# Cost-effectiveness Analysis of Abacavir-based and Tenofovir-based regimens in Singapore Ong Y.Y.<sup>1</sup>, Hoo G.<sup>1</sup>, Law H.L.<sup>1</sup>, Ng O.T.<sup>2</sup> <sup>1</sup>Department of Pharmacy, Tan Tock Seng Hospital, Singapore <sup>2</sup>Department of Infectious Disease, Tan Tock Seng Hospital, Singapore



### Introduction

- Human immunodeficiency virus (HIV) infection has evolved into a chronic disease with the use of highly active antiretroviral therapy (HAART) <sup>(1,2)</sup>. As life expectancy of HIV-positive individuals improves, the duration of treatment increases. Cost of treatment will be a major factor affecting the choice of HAART.
- In addition to cost of medications, the cost of monitoring of short- and long-term side effects will also contribute to the total treatment cost of this disease.
- For management of HIV, the need for long-term treatment, good adherence and a low rate of antiviral resistance should also be evaluated in terms of lifetime costs.

Demographics Information		ABC-based regimen N (%)		TDF-based regimen N (%)	
		Male (n=105)	Female (n=10)	Male (n=105)	Female (n=10)
Age	Mean age (years) ± S.D.	46 ± 14.0			
Race	Chinese	95 (82.6)		94 (81.7)	
	Malay	14 (12.2)		13 (11.3)	
	Indian	3 (2.6)		6 (5.2)	
	Others	3 (2.6)		2 (1.7)	
Pharmacological	NRTI <sup>1</sup> /NNRTI <sup>2</sup>	97 (84.3)		92 (80.0)	
	NRTI/PI <sup>3</sup>	15 (13.1)		20 (17.4)	

# **Objectives**

To assess the cost-effectiveness of abacavir (ABC)-based and tenofovir (TDF)based regimens for treatment of HIV in Singapore, taking into account:

# (1) Cost of drugs alone(2) Cost of drugs and monitoring

## Methods

- Study design: This was a single-centre, retrospective study, conducted at the Communicable Disease Centre (CDC), Tan Tock Seng Hospital (TTSH). This study was approved by National Health Group Institutional Review Board.
- Inclusion criteria: (1) Have a positive western blot test and diagnosed with HIV by a medical doctor, (2) above 21 years of age, (3) currently on follow up at CDC, and (4) prescribed with either ABC-based or TDF-based regimens for at least 48 weeks.
- Exclusion criteria: (1) Patients who have less than two viral load (VL) readings,
  (2) diagnosed with HIV before the year 2006, (3) incomplete or missing case notes.
- $\blacktriangleright$  Matching criteria: Patients from the ABC group were matched with patients from

NRTI/INSTI <sup>4</sup>	3 (2.6)	3 (2.6)
Adherence (>95%)	107 (93)	105 (91.3)

<sup>1</sup>NRTI (Nucleot(s)ide reverse transcriptase inhibitors): Lamivudine, emtricitabine <sup>2</sup>NNRTI (Non-nucleotide reverse transcriptase inhibitors): Efavirenz, nevirapine <sup>3</sup>PI (Protease inhibitors): Ritonavir-boosted atazanavir, ritonavir-boosted lopinavir <sup>4</sup>INSTI (Integrase inhibitors): Raltegravir

Table 1. Patient characteristics (N=230)

#### p = 0.029 p = 0.006100 94 (86.2) 89 (78.1) 88 (74.1) 90 80 68 (60.7) 70 60 50 40 30 20 10 W12 to W36 W36 to W60 TDF ABC

#### Figure 1. Effectiveness of regimen at Week 24 and Week 48

#### Table 2: Incremental cost-effectiveness results

		Effectiveness measure	Difference in costs $(C_a - C_t)^a$	Difference in effectiveness $(E_a - E_t)^b$	Costs of medications only
Cm	Costs of medications only	Undetectable viral load (week 12 to week 36) <sup>c</sup>	-2309.33	0.173	-13348.72 Dominant
		Undetectable viral load (week 37 to week 60) <sup>d</sup>	-2309.33	0.121	-19085.37 Dominant
Un vira mo	Jndetectable viral load (3 to 9 nonths) <sup>c</sup>	Undetectable viral load (week 12 to week 36) <sup>c</sup>	-2361.46	0.173	-13650.05 Dominant
		Undetectable viral load (week 37 to week 60) <sup>d</sup>	-2361.46	0.121	-19516.20 Dominant

TDF group according to: (1) age group, (2) gender, (3) remaining two HAART agents prescribed, (4) body mass index (BMI), and (5) race.

Data analysis: Effectiveness is defined as percentage of patients who achieved undetectable VL at the period between week 24 and week 48. An incremental cost-effectiveness ratio (ICER) analysis was performed to evaluate the costeffectiveness between ABC-based and TDF-based regimens between the two evaluation periods. Costs were reported in Singapore dollars (S\$).

# **ICER:** $(C_a - C_t)/(E_a - E_t)$

- C<sub>a</sub>: total cost in the ABC group
- C<sub>t</sub>: total cost in the TDF group
- E<sub>a</sub>: effectiveness in the ABC group
- $E_t$ : effectiveness in the TDF group
- NOTE: 1<sup>st</sup> ICER: drug cost alone, 2<sup>nd</sup> ICER: drug cost and monitoring costs

# Results

- 230 patients were analyzed (82.2% Chinese, 91.3% male, age: 46.0 ± 13.0 years old), 115 patients in each group [Table 1].
- The most commonly used combinations were ABC, lamivudine (3TC) and

- <sup>a</sup> Difference in the total annualized drug cost between treatment groups [i.e. the cost in the ABC group (C<sub>a</sub>) minus the cost in the TDF group (C<sub>t</sub>)]. This is the numerator of the ICER.
- <sup>b</sup> Difference in the effectiveness endpoint between treatment groups [i.e. the proportion of patients achieving the endpoint in the ABC group (E<sub>a</sub>) minus the proportion of patients achieving the endpoint in the TDF group (E<sub>t</sub>)]. This is the denominator of the ICER.
- <sup>c</sup> Proportion of subjects achieving a undetectable viral during the evaluation period (weeks 12 to 36). <sup>d</sup> Proportion of subjects achieving a undetectable viral during the evaluation period (weeks 37 to 60).

## Discussion

- ABC-based regimen was found to be more cost effective than TDF-based regimen in HIV patients in Singapore for both evaluation periods (week 24 and week 48), regardless whether only cost of medications or cost of medications and monitoring were considered.
- Exclusion of some monitoring costs such as genotyping for HLAB\*5701 for ABC could make it even more cost-effective, especially for our Asian population (excluding Indians)<sup>(3)</sup>.
- One of the limitations we faced included the presence of external buyers' club where patients were able to obtain medications at cheaper prices or fixed-dose combinations that are not available at TTSH. Additional costs such as physician office visits and hospitalization costs for opportunistic infections were not included in the calculation of ICERs.

# efavirenz (EFV) (76.5%), and TDF, 3TC and EFV (78.2%); followed by ABC, 3TC, ritonavir boosted-atazanavir (ATV/r) (13.0%) and TDF, 3TC and ATV/r (11.3%) [Table 1].



- Majority of the patients were at least 95% adherent to their medication regimen (93.0% and 91.3% for ABC and TDF group respectively) [Table 1]. For both evaluation periods, more patients in the ABC group obtained undetectable VL (77.4% vs 59.1% and 81.7% vs 76.5%) [Figure 1].
- The ICER value was –S\$13348.72 for the period of week 24 (week 12 to 36) and –S\$19085.37 for the period of week 48 (week 37 to week 60) [Table 2].
- As public healthcare expenditure increase, this knowledge may be useful to physicians, policy makers, and tax payers in their efforts at making clinically appropriate yet cost-conscious decisions.



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