

Japan Tobacco Inc. Clinical Development as of February 4, 2016

<In-house development>

Code (Generic Name)	Potential Indication/Dosage form	Mechanism		Phase	Note
JTK303(elvitegravir)/ cobicistat/emtricitabine/ tenofovir alafenamide fumarate	HIV infection /Oral	HIV integrase inhibitor/ Nucleoside reverse transcriptase inhibitor	Suppresses blood HIV levels by inhibiting the activities of integrase and reverse transcriptase, enzymes involved in the replication of HIV.	Preparing to file (Japan)	JTK-303(elvitegravir); In-house Cobicistat, Emtricitabine, Tenofovir Alafenamide; In-license (Gilead Sciences)
emtricitabine/ tenofovir alafenamide fumarate	HIV infection /Oral	Nucleoside reverse transcriptase inhibitor	Suppresses blood HIV levels by inhibiting the activity of reverse transcriptase, an enzyme involved in the replication of HIV.	Preparing to file (Japan)	In-license (Gilead Sciences)
JTT-851	Type 2 diabetes mellitus /Oral	G protein-coupled receptor 40 agonist	Decreases blood glucose by stimulation of glucose-dependent insulin secretion.	Phase2 (Japan) Phase2 (Overseas)	In-house
JTZ-951	Anemia associated with chronic kidney disease /Oral	HIF-PHD inhibitor	Increases red blood cells by stimulating production of erythropoietin, an erythropoiesis- stimulating hormone, via inhibition of HIF-PHD.	Phase2(Japan) Phase1(Overseas)	In-house
JTE-052	Autoimmune/allergic diseases /Oral, Topical	JAK inhibitor	Suppresses overactive immune response via inhibition of Janus kinase (JAK) related to immune signal.	Phase2(Japan)	In-house
JTE-051	Autoimmune/allergic diseases /Oral	Interleukin-2 inducible T cell kinase inhibitor	Suppresses overactive immune response via inhibition of the signal to activate T cells related to immune response.	Phase1(Overseas)	In-house
JTE-151	Autoimmune/allergic diseases /Oral	ROR γ antagonist	Suppresses overactive immune response via inhibition of ROR γ related to Th 17 activation.	Phase1(Overseas)	In-house
JTT-251	Type 2 diabetes mellitus /Oral	PDHK inhibitor	Decreases blood glucose by activation of pyruvate dehydrogenase (PDH) related to carbohydrate metabolism.	Phase1(Overseas)	In-house
JTK-351	HIV infection /Oral	HIV integrase inhibitor	Suppresses blood HIV levels by inhibiting the activity of integrase, an enzyme involved in the replication of HIV.	Phase1(Japan)	In-house

Clinical trial phase presented above is based on the first dose.

<Licensed compounds>

Compound (JT's code)	Licensee	Mechanism		Note
trametinib	Novartis	MEK inhibitor	Inhibits cellular growth by specifically inhibiting the activity of MAPK/ERK Kinase (MEK1/2).	<u>Melanoma</u> Japan marketing approval submitted
Anti-ICOS monoclonal antibody	MedImmune	ICOS antagonist	Suppresses overactive immune response via inhibition of ICOS which regulates activation of T cells.	
JTE-052	LEO Pharma	JAK inhibitor	Suppresses overactive immune response via inhibition of Janus kinase (JAK) related to immune signal.	

Updates since the previous announcement on November 4, 2015:

<Licensed compounds>

- Gilead Sciences announced that an anti-HIV single-tablet regimen Genvoya (elvitegravir/cobicistat/emtricitabine/tenofovir alafenamide) has been approved by the U.S. FDA and the European Commission. (November 5 and 23, 2015)