countries.

Priority Registration

A comprehensive, multi-country registration effort supports sustainable availability of our branded HIV medicines where there is a public health need. Our Program has prioritized registration efforts in those countries with the greatest HIV burden, economic vulnerability, and immediate need for third-line medications, such as PREZISTA® and INTELENCE®. In SSA, we target 20+ countries for priority registration. For those countries with limited public health need, and where local regulations allow, our products may be available through pre-approval access mechanisms. The 300mg and 600mg tablet of PREZISTA® and 100mg tablet of INTELENCE® have already received regulatory approvals in many countries in SSA. We have begun filings in the region for the 400mg tablet of PREZISTA® for use in treatment-experienced adult patients, and the 25mg tablet of INTELENCE® and 75mg and 150mg tablet of PREZISTA® for use in treatment-experienced pediatric patients.

EDURANT™ received its first regulatory approvals in the United States and Europe in 2011. This is an important first step, as many resource-limited countries require EMA and/or
## Summary of Branded Product Registration Status (as of April 2014)

<table>
<thead>
<tr>
<th>Tablet Strength</th>
<th>PREZISTA® (darunavir)</th>
<th>INTELENCE® (etravirine)</th>
<th>EDURANT™ (rilpivirine)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Tablet Strength</td>
<td>Adult (300mg and/or 600mg)</td>
<td>Adult (400mg)</td>
<td>Pediatric (75mg and 150mg)</td>
</tr>
</tbody>
</table>

### SSA & LDCs

| Approvals | 21 | 1 | 3 | 18 | 0 | 3 |
| Approvals Pending | 1 | 3 | 7 | 3 | 3 | 10 |

### LMICs

| Approvals | 15 | 11 | 6 | 13 | 2 | 8 |
| Approvals Pending | 0 | 1 | 5 | 2 | 6 | 5 |

FDA approval documentation to submit local regulatory dossiers. Regulatory filings for EDURANT™ are ongoing worldwide. In December 2013, EDURANT™ was approved in South Africa — its first regulatory approval in SSA.

We have reached a significant milestone in our efforts to provide affordable and sustainable access to quality HIV medicines, as three tablet strengths of our HIV medicines PREZISTA® and two tablet strengths of INTELENCE® have been included in the World Health Organization (WHO) List of Prequalified Medicinal Products. Having our HIV medicines on this list facilitates both the registration and procurement processes.

### Reduced Special Effort Access Pricing

We offer our branded HIV medicines at Special Effort Access Prices in SSA and LDCs and at reduced tiered prices in other lower-middle-income countries (LMICs) in the rest of the Program territory. These prices are significantly reduced from those in the US and Europe. We are committed to lowering prices, as volume or cost-savings in manufacturing allow.

- **In October 2011,** we reduced the Special Effort Access Price of PREZISTA® by 26% — to US$2.22 (ex-factory) for the 1200mg daily dose.

- **In July 2012,** we reduced the Special Effort Access Price of INTELENCE® by 52% — to US$1.20 (ex-factory) for the 400mg daily dose.

- **In June 2013,** we announced a reduced tiered pricing approach for EDURANT™, offering a price of US$0.11 per patient for the 25mg daily dose ($40/pppy) in SSA and LDCs and US$0.16 ($60/pppy) in other countries within the Program’s territory.

Beyond our Program’s territories, we provide reduced prices for our branded HIV medicines through differential pricing that considers local economic development, HIV treatment programs, and public health need. Prices in individual countries are subject to local pricing and reimbursement discussions.

### Medical Education & Clinical Research

Our HIV medicines require physicians to understand how to manage patients who have failed previous HIV treatment regimens. To assist with this, we support medical education on appropriate and safe use of HIV medicines, including accredited workshops on HIV treatment resistance and use of HIV third-line treatment. These workshops have reached more than 1,000 physicians in SSA countries and India. Additional workshops are planned for 2014. In December 2013, we launched Janssen eMedical Advisor, a new interactive training tool to help address the limited knowledge of Janssen’s HIV medicines among physicians in SSA.

We also support or conduct clinical research on the appropriate and safe use of our approved HIV medicines in resource-limited countries.

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*Antiretroviral Therapy for HIV Infection in Adults and Adolescents: Recommendations for a public health approach: 2010 revision. World Health Organization.*
We share clinical data to support inclusion in global, regional, and national HIV treatment guidelines, which are essential in designing effective HIV treatment strategies. In 2010, PREZISTA® and INTELENCE® were mentioned in WHO Guidelines for HIV Treatment as potential candidates for use in highly treatment-experienced adult and pediatric patients (“third-line”). They are not included in the guidelines for first- or second-line treatment.

**Pediatric HIV Treatment**

Our pediatric tablet formulations of PREZISTA® (75mg and 150mg tablets) are approved in the United States and Europe for treatment-experienced patients six to 18 years old, and regulatory submissions in resource-limited countries are underway. In December 2011, a twice daily regimen of PREZISTA® 100mg/ml oral suspension for use in treatment-experienced patients three to six years of age was approved by the FDA. In 2012, PREZISTA® 75mg and 150mg tablets, in combination with low-dose ritonavir and other antiretrovirals, were included in the WHO List of Prequalified Medicinal Products for treatment-experienced pediatric patients. In March 2012, INTELENCE® 25mg scored and dispersible tablet for treatment-experienced children and adolescents aged six to 18 years of age received FDA approval. In October 2013, this formulation was also included in the WHO List of Prequalified Medicinal Products. We are currently investigating a potential pediatric formulation of our HIV medicine EDURANT™.

In December 2013, we announced a first-of-its-kind donation program to improve access to our HIV medicines for children and adolescents failing HIV treatment in SSA. We are working with a number of organizations to fully maximize the impact and reach of this initiative.

**HIV Resistance and Diagnostics**

Antiretrovirals, or HIV medicines, are designed to stop the HIV virus from replicating. The HIV virus replicates frequently, which can lead to mistakes, or mutations, in the virus sequence. Some of these mutations can enable the virus to become resistant to one or more HIV medicines. This means the medicine no longer works and the virus starts replicating again, which is called “HIV drug resistance.” Understanding how HIV medicines work — and why they fail — is essential to providing effective and optimal HIV treatment. Janssen Diagnostics has long supported and provided technical assistance on resistance diagnostics, surveillance, and monitoring to better understand HIV resistance and resistance patterns in resource-limited settings. As a result, Janssen Diagnostics has built a database with the world’s largest collection — approximately 500,000 — of HIV virus sequences of which approximately 5% are non-B subtype, the most prevalent genotype in resource-limited countries. Janssen Diagnostics shares the information in this database through collaborations with research groups worldwide.

**HIV Prevention**

**Partnerships for HIV Prevention**

We are working to prevent the spread of HIV, and reduce the burden of HIV on women and their families. Microbicides are an innovative HIV prevention tool being investigated in various forms such as sustained-release vaginal rings to help prevent sexual transmission of HIV. The Company led the way in antiretroviral microbicides research. In 2004, it formed one of the very first public-private collaborations in the microbicides field with the International Partnership for Microbicides (IPM). The program provided IPM with a royalty-free license for TMC120 (dapivirine) to develop, manufacture, and distribute the compound as a vaginal microbicide in developing countries. The monthly dapivirine ring is now being evaluated in two parallel efficacy trials in multiple countries in Africa, with results expected in 2015.

**Rilpivirine (RPV, TMC278) Long-Acting**

Janssen recognizes that a long-acting, low-dose HIV medicine may someday play a role in pre-exposure prophylaxis (PrEP). Exploratory studies of an investigational long-acting, injectable formulation of rilpivirine (RPV-LA) are currently underway. A study presented at the 19th Conference on Retroviruses and Opportunistic Infections (CROI) in 2012, shared data supporting ongoing exploration of RPV-LA as a potential PrEP agent. The RPV-LA project is currently in

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the early stages of development and additional studies are needed to determine the next steps in the development of RPV-LA as a potential PrEP candidate.

**Opportunistic Infections**

**MicMAT (TIBOZOLE™)**

Oral candidiasis or “thrush” is the most common and debilitating opportunistic infection in HIV patients not receiving HIV medicines in the developing world. TIBOZOLE™ (miconazole nitrate muco-adhesive buccal tablet), also known as MicMAT, indicated for the treatment of oral candidiasis, is designed specifically to meet the unique needs of PLWHA in the developing world. More than 3.5 million MicMAT treatments have been made available through not-for-profit prices and donation programs.

**Our Products**

1. **PREZISTA® (darunavir)**

PREZISTA®, co-administered with 100mg ritonavir (PREZISTA®/rtv), and with other antiretroviral agents, is indicated for the treatment of human immunodeficiency virus (HIV) infection in antiretroviral treatment-experienced adult and pediatric (ages six to <18 years) patients, such as those with HIV-1 strains resistant to more than one protease inhibitor. This indication is based on week 24 analyses from two controlled clinical trials in treatment-experienced, HIV-1 infected patients, where PREZISTA®/rtv showed a significantly greater reduction of plasma HIV RNA levels and greater increase in CD4+ cell counts when compared to a protease inhibitor (PI) regimen of choice, each given in combination with other antiretroviral agents. Additional data is available from open label studies. Clinical studies on the use of PREZISTA®/rtv in antiretroviral treatment-naïve adult patients are ongoing.

2. **INTELENCE® (etravirine)**

INTELENCE®, in combination with a boosted protease inhibitor and other antiretroviral medicinal products, is indicated for the treatment of human immunodeficiency virus type 1 (HIV-1) infection in antiretroviral treatment-experienced adult patients and in antiretroviral treatment-experienced pediatric patients from six years of age. This indication is based on week 48 analyses from two randomized, double-blind, placebo-controlled Phase III trials in treatment-experienced patients with NNRTI resistance (present at screening and/or archived) and protease inhibitor (PI) resistance, where INTELENCE® administered with a background regimen (BR) was statistically superior to placebo with a BR in terms of the proportion of patients achieving a confirmed undetectable viral load (<50 HIV-1 RNA copies/ml) and the increase in CD4 cell counts from baseline. Treatment history and, when available, resistance testing should guide the use of INTELENCE®. The use of other active antiretroviral agents with INTELENCE® is associated with an increased likelihood of treatment response. The risks and benefits of INTELENCE® have not been established in treatment-naïve adult patients.

3. **EDURANT™ (rilpivirine)**

EDURANT™, in combination with other antiretroviral medicinal products, is indicated for the treatment of human immunodeficiency virus type 1 (HIV-1) infection in antiretroviral treatment-naïve adult patients with a viral load ≤ 100,000 HIV-1 RNA copies/ml. As with other antiretroviral medicinal products, genotypic resistance testing should guide the use of EDURANT™.