



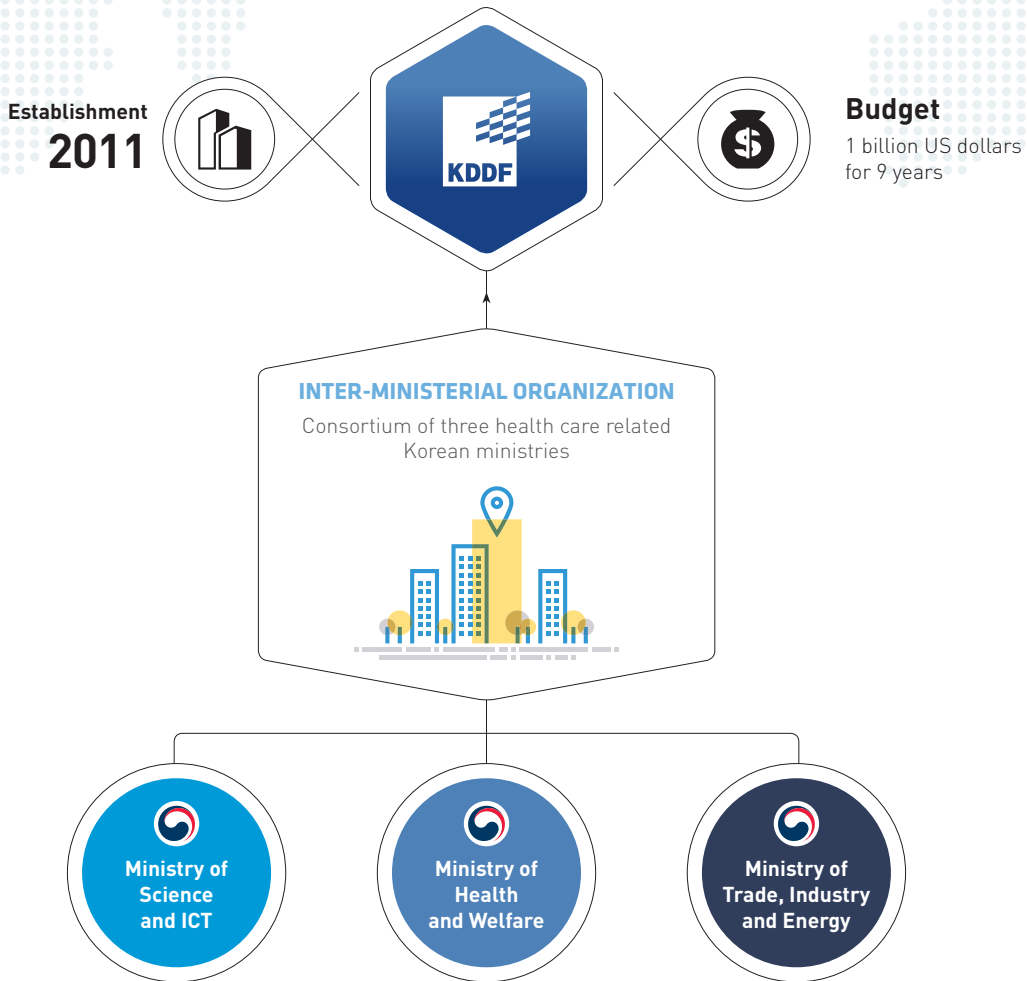
KOREA DRUG DEVELOPMENT FUND

ABOUT KDDF

Korea Drug Development Fund (KDDF) is a government funded organization with one billion USD budget over nine years period of time to accelerate innovation activities in Korean pharmaceutical R&D communities.

KDDF has

- Top-notch proposal selection system
- Value focused project management system
- Large pool of excellent domestic and foreign experts in various drug development field
- International and domestic network in business development field
- More than 136 pipeline in various therapeutic areas from lead stage to clinical trial stage

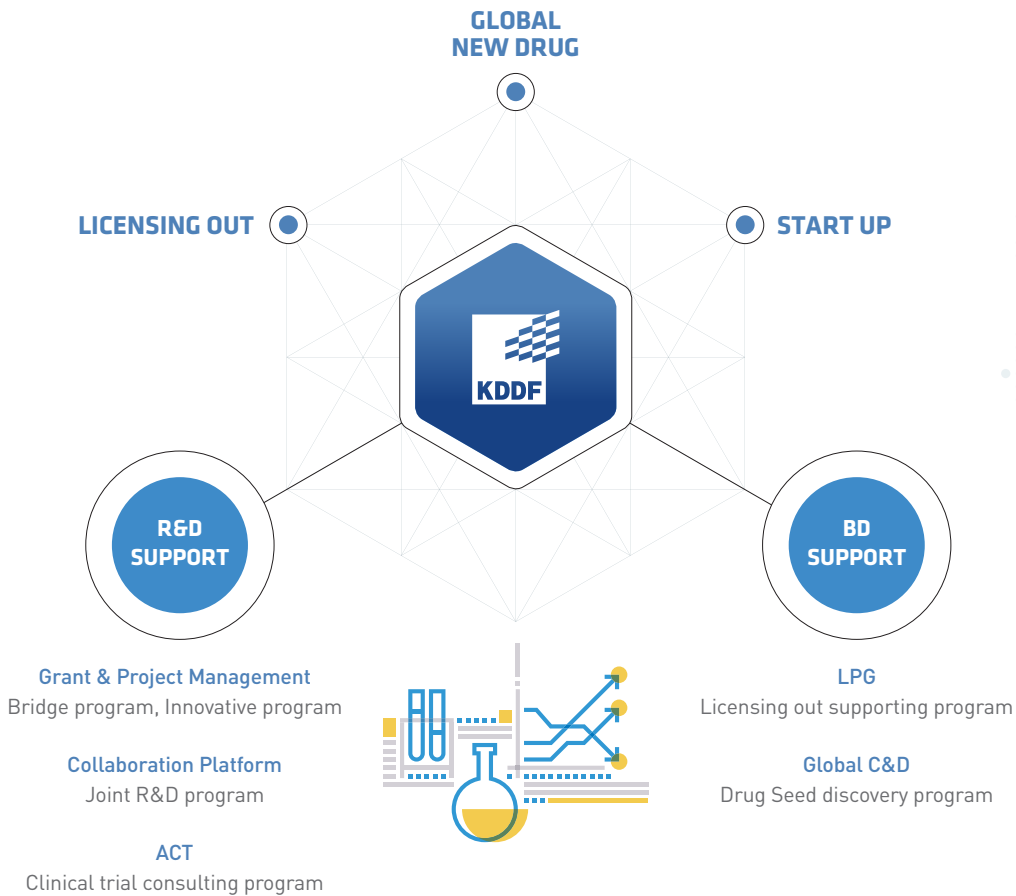


BUSINESS MODEL

KDDF selects and manages portfolio in alignment with Global Big Pharmaceutical companies' interests who are capable of commercializing the asset globally.

What we do

KDDF funds and manages the innovative drug development projects from lead to clinical stage, covering academia, research institute, biotech, and pharmaceutical companies. KDDF does not only fund drug development project, but also supports business development, including out-licensing of the funded projects and in-sourcing drug seeds from abroad.

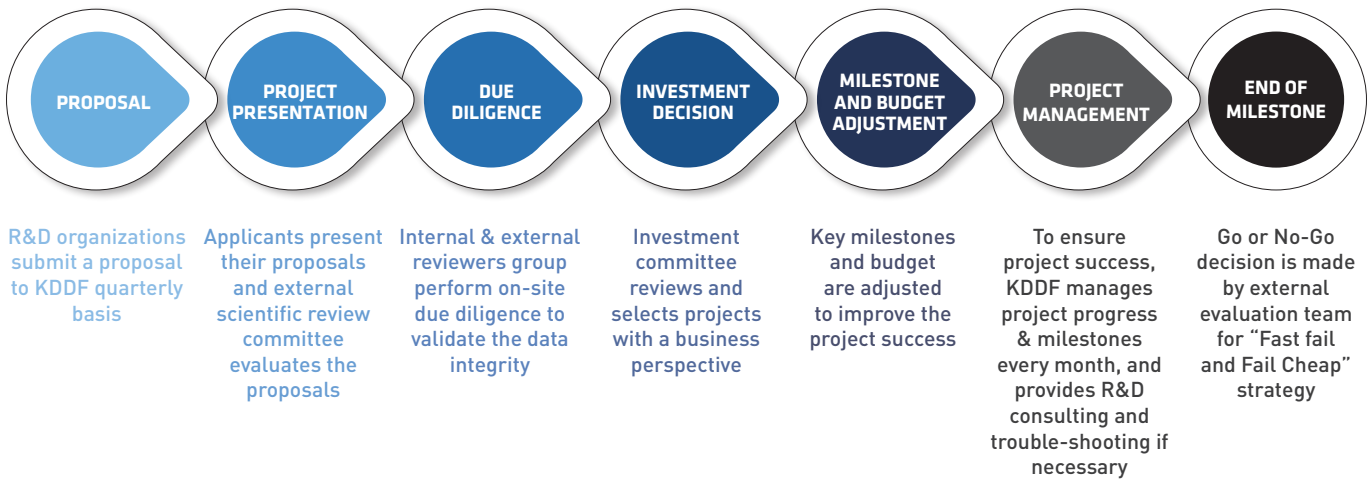


R&D PROGRAM

KDDF has three R&D tracks and one consulting program.

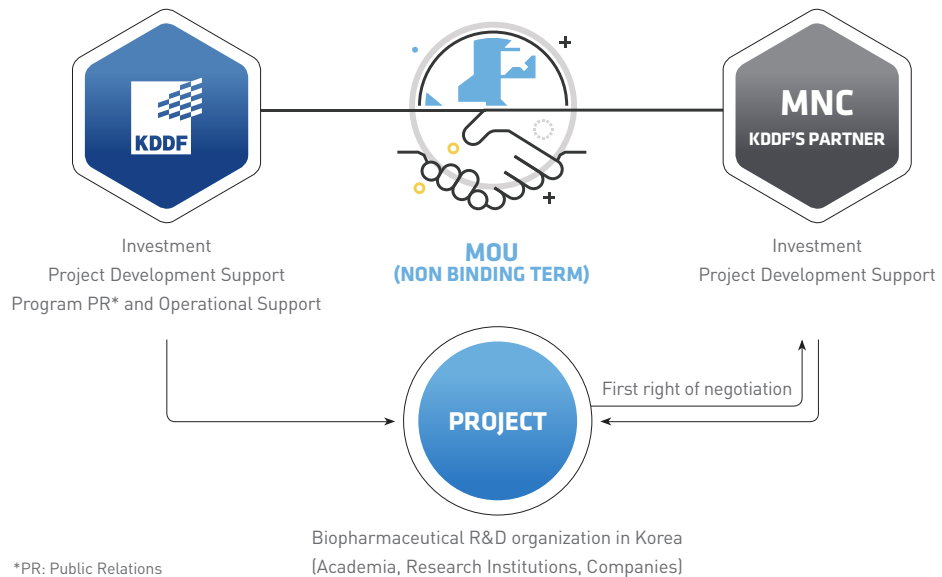
Innovative Track

Innovative track is a bottom up model to find and support novel and innovative drug candidates in Korea. KDDF calls for proposals to Korean drug development communities and selects program through science and investment committee.



Joint R&D Track

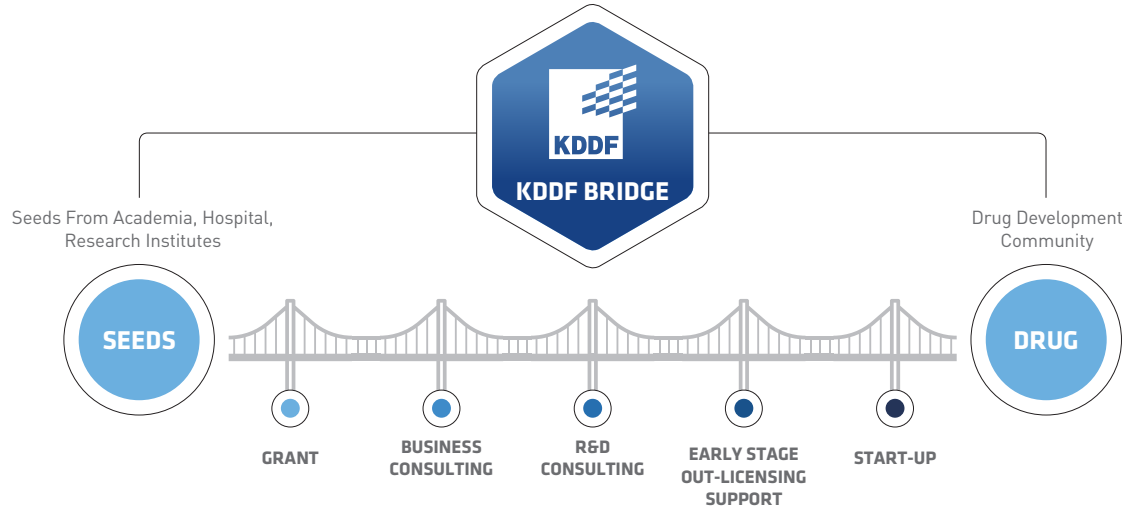
Joint R&D track is a top-down collaboration model, partnered with Global Big Pharmaceutical companies. Call for proposal of Joint R&D Track specifically focuses on certain disease area and mode of action requested by the partner. Through this program the partner is able to find projects to enrich their pipeline.



*PR: Public Relations

BRIDGE Track

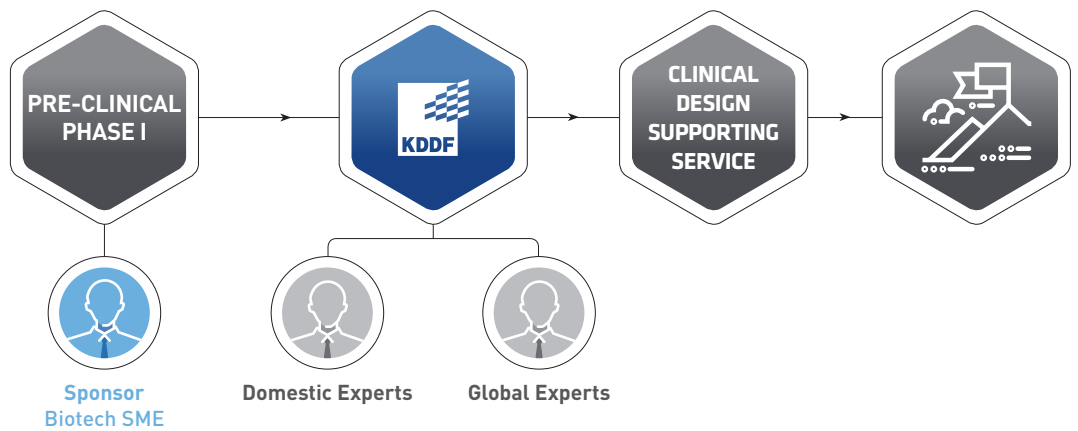
BRIDGE Track is specially designed to bridge very early stage drug discovery to clinical development. In BRIDGE Track, KDDF will co-manage the funded project from the very start to the commercialization, by providing not only grant but also proficient development consulting services, utilizing all the public drug development resources.



ACT

(Advancing Clinical Trial)

- ACT program offers significant benefits for biotech and pharmaceutical companies looking to reduce clinical attribution rate and to increase success rate for their clinical program.
- In-depth clinical trial protocol design consulting service provided by top class clinical experts.
- Advising on study design including appropriate endpoints, statistical methods for analysis, sample size calculations, surrogate biomarkers.



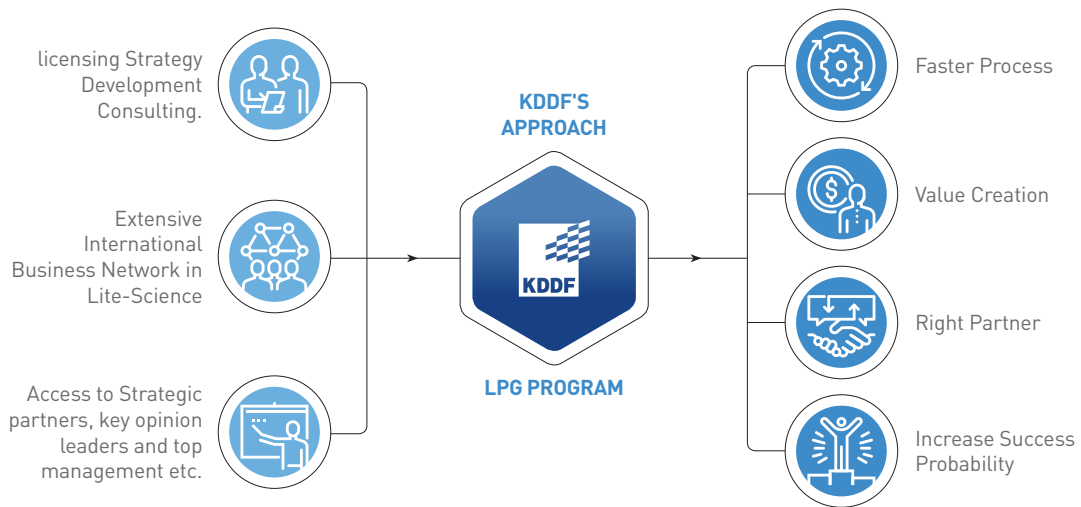
BUSINESS SUPPORTING PROGRAM

KDDF does not only work as a granting agency for new drug development organizations, but also supports their business development through LPG program and Global C&D program.

LPG

(Licensing Partnering for Globalization)

- Business development consulting services for Korean drug development communities.
- Acceleration of licensing activities via KDDF's global network, with consulting service covering all business areas necessary, including from evaluation of licensing value to finding out fittest licensee.



Global C&D

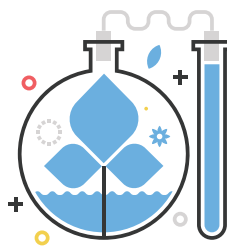
C&D program is to improve R&D productivity by providing open innovation platform to Korean drug development communities, facilitating their in-sourcing novel drug seeds from worldwide.

Pipeline Sourcing



If your seed has a potential, KDDF is able to connect the seed to appropriate development organizations

Farming



Korean Pharmaceutical industry incubate and develops the seed
Biotech, Research Institutes, Pharma Companies, Research Grant (KDDF)

Harvesting



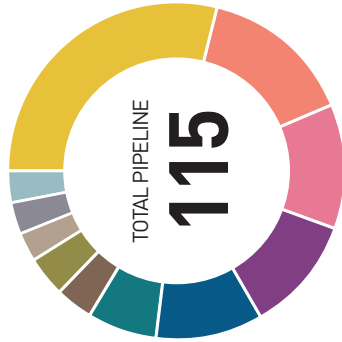
Value added and Global Drug Project
Reverse licensing, Value creation, IPO, M&A, etc.

R&D PIPELINE

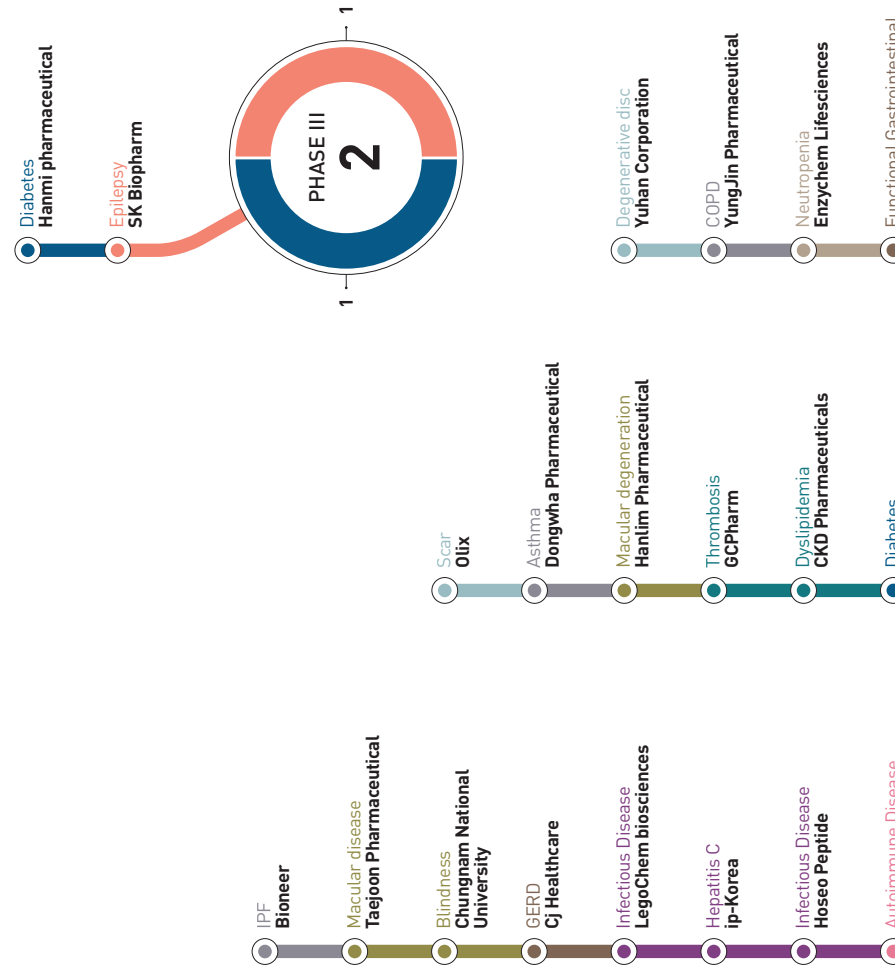
R&D PIPELINE

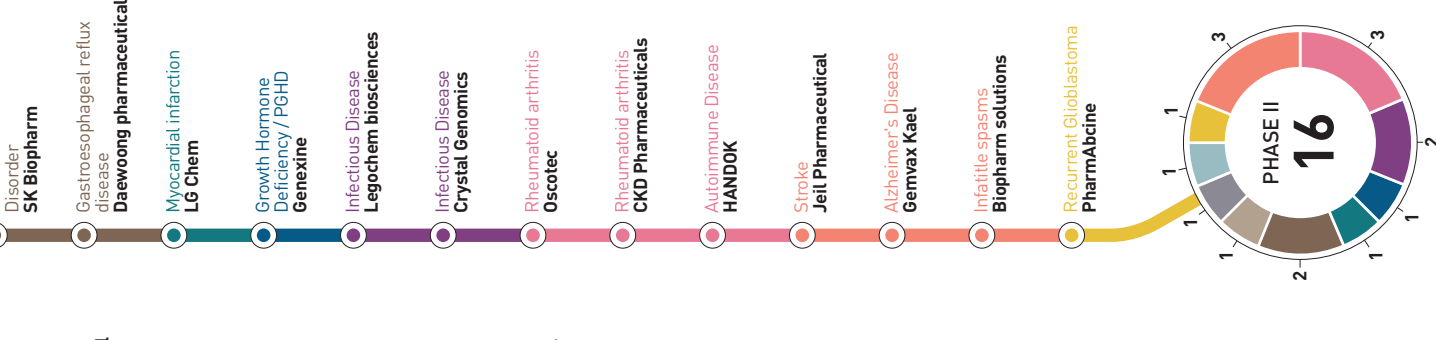
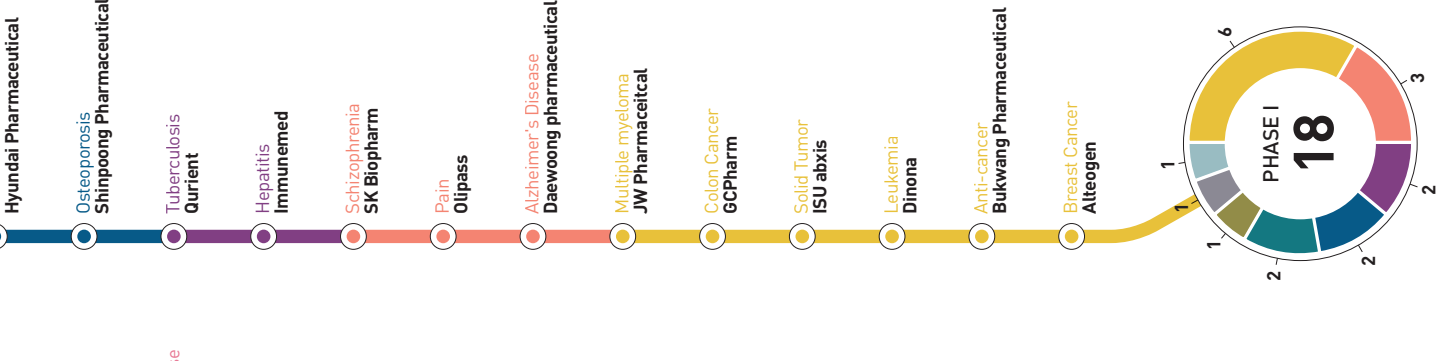
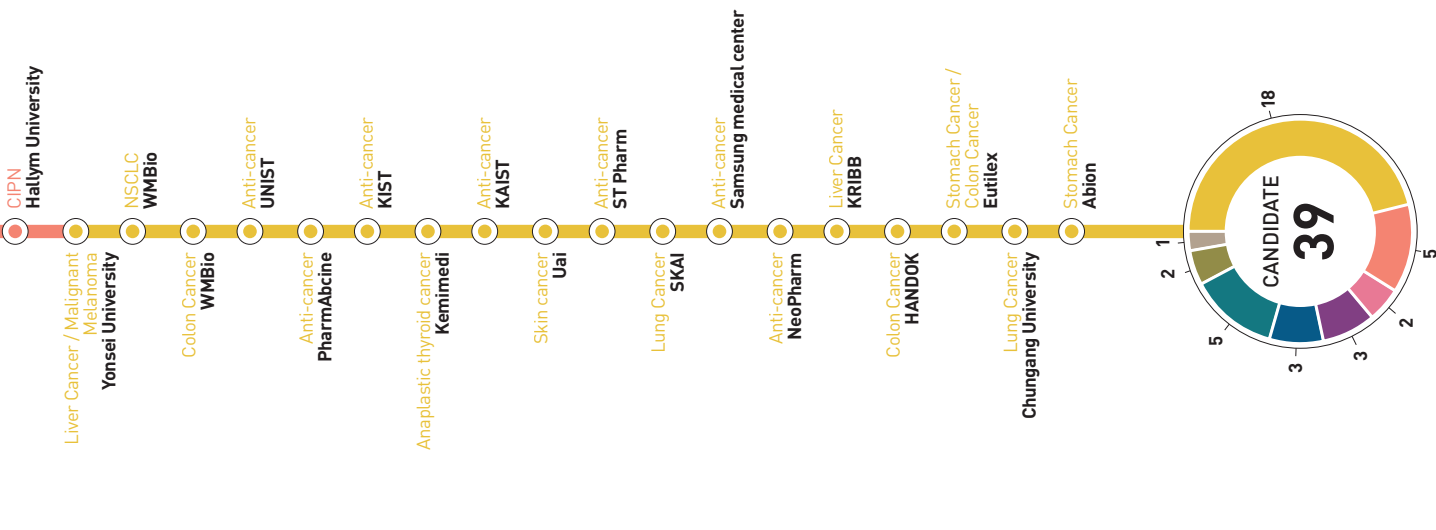
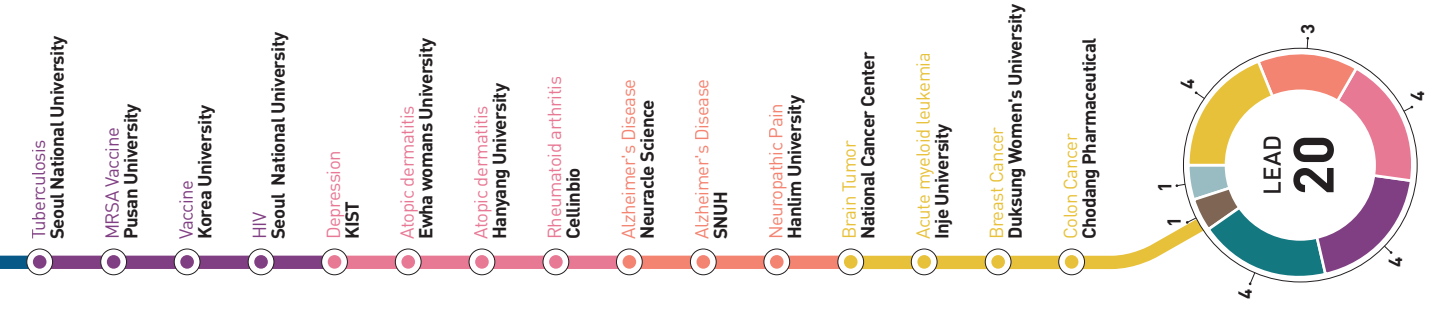
KDDF covers broad range of drug development fields.

For more information, please contact us.
 - <http://eng.kddf.org>
 - kddf@kddf.org



Duchenne Muscular Dystrophy	Korea University
NASH	Asan Medical Center
Diabetes	Ewha womans University
Diabetes	GIST
Osteoporosis	Yonsei University
Diabetes	Medytox





ACCOMPLISHMENT

KDDF has been supporting new drug development throughout all value chains from discovery to clinical trials. As of April 2018, 27 pipeline entered into licensing agreement, of which deal value is totaled more than 4.5 billion USD.



Global Licensing out

Genexine, Cj Healthcare, Hanmi Pharm, LecoChem BIO, AbClon, hanAll-biopharma, Queient, PharmAbcine, GCPharma, Dong-A ST, Dinona

US-FDA ODD*

Qurient, LegoChem BIO, PharmAbcine, BioPharmSolutions, Enzychem Lifescience

* ODD: Orphan Drug Designation

CONTENTS

CANCER

- 12 Unist
- 13 Yonsei University
- 14 WMBio
- 15 Kemimedi
- 16 KAIST
- 17 PharmAbcine
- 18 WMBio
- 19 KIST
- 20 SKAI
- 21 Kwangwoon University
- 22 AbClon
- 23 Hanyang University
- 24 CKD Pharmaceuticals
- 25 Abclon
- 26 Cellbion
- 27 Alteogen
- 28 ISU abxis
- 30 JW Pharmaceutcal
- 31 GCPharm
- 32 PharmAbcine

CNS DISEASE

- 33 Neuracle Science
- 34 KIST
- 35 ABL bio
- 36 ADEL
- 37 Donga ST
- 38 Olipass
- 39 SK Biopharm
- 40 Jeil Pharmaceutical
- 41 BiopharmSolutions
- 42 Gemvax Kael
- 43 SK Biopharm

IMMUNE DISEASE

- 44 Ewha womans University
- 45 Sookmyung women's Univeristy
- 46 Y-Biologics
- 47 JW Pharmaceutcal
- 48 CKD Pharmaceuticals
- 49 Oscotec

INFECTIOUS DISEASE

- 50 Pusan University
- 51 Seoul National University
- 52 Korea University
- 53 Konkuk University
- 54 Ip-Korea
- 55 Immunemed
- 56 Qurient
- 57 Legochem biosciences
- 58 Legochem biosciences
- 59 Crystal Genomics

METABOLIC DISEASE

- 60 Ewha womans University
- 61 GIST
- 62 Aptamer Science
- 63 Hyundai Pharmaceutical
- 64 Shinpoong Pharmaceutical
- 65 Genexine
- 66 Hanmi pharmaceutical

CARDIOVASCULAR DISEASE

- 67 Shinpoong Pharmaceutical
- 68 KRICT

GASTROINTESTINAL DISEASE

- 69 Asan Medical Center
- 70 Daewoong pharmaceutical

OPHTHALMIC DISEASE

- 71 HANDOK
- 72 Chungnam national university
- 73 Taejoon Pharmaceutical

HEMATOLOGIC DISEASE

- 74 Alteogen
- 75 Enzychem Lifesciences

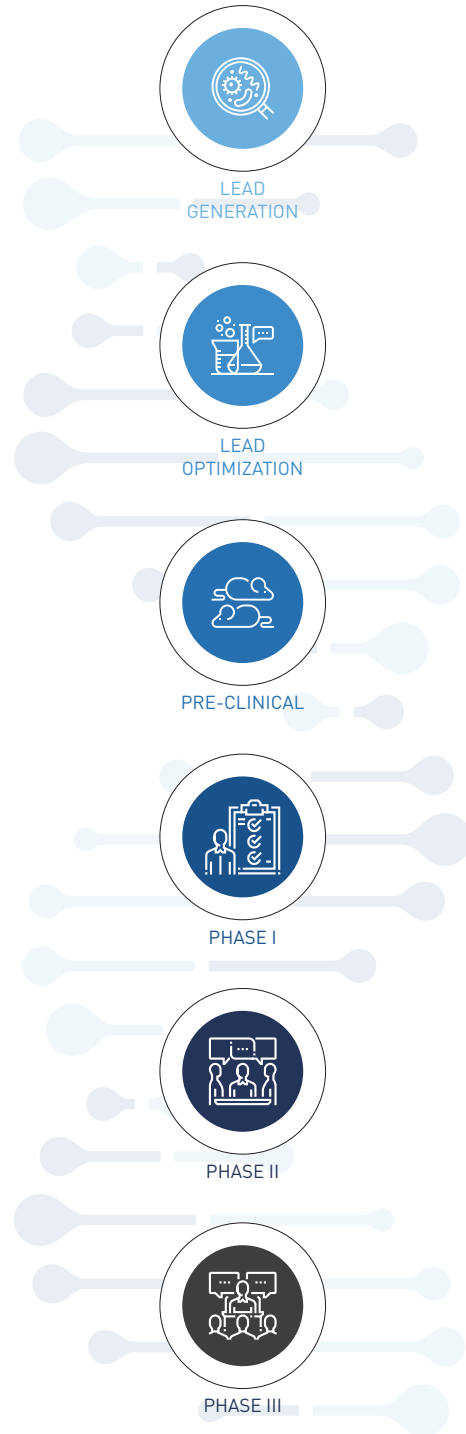
RESPIRATORY DISEASE

- 76 Bioneer
- 77 Dongwha Pharmaceutical

OTHERS

- 78 Olix

From Discovery to Clinical stage
From Academia, research institutes, to Biotech and Pharmaceutical companies



Development of cancer therapeutics targeting mitochondrial TRAP1

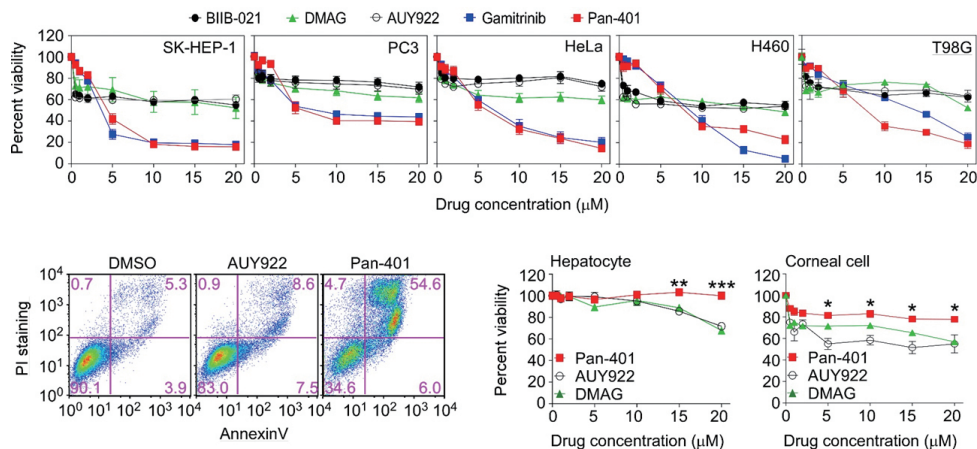
12

KOREA DRUG DEVELOPMENT FUND

Ulsan National Institute of Science and Technology



Product Type	Chemical Product
Indication	1st indication: Prostate cancer, Prostatic Neoplasms (MeSH term) 2nd indication: Brain tumor, Brain Neoplasms (MeSH term)
Target	TNF Receptor associated Protein 1 (TRAP1)
MoA (Mechanism of Action)	Mitochondria permeable TRAP1 inhibitor DN401 accumulates inside the mitochondria and inhibits tumor-supporting chaperone activities of TRAP1.
Differentiation Point	First In Class The subcellular distribution of the drug has been optimized to efficiently inhibit the mitochondrial target TRAP1.
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous
Data	DN401: A purine derivative, intramitochondrial accumulation without the mitochondrial delivery moiety. More potent anticancer activity compared with conventional HSP90 inhibitors.



Patent Position PCT/KR2017/007907

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YSC-02: adenovirus-based anticancer drug with enhanced cancer cell killing and immune activation

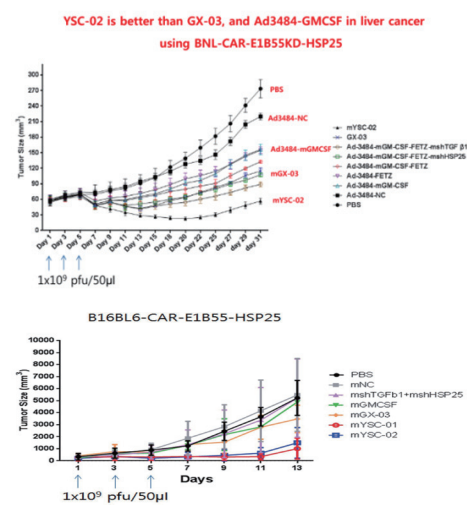
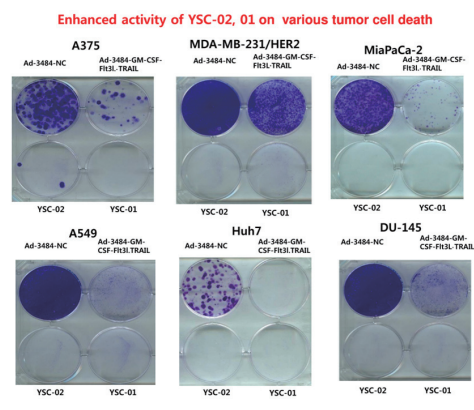
Yonsei University



13
KOREA DRUG DEVELOPMENT FUND

Product Type	shRNAs, cytokines
Indication	1st indication: HCC, Neoplasms (MeSH term) 2nd indication: TNBC, Neoplasms (MeSH term)
Target	TGF-beta, HSP27
MoA (Mechanism of Action)	tumor lysis by oncolytic virus, anti tumor immunity/apoptosis
Differentiation Point	Best In Class
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous

Data



Patent Position 10-2016-0166171/Korea, PCT/KR2016/014325 (10-2016-0166171) and PCT (PCT/KR2016/014325)

Development of a novel small molecule inhibitor for treatment of Erbitux resistant colon cancer

14

KOREA DRUG DEVELOPMENT FUND

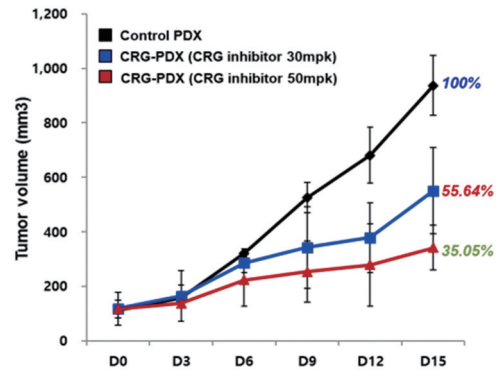
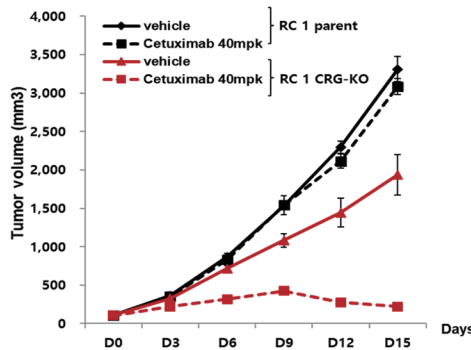
Wellmarkerbio Co., Ltd.



WMBIO

Product Type	Chemical Product
Indication	1st indication: Colon Cancer, Colonic Neoplasms (MeSH term) 2nd indication: Breast Cancer, Breast Neoplasms (MeSH term)
Target	Undisclosed
MoA (Mechanism of Action)	Binding to CRG (Cetuximab-Resistant Gene)
Differentiation Point	First In Class Predictive biomarker for treatment of colon cancer Overcoming resistance of Cetuximab in treatment of colon cancer
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Oral

Data Target validation by CRG-Knock out animal model CRG activated patients' derived colon cancer mouse model shows strong anti-cancer effect of WM-S1-001



2018 R&D Pipeline



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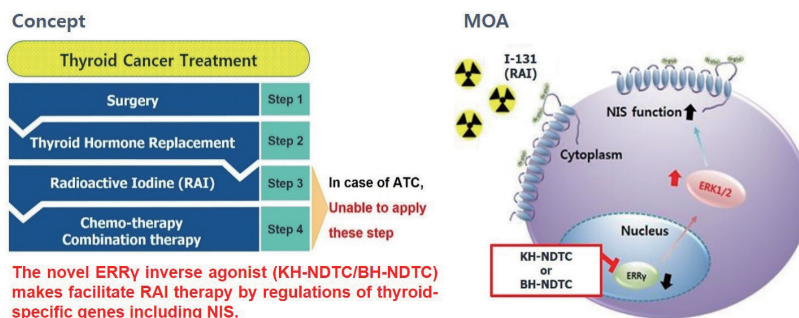
Verify candidate for pre-clinical study in Anaplastic Thyroid Cancer (ATC)

KEMIMEDI Co., Ltd.



15
KOREA DRUG DEVELOPMENT FUND

Product Type Chemical Product
Indication 1st indication: Anaplastic Thyroid Cancer, Thyroid Carcinoma, Anaplastic (MeSH term)
Target Estrogen-related receptor gamma (ERRγ)
MoA (Mechanism of Action)

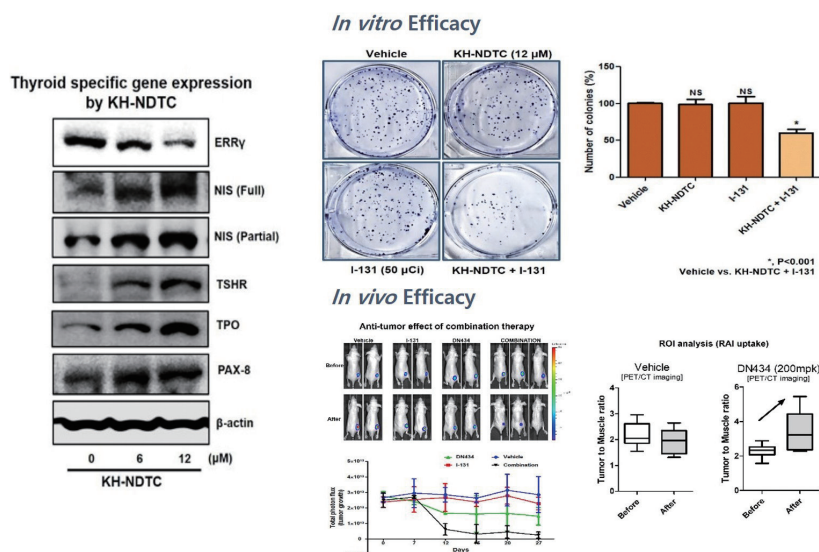


Differentiation Point It will be possible to expand indications to other types of thyroid cancer and to decrease the side effects of radioiodine by reduction of its dose.

Current Development Stage Lead Optimization (Lead to Candidate)

Route of Administration Undecided

Data



Patent Position 10-1819639 / Korea, 10-1835133 / Korea

2018 R60 Pipeline

A novel RNA oligonucleotide developed for cancer therapy

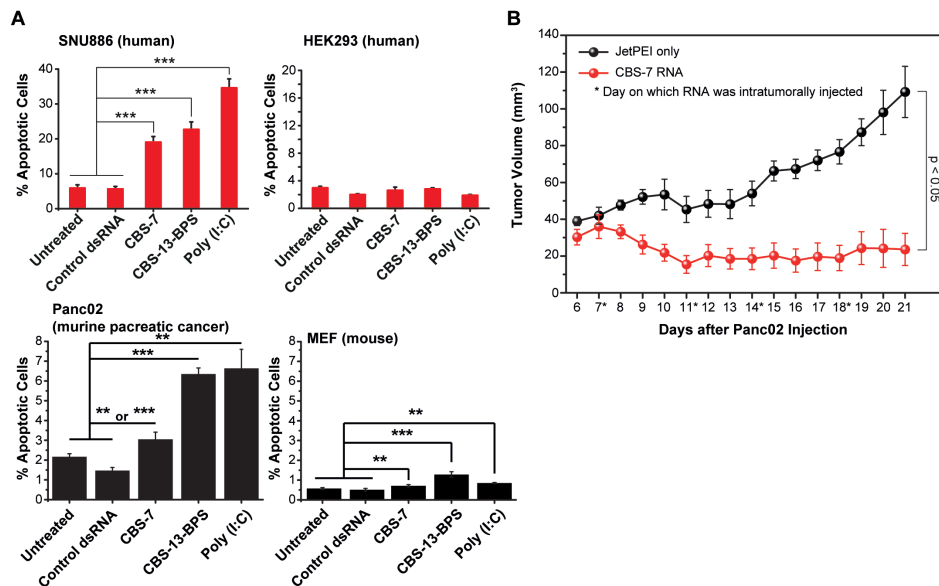
16

KOREA DRUG DEVELOPMENT FUND

Korea Advanced Institute of Science and Technology (KAIST)



Product Type	Aptamer (RNA oligonucleotide)
Indication	1st indication: Cancer, Neoplasms (MeSH term)
Target	Retinoic acid-inducible gene I (RIG-I) (putative)
MoA (Mechanism of Action)	Developed a novel RNA oligonucleotide that specifically induces apoptosis of a tumor
Differentiation Point	First In Class A systematically designed RNA oligonucleotide that shows comparable anti-tumor efficacy to poly I:C
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Parenteral-Intratumoral
Data	<ul style="list-style-type: none"> - Developed RNA oligonucleotide (CBS RNA) is 5'-OH dsRNA. It shows selective anti-tumor efficacy in human liver cancer (SNU886) and mouse pancreatic cancer cells (panc02). (Fig. A) - CBS RNA showed anti-tumor activity against mouse pancreatic cancer (panc02) in vivo (Fig. B). RNA was administered via intratumoral injection.



Patent Position PCT/KR2016/011571

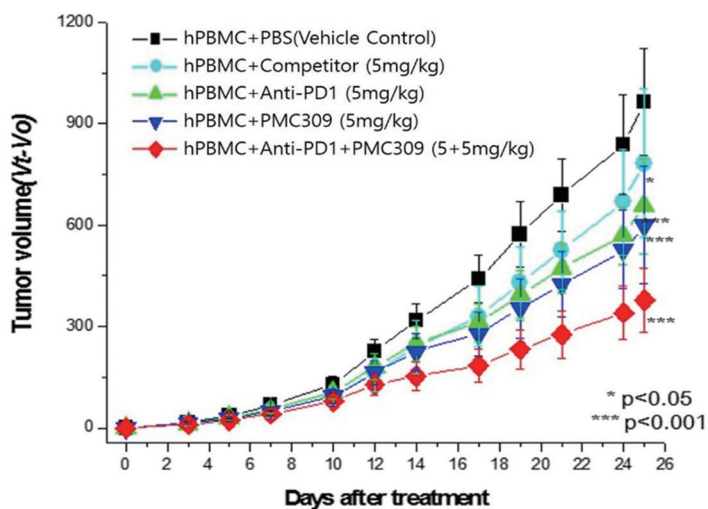
2018 R&D Pipeline

Development of immuno-oncology drug PMC-309 targeting VISTA known as a negative checkpoint regulator

PharmAbcine



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: NSCLC, carcinoma, non-small cell lung (MeSH term) 2nd indication: breast cancer, Breast Neoplasms (MeSH term)
Target	VISTA (v-domain Ig suppressor of T-cell activation)
MoA (Mechanism of Action)	Increase immunity against tumor by binding to VISTA
Differentiation Point	Best In Class fully human, best immune activation activity
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous
Data	- stronger anti-tumor activity than competitor in humanized mouse model - synergic anti-tumor activity with anti-PD1 antibody



Development of new mechanism based therapeutic antibodies in non-small cell lung cancer patients

18

KOREA DRUG DEVELOPMENT FUND

Wellmarkerbio Co., Ltd.



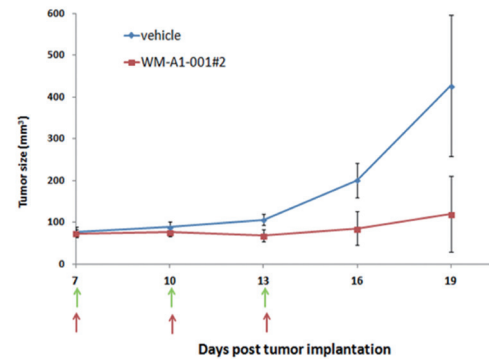
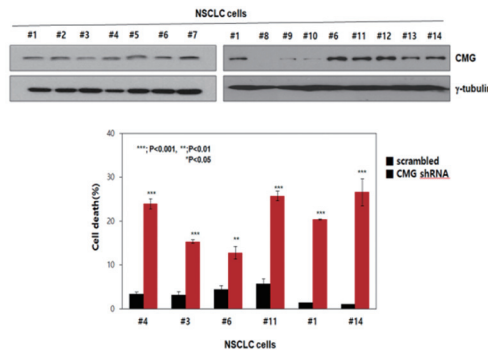
WMBIO

Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Lung cancer, Lung Neoplasms (MeSH term) 2nd indication: Liver cancer, Liver Neoplasms (MeSH term)
Target	Undisclosed
MoA (Mechanism of Action)	Binding to CMG (Cancer Immunotherapy-related gene)
Differentiation Point	First In Class
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intraperitoneal injection (IP)

Data

Target validation
: knockdown of CMG

Syngeneic model:
in vivo efficacy for WM-A1-001#2

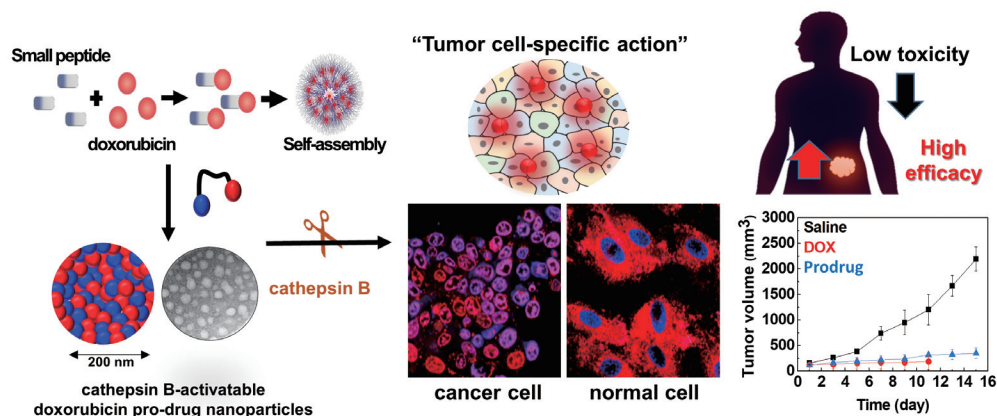


Development of nano-sized anti-cancer prodrug capable of tumor enzyme (cathepsin B)-specific activation

Korea Institute of Science and Technology (KIST)



Product Type	Peptide Product (A self-assembly tumor specific drug conjugate)
Indication	1st indication: Colorectal cancer, Colorectal neoplasms (MeSH term) 2nd indication: Breast cancer, Breast neoplasms (MeSH term) 3rd Indication: Pancreatic cancer, Pancreatic neoplasms (MeSH term)
Target	DNA in tumor cells (doxorubicin based)
MoA (Mechanism of Action)	1. Self-assembly small peptide-based nanoparticles without any nanocarriers 2. Passive accumulation effect in the tumor site 3. Tumor cell specific activation by cathepsin B (biomarker) 4. Doxorubicin-based strong cytotoxic effect on cancer cells
Differentiation Point	First In Class This is the first low molecular weight drug candidate based on self-assembly and cathepsin B-specific action among doxorubicin prodrugs
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous
Data	1. Long term administration due to low toxicity by prodrug-type 2. Delivery to tumor tissue by the passive accumulation effect of nanoparticles 3. Simple synthesis and predicted efficacy with a clear biomarker 4. Having the advantages of cytotoxic and target anticancer agents



Patent Position

KR 10-2017-0121169

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A novel anti-angiogenic human monoclonal antibody specifically targeting CLEC14a

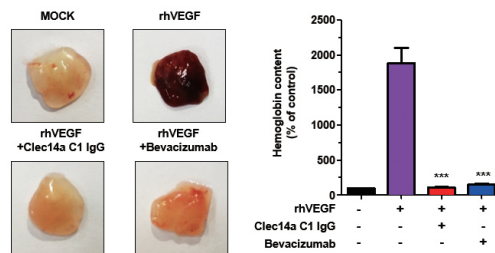
20

KOREA DRUG DEVELOPMENT FUND

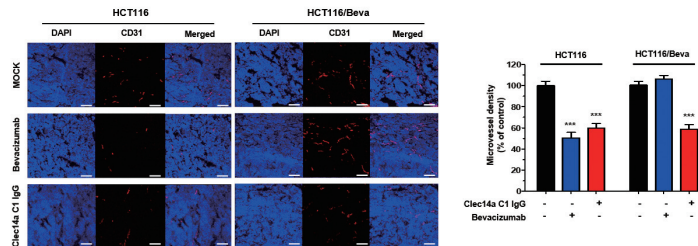
Scripps Korea Antibody Institute



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Cancer, Neoplasms (MeSH term) 2nd indication: Wet age-related macular degeneration, Macular Degeneration (MeSH term)
Target	CLEC14a (Tumor endothelial marker)
MoA (Mechanism of Action)	1) Internalization-dependent down-regulation of CLEC14a on the surface of endothelial cells 2) Inhibition of CLEC14a-CLEC14a protein interactions → Inhibition of endothelial cell-cell contacts → Inhibition of angiogenesis
Differentiation Point	First In Class This mAb has lower toxicity and anti-angiogenic efficacy similar to Avastin
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Intravenous
Data	1) Inhibition of VEGF-dependent and tumor angiogenesis



2) Inhibition of tumor angiogenesis induced by Avastin-resistant tumor



Patent Position 9,751,933/USA, 6129305/Japan, 10-1760465-0000/Korea, 201380043426.9/China, 10-2016-0115577/PCT

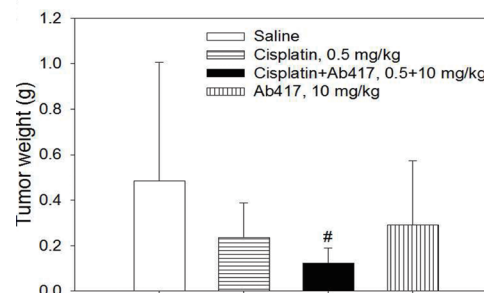
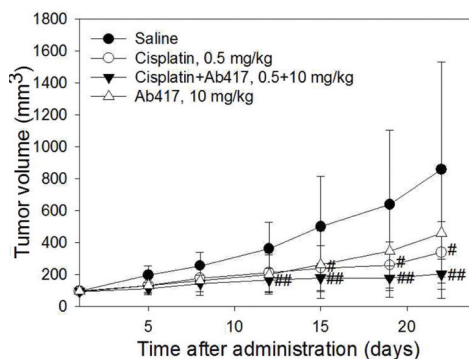
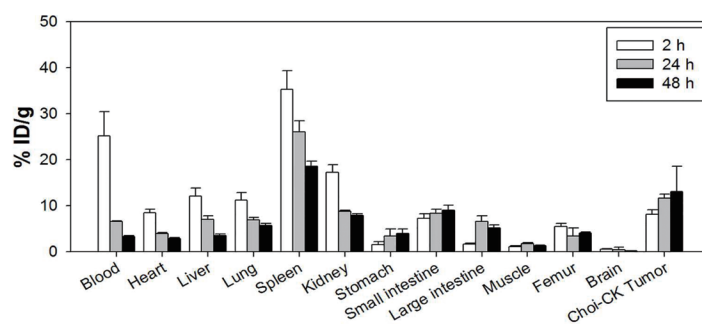
2018 R&D Pipeline

Preliminary non-clinical study of a therapeutic antibody candidate for the treatment of cholangiocarcinomas

Kangwon National University



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Cancers, Neoplasms (MeSH term) 2nd indication: Cholangiocarcinomas, Cholangiocarcinomas (MeSH term)
Target	L1CAM
MoA (Mechanism of Action)	Inhibition of tumor growth by internalization of L1CAM in the cells and down-regulation of membrane L1CAM
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Intravenous
Data	: No adverse effect in single-dose toxicity study in mice. Cross-reactive with mouse L1CAM



Patent Position 10-2007-0084868/ Korea; 10-2013-0140237/Korea; 10-2018-0000381/Korea

A Novel Therapeutic Antibody targeting HER2

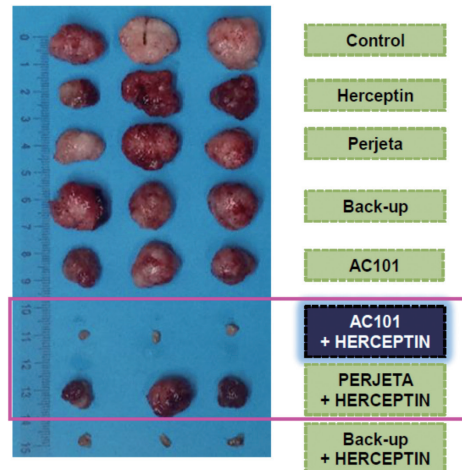
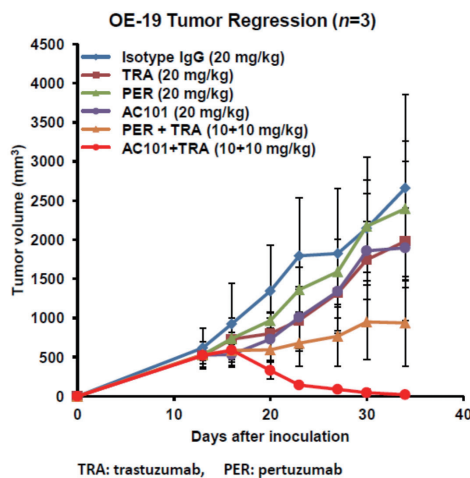
22

KOREA DRUG DEVELOPMENT FUND

AbClon Inc.



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Gastric Cancer, Stomach Neoplasms (MeSH term) 2nd indication: Breast Cancer, Breast Neoplasms (MeSH term)
Target	Human epidermal growth factor receptor 2 (HER2)
MoA (Mechanism of Action)	The drug shows superior efficacy in combination with trastuzumab against HER2(+) cancers. Synergistic effect of the drug with trastuzumab due to increased apoptosis, increased cell cycle arrest, inhibition of both homo- & hetero- dimerization effect.
Differentiation Point	Best In Class The drug can overcome limitation of Herceptin mono-therapy. The anti-cancer activity of the drug is superior to the combination of Herceptin and Perjeta when the drug is combined with Herceptin.
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Intravenous
Data	<ul style="list-style-type: none"> ▪ Combination treatment of AC101 with trastuzumab (Herceptin®, TRA) in HER2 positive gastric cancer cell (OE-19) xenografted mice shows superior tumor regression efficacy to the monotherapy of trastuzumab, pertuzumab (Perjeta®, PER) or the combination of the both (PER + TRA).



Patent Position PCT/KR2014/004317

2018 R&D Pipeline

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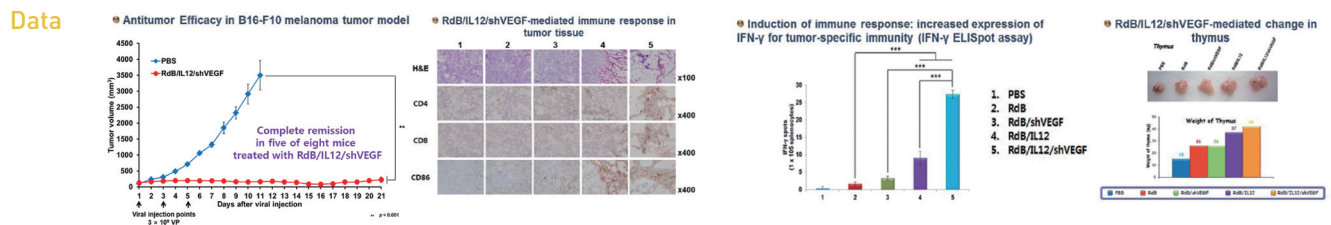
(+82)2-2109-1366

Development of gene therapeutic-mediated combination therapy regimen targeting lung cancer

Hanyang University, Gene Therapy Lab.



Product Type	Oncolytic adenovirus
Indication	1st indication: Lung cancer 2nd indication: Breast cancer, Prostate cancer
Target	Hypoxia, VEGF, Immune surveillance
MoA (Mechanism of Action)	<ul style="list-style-type: none"> • Oncolytic adenovirus kill cancer cell specifically and multiply therapeutic transgene shVEGF and IL-12. • Novel promoter enhances induces cancer-specific killing effect • shVEGF inhibits angiogenesis, metastasis and proliferation of cancer cell. • shVEGF reverses immunosuppressive tumor microenvironment. • IL-12 decreases the tumor metastasis by regulating tumor microenvironment. • IL-12 increases infiltration of CD4+ T, CD8+ T, natural killer, and dendritic cells in tumors tissues • IL-12 upregulates IFN-γ to shift the T cell response toward the Th1 immunity, ultimately enhancing anti-tumor immune response.
Differentiation Point	<ul style="list-style-type: none"> • Targeting multiple carcinogenic pathway through novel strategy • No observable cytotoxicity or side effect in normal cells while preferentially treating tumor cells • Significantly lower production cost in comparison with competing product • Synergistic antitumor efficacy in combination with standard therapies (chemotherapy, radiotherapy, antibody, immune cells)
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Intratumoral



Patent Position

- PCT/KR2016/009717 (IL12/shVEGF)
- PCT/KR2004/000427 (mTERT)
- PCT/KR2011/004693 (E2F+TERT)

Pre-Clinical studies of a novel anti-cancer bispecific antibody CKD-702

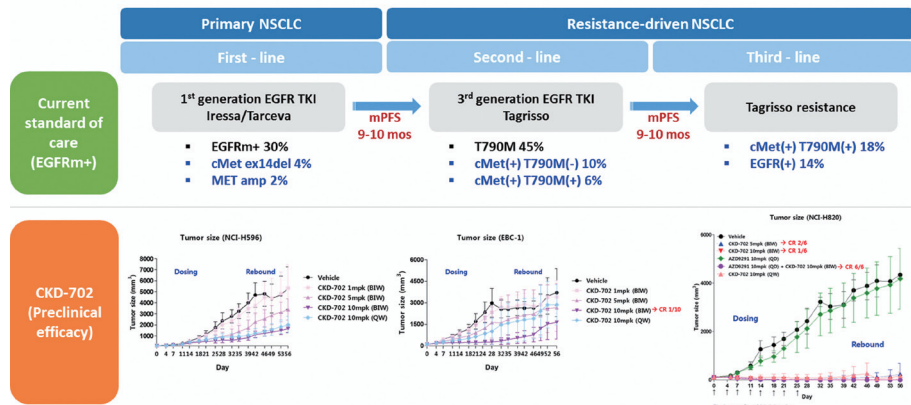
24

KOREA DRUG DEVELOPMENT FUND

Chong Kun Dang Pharmaceutical Corporation



Product Type	Immunoglobulin Product (mAb), bispecific antibody
Indication	1st indication: Non-small cell lung cancer, Carcinoma, Non-Small-Cell Lung (MeSH term) 2nd indication: Gastric cancer, Glioblastoma, Stomach Neoplasms, Glioblastoma (MeSH term)
Target	Mesenchymal to Epithelial Transition factor receptor (cMET) or Hepatocyte growth factor receptor (HGFR) and Epidermal growth factor receptor (EGFR)
MoA (Mechanism of Action)	CKD-702 binds and internalizes both cMET and EGFR → Degrades the receptors and effectively blocks downstream signaling pathways → Superior antitumor activity
Differentiation Point	Best In Class Superior antitumor activity in multiple NSCLC, colon and gastric cell lines or NSCLC tumor xenograft models. Treatment of cynomolgus monkeys with CKD-702 resulted in a low toxicity profiles, including skin rash observed with other EGFR-directed agents.
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Intravenous
Data	CKD-702 opportunities in NSCLC: - In first-line therapy, patients with EGFR wild type and cMET ex14 deletion - In second and third line treatment, patients with cMET amplification and acquired resistance to EGFR tyrosine kinase inhibitors



Patent Position 10-2017-0067106 (Korea)

EunJu Jeon

ejjeon@ckdpharm.com

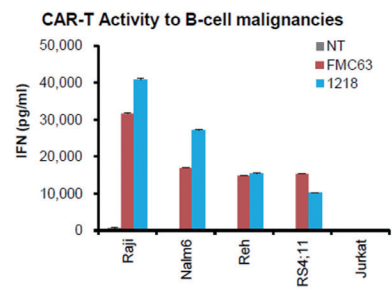
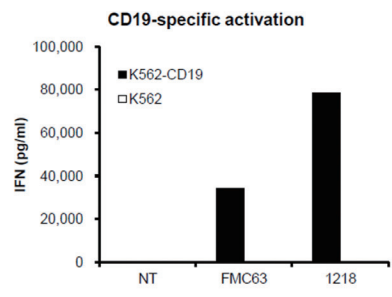
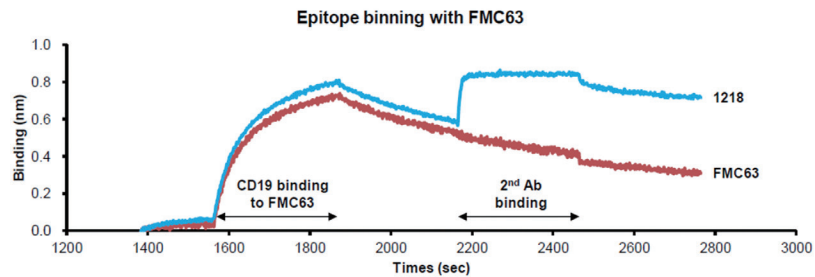
(+82)31-340-1421

Development of CD19-targeting CAR-T therapeutics

AbClon Inc.



Product Type	Cell Therapy
Indication	1st indication: Acute Lymphoid Leukemia, Precursor Cell Lymphoblastic Leukemia-Lymphoma (MeSH Term) 2nd indication: B-cell lymphoma, Lymphoma, B-Cell (MeSH term)
Target	B-lymphocyte antigen CD19 (CD19)
MoA (Mechanism of Action)	Favorable formation of immune synapse of the CAR-T using new CD19 antibody with novel epitope can increase the efficacy whereas it can reduce adverse effects.
Differentiation Point	The drug can overcome limitations such as resistance and immunogenicity of marked CD19 targeting CAR-T on the basis of novel epitope.
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravascular
Data	<ul style="list-style-type: none"> ▪ New CAR-T is constructed using novel CD19 scFv (1218). ▪ The drug has different epitope compared to marked CAR-T using chimeric scFv (FMC63). ▪ The drug has increased activity compared to FMC63 CAR-T.



Patent Position 10-2017-0166969/KR

Development of a PSMA(Prostate Specific membrane antigen)-targetting therapeutic radiopharmaceutical

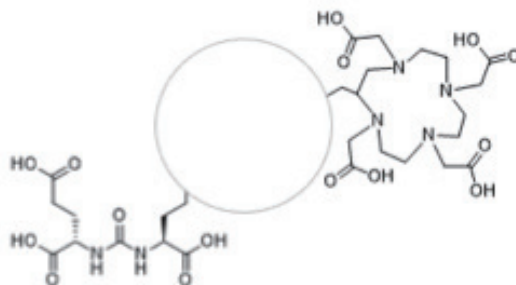
26

KOREA DRUG DEVELOPMENT FUND

Cellbion Co. Ltd.

CellBion

Product Type	Peptide Product
Indication	1st indication: Prostate cancer, Prostatic Neoplasms (MeSH term) 2nd indication: Metastatic cancer, Prostatic Neoplasms, Castration-Resistant (MeSH term)
Target	Prostate Specific Membrane Antigen (PSMA)
MoA (Mechanism of Action)	Prostate-specific membrane antigen (PSMA), also known as folate hydrolase I (FOLH1) or glutamate carboxypeptidase II (GCP II), is a transmembrane, 750 amino acid, type II glycoprotein that is primarily expressed in normal human prostate epithelium but is overexpressed in prostate cancer, including metastatic disease. Lu-177 has a relatively long physical half-life of 6.73 days. It is these physical properties that allow for the delivery of high activities of Lu-177 PSMA to prostate cancer cells.
Differentiation Point	Best In Class This compound is composed of Glutamate-Urea-Lysine moiety known as a pharmacophore for PSMA inhibitors, and a chelating agent for a radioisotope connected by a linker. This linker is very distinctive to make this molecule stable and safe due to its simple chemical structure compared to those of other products. This reference-based data is being investigated by performing some experiments to confirm the advantage of our chemical.
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Intravenous
Data	Structure of PSMA-GUL-DOTA



PSMA-GUL-DOTA(therapeutic)

- α -Therapy(^{225}Ac -labelled)
- β -Therapy(^{177}Lu -labelled)

Patent Position PCT/KR2016/012849

TaeRahk Kim

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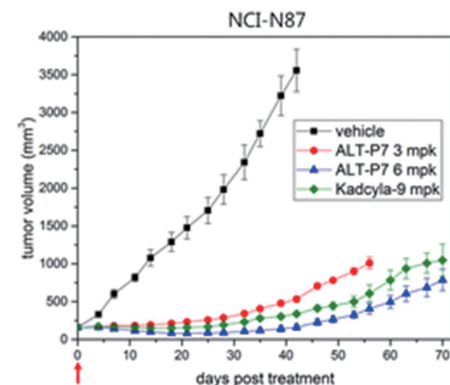
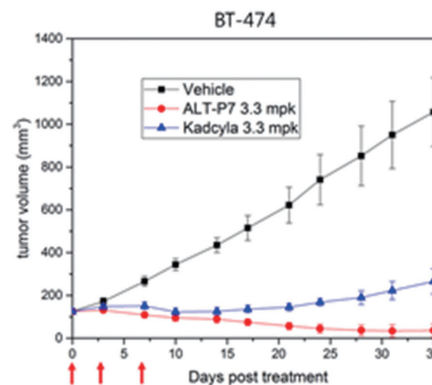
[+82]2-743-3311

Phase I clinical study of HER2-positive breast cancer by an antibody-drug conjugate ALT-P7

Alteogen, Inc.



Product Type	Immunoglobulin Product (mAb) Antibody-drug conjugate (ADC)
Indication	1st indication: Her2-positive breast cancer, Breast neoplasms (MeSH term) 2nd indication: Her2-positive gastric cancer, Stomach neoplasms (MeSH term)
Target	Human epidermal growth factor receptor 2 (Her2)
MoA (Mechanism of Action)	Antibody-based Her2-positive cancer cell targeting followed by a payload-dependent cancer cell death
Differentiation Point	Best In Class -Drug conjugation at a specific site of the antibody allows ALT-P7 a structural stability and low in vivo toxicity, compared to non-specifically conjugated ADC. -Cleavable linker permits ALT-P7 the bystander killing effect and shows superior efficacy to non-cleavable linker-adopted ADC in Herceptin-resistant in vivo models.
Current Development Stage	Phase I
Route of Administration	Parenteral-Intravenous
Data	<ol style="list-style-type: none"> Superior in vivo efficacy in mouse xenograft studies. <ul style="list-style-type: none"> - Higher in vivo efficacy and a complete regression of tumor in BT-474 breast carcinoma xenograft model. - Better in vivo efficacy even in lower dose in NCI-N87 gastric carcinoma xenograft model. Phase I study is underway for the safety, tolerability, pharmacokinetics, and effective dose determination on a first-in-human setting.



ISU104, A fully human IgG1 monoclonal antibody blocking human ErbB3

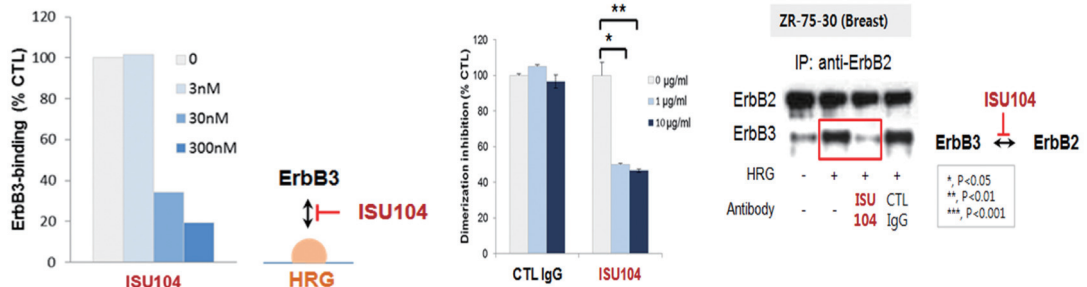
28

KOREA DRUG DEVELOPMENT FUND

ISU ABXIS Co., Ltd



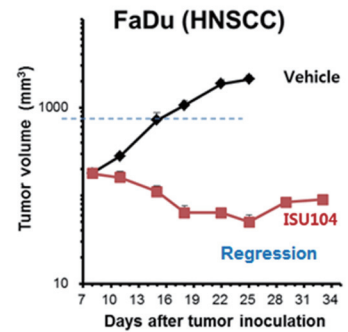
Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Head and Neck Cancer, Head and Neck Neoplasms (MeSH term) 2nd indication: Breast Cancer, Breast Neoplasms (MeSH term)
Target	Human epidermal growth factor receptor 3 (ErbB3)
MoA (Mechanism of Action)	Anti-cancer mAb via blocking ErbB3 signaling
Differentiation Point	Best In Class - Dual blocker (ligand binding & receptor dimerization) - Tumor growth regression (TGR) in HNSCC, →50% tumor growth inhibition (TGI) (→10 sc/orthotopic xenograft & syngeneic models). - Confirmed ErbB3 upregulation by Erbitux-resistance (Bypass MOA for SOC resistance) - TGR (Erbitux-resistant xenograft model) & stronger TGI by various combination w/ SOC (HNSCC)
Current Development Stage	Phase I
Route of Administration	Parenteral-Intravenous
Patent Position	WO2017-099362 A1 / PCT, KR 10-2017-0067637 / Korea, KR 10-2017-0067638 / Korea
Data	ISU104 is a specific dual blocker for ligand binding and ErbB3 dimerization <ul style="list-style-type: none"> • Ligand Blocker • Dimerization Blocker



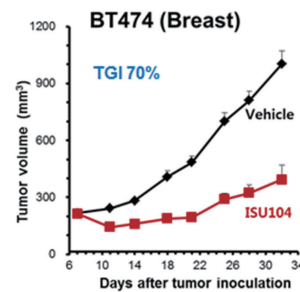
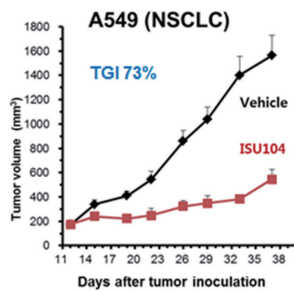
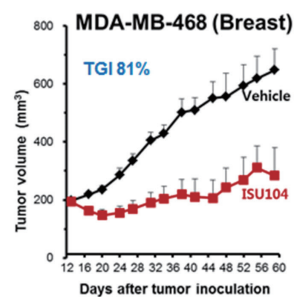
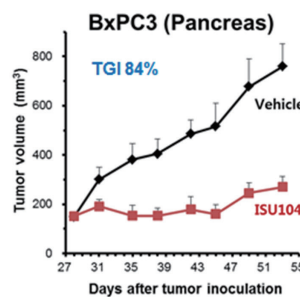
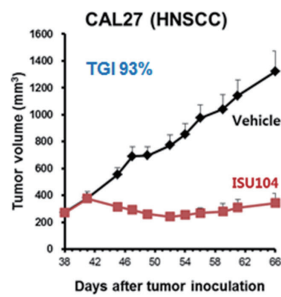
Data

Efficacy in vivo : Tumor growth in mice

- Tumor growth regression (TGR) in HNSCC

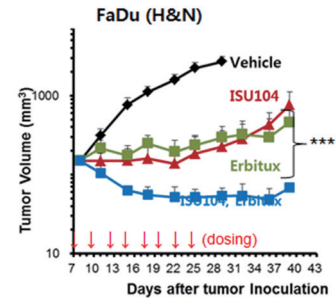
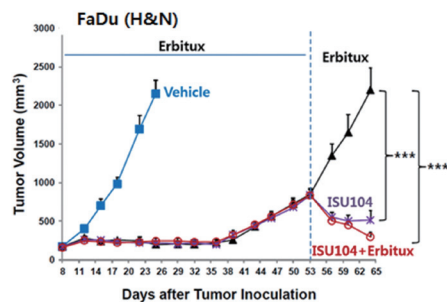


- Tumor growth inhibitions (TGI) in various xenografts & syngeneic models



Resistance Overcome: Tumver growth & Cell proliferation

- TGR in Erbitux-resistant H&N cancer model
- TGR by combination of ISU104 w/ Erbitux



CWP232291

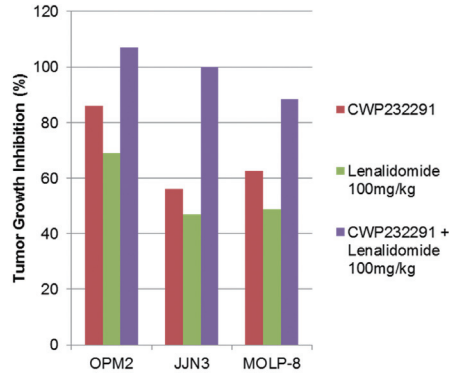
First-in-class Wnt signaling pathway inhibitor

30
KOREA DRUG DEVELOPMENT FUND

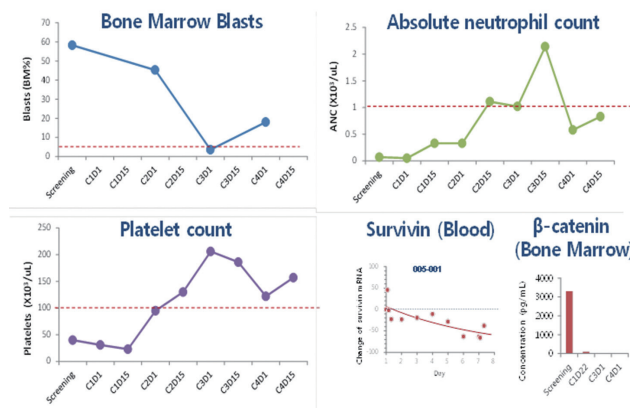
JW Pharmaceutical



Product Type	Chemical Product
Indication	1st indication: Relapsed/Refractory Multiple Myeloma, Multiple Myeloma (Mesh term) 2nd indication: Relapsed/Refractory Acute Myeloid Leukemia, Leukemia, Myeloid, Acute (MeSH term)
Target	Wnt signaling pathway
MoA (Mechanism of Action)	Inhibit Wnt signaling pathway by disrupting the unfolded protein response and endoplasmic reticulum stress, resulting in tumor-selective apoptosis
Differentiation Point	CWP232291 showed single agent efficacy and good candidate for combo. with SOC
Current Development Stage	Phase I b
Route of Administration	Parenteral-Intravenous
Data	Synergistic effect with SOC in Multiple Myeloma xenograft model



Single-agent efficacy in RR-AML patients with excellent tolerability



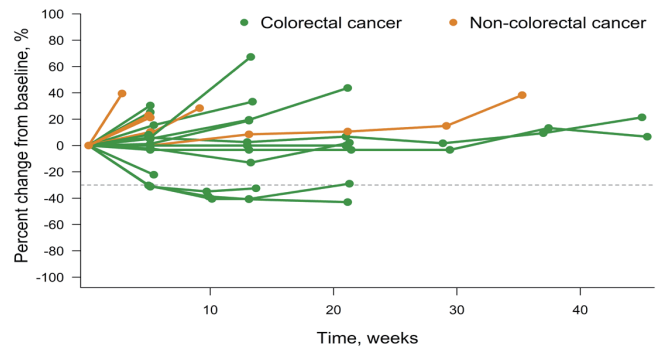
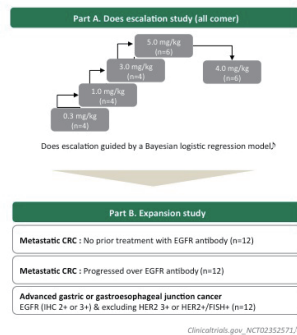
2018 R&D Pipeline

Development of a novel EGFR-targeted antibody

GCPharma



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Colorectal cancer, Colorectal Neoplasms (MeSH term) 2nd indication: Gastric cancer, Stomach Neoplasms (MeSH term)
Target	Epidermal Growth Factor Receptor
MoA (Mechanism of Action)	Binds to EGFR and interrupts the binding of EGFR ligands to EGFR
Differentiation Point	Best In Class Different binding epitope/More efficient inhibition of EGFR ligand binding to EGFR esp. high-affinity ligands
Current Development Stage	Phase I b
Route of Administration	Parenteral-Intravenous
Data	Phase 1 dose escalation study indicated that weekly infusion of GC1118 is tolerable and safety Partial response was observed from dose level of 4 and 5 mg/kg and best disease control rate was 67% In Expansion study 58.3% of DCR was observed in CRC patients with no prior EGFR therapeutics and 1 out of 12 gastric cancer patients showed PR Phase 1b/2a is open to evaluate the safety and efficacy of GC1118 in combination with FOLFIRI or irinotecan



Patent Position

WO2011/040668, WO2013/147509

Clinical Phase IIa Trial with Tanibirumab, a Novel Anti-Cancer Antibody Therapeutics, in Recurrent Glioblastoma Patients

32

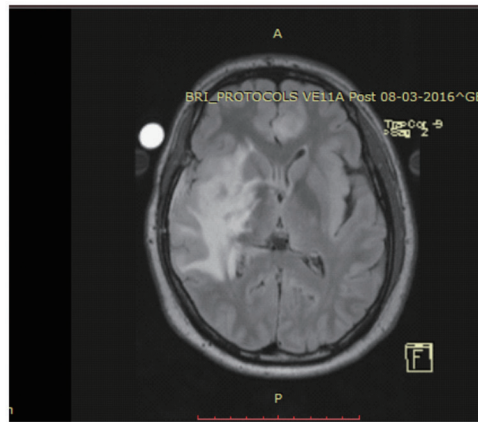
KOREA DRUG DEVELOPMENT FUND

PharmAbcine

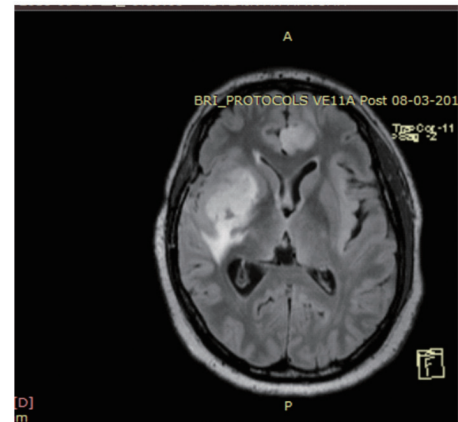


Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: recurrent glioblastoma, glioblastoma (MeSH term) 2nd indication: breast cancer, breast neoplasms (MeSH term)
Target	VEGFR-2
MoA (Mechanism of Action)	Inhibit tumor angiogenesis by blocking VEGFs binding to VEGFR-2
Differentiation Point	Best In Class Fully human, good safety and activity
Current Development Stage	Phase II a
Route of Administration	Parenteral-Intravenous
Data	25 % disease control rate (upto 16 cycles (1.5 years)) in rGBM phase IIa 42 % patients showed relief of edema in rGBM phase IIa (figure) 16.7 % patients showed steroid use reduction in rGBM phase IIa

Before



After



Patent Position WO2008153237

WeonSup Lee

weonsup.lee@pharmabcine.com

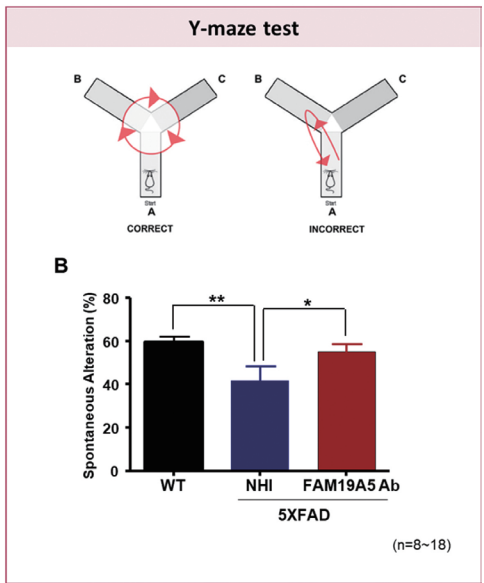
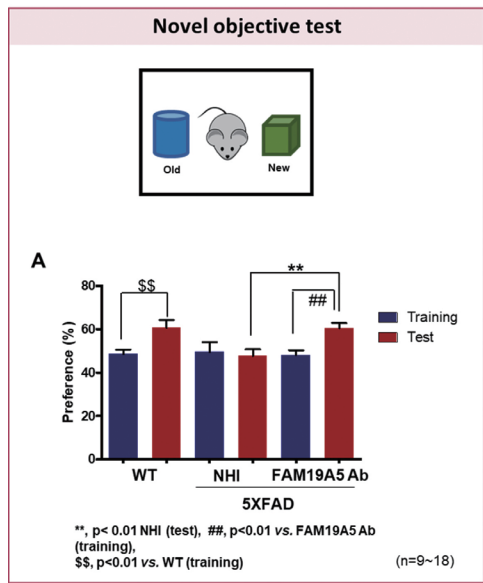
(+82)43-235-2017(ext 3001)

Development of therapeutic antibody for Alzheimer's disease by targeting glial scar formation

Neuracle Science Co., Ltd.



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Alzheimer`s disease, Alzheimer Disease (MeSH term)
Target	Family with similarity 19, member A5 (FAM19A5)
MoA (Mechanism of Action)	Glial scar remodeling and vessel normalization through increased immune activity
Differentiation Point	First In Class
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Parenteral-Intravenous
Data	FAM19A5 antibody efficacy test with Alzheimer`s disease animal model



Patent Position US9579398B2

Novel Anti-depressant by triple reuptake inhibition

34

KOREA DRUG DEVELOPMENT FUND

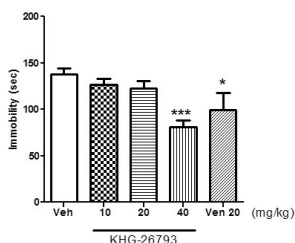
Korea Institute of Science and Technology



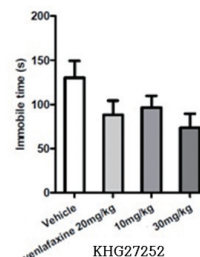
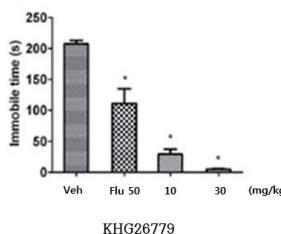
Product Type	Chemical Product
Indication	1st indication: anti-depressant, Nervous System Diseases (MeSH term)
Target	Reuptake inhibition of neurotransmitters
MoA (Mechanism of Action)	Reuptake inhibition of neurotransmitters
Differentiation Point	Reuptake inhibition of three neurotransmitters (dopamine, serotonin, norepinephrine) simultaneously
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Oral
Data	- Excellent reuptake inhibitory activities against three neurotransmitters - hERG, CYP, Microsomal stability, BBB, PK

In vivo

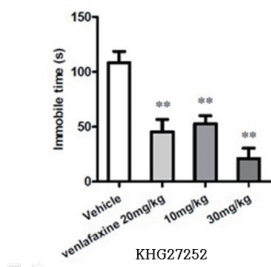
Forced Swimming Test (FST)



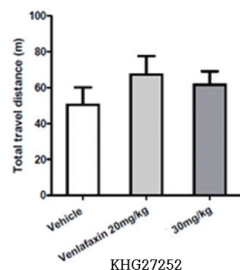
*p<0.05, ***p<0.001 compared with the Veh group.



Tail suspension test (TST)



Locomotion test



Patent Position

PCT/KR2012/007076, KR10-1561992, KR10-1651994

2018 R&D Pipeline

HohGyu Hahn

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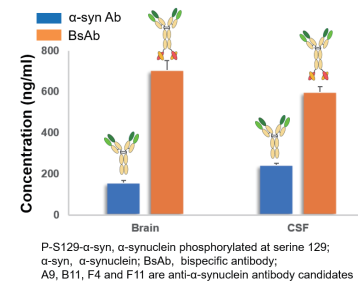
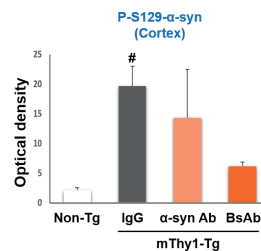
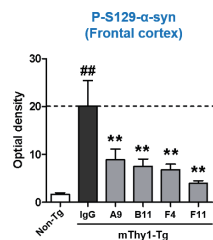
(+82)2-958-5139

Bispecific Antibody Targeting Pathological α -synuclein with Improved Blood-brain Barrier Penetration

ABL Bio



Product Type	Immunoglobulin Product (mAb), Bispecific antibody
Indication	1st indication: Parkinson's Disease, Parkinson Disease (MeSH term) 2nd indication: Multiple Systems Atrophy, Multiple Systems Atrophy (MeSH term)
Target	- Aggregated forms of α -synuclein - A receptor on surface of brain endothelial cells
MoA (Mechanism of Action)	1. Increase α -synuclein clearance, inhibit cell-to-cell transmission of α -synuclein 2. BBB penetration: receptor-mediated transcytosis
Differentiation Point	First In Class The drug is bispecific antibody targeting α -synuclein aggregate with improved BBB penetration
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous
Data	<ul style="list-style-type: none"> • Anti-α-synuclein antibody binds selectively to α-synuclein aggregates in vitro. • Bispecific antibody composed of the anti-α-synuclein antibody above and BBB shuttle enters into rat CSF and brains better than anti-α-synuclein antibody. • Bispecific antibody shows better therapeutic efficacy in animal model of Parkinson's disease by clearing aggregated α-synuclein more than anti-α-synuclein antibody and control IgG.



Tau Antibody development for Alzheimer's Disease

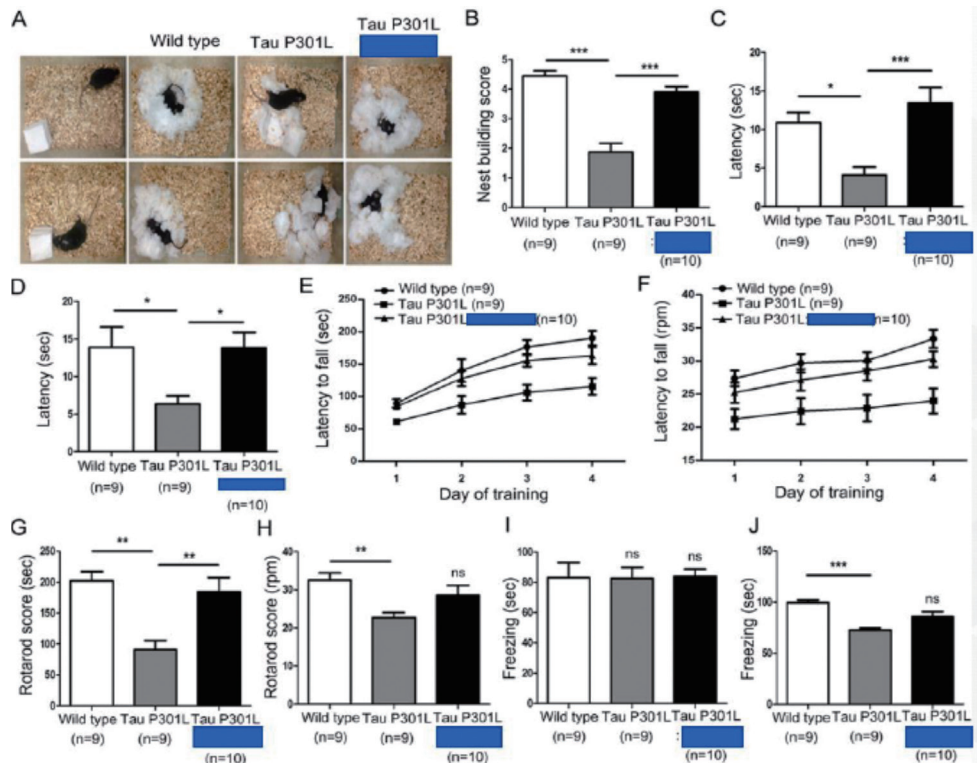
36

KOREA DRUG DEVELOPMENT FUND

ADEL, Inc.



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Alzheimer's disease, Alzheimer Disease (MeSH term) 2nd indication: Tauopathy, Tauopathies (MeSH term)
Target	Tau(specific epitope with post-translational modification)
MoA (Mechanism of Action)	anti-aggregation & anti-propagation of pathologic Tau protein, etc.
Differentiation Point	First therapeutic antibody targeting the specific epitope of Tau
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous
Data	in vivo efficacy of Tau antibody



Patent Position PCT/KR2017/015137

SeungYong Yoon

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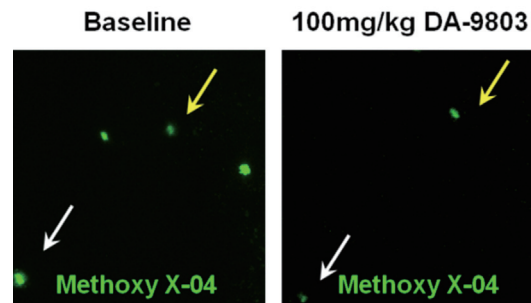
(+82)2-3010-4241

Novel Botanical Drug Development for Alzheimer's disease in the US

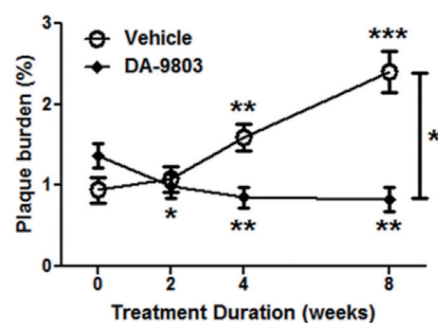
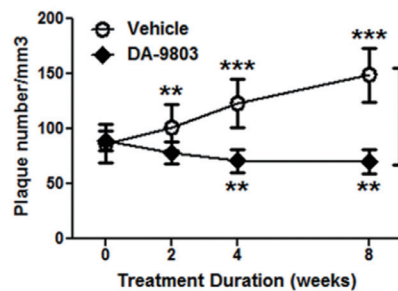
Dong-A ST



Product Type	Botanical drug
Indication	1st indication: Alzheimer disease, Nervous System Diseases (MeSH term)
Target	Multi-Target (A β , Tau, AChE)
MoA (Mechanism of Action)	1) Disease-treating via removal of disease-causing source A β , ptau 2) Improving cognitive ability via AChE inhibition 3) Neuroprotection via NGF
Differentiation Point	Multi-function (Disease modifying and symptomatic effects)
Current Development Stage	Pre-Clinical
Route of Administration	Undecided
Data	Amyloid beta research (Brain of APP/PS1 mouse) (Alzheimers Res Ther. 2018 Jan 29;10(1):11)



Red dextran : Blood vessel labeling reagent
Methoxy X-04 : Fluorescent Amyloid beta



Patent Position

PCT/KR-2015-013134 , PCT/KR2015/013136

Novel Pain Killer SCN9A Antisense Oligonucleotide

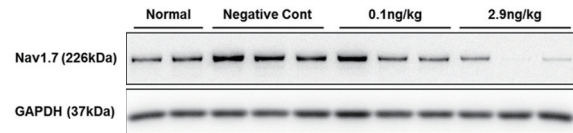
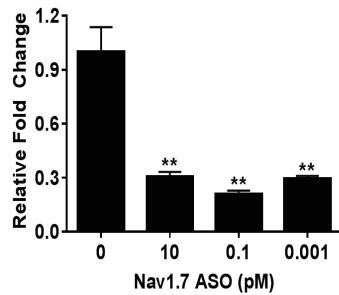
38

KOREA DRUG DEVELOPMENT FUND

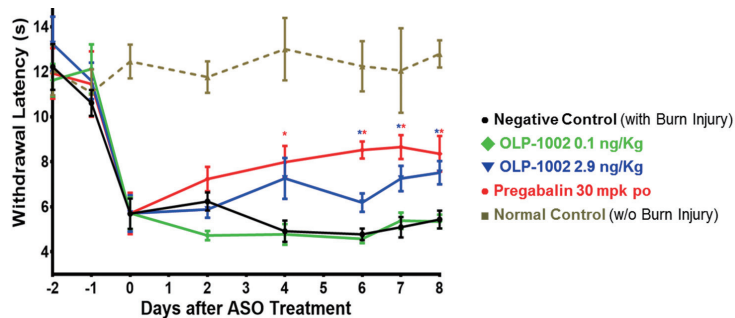
OliPass Corporation



Product Type	RNAi, Antisense oligonucleotide
Indication	1st indication: Suffering, Physical, Pain (MeSH term)
Target	Sodium Voltage-Gated Channel Alpha Subunit 9 / SCN9A
MoA (Mechanism of Action)	OliPass PNA tightly binds to a specific region in pre-mRNA and potently induces exon skipping
Differentiation Point	Improvements in cell permeability will lead to lower doses and better therapeutic ratios
Current Development Stage	Phase I
Route of Administration	Parenteral-Subcutaneous
Data	SCN9A mRNA Level Nav 1.7 expression in DRG (ex Vivo)



In vivo- Burn Model Pain Behavior



Patent Position PCT/KR2009/001256, PCT/WO02009/113828

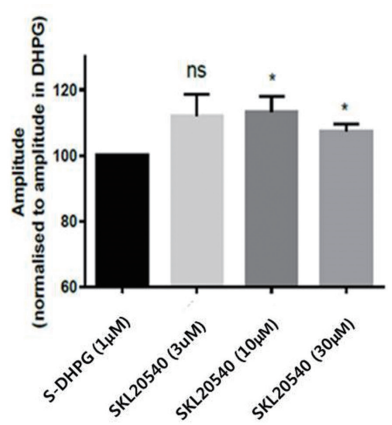
2018 R&D Pipeline

A New Generation of Anti-Psychotics Specialized in Schizophrenia with the improvement of Negative Symptom

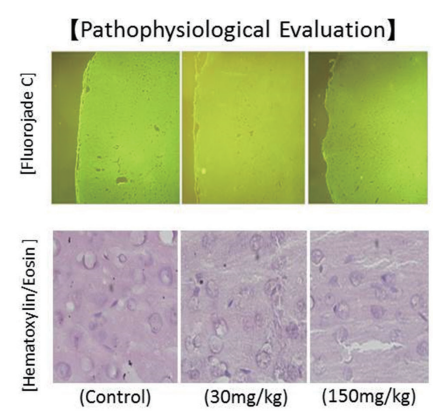
SK biopharmaceuticals



Product Type	Chemical Product
Indication	1st indication: Schizophrenia, Schizophrenia (MeSH term)
Target	Metabotropic glutamate receptor subtype 5 (mGluR5)
MoA (Mechanism of Action)	Positive modulator of mGluR5 by binding the allosteric site thereof without the agonistic character to activate NMDA receptor
Differentiation Point	<p>First In Class</p> <p>The compound is a non dopamined-based drug having the therapeutic effect on the netive symptom as well as the positive symptom and mitigating metabolic side effects caused by the currently marketed drugs</p>
Current Development Stage	Phase I
Route of Administration	Oral
Data	<ul style="list-style-type: none"> - Confirmed that NMDAR-mediated fEPSP at CA region of hippocampus was enhanced - Confirmed that there was no significant toxicities which are related to the target receptor



A paired, one-tailed t-test (S-DHPG(N=3), 3µM(N=5), 10µM(N=7), 30µM(N=3))



Phase 2A Clinical Study of JPI-289 for the Treatment of Stroke

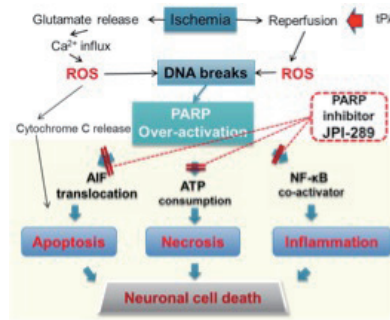
40

KOREA DRUG DEVELOPMENT FUND

JEIL Pharmaceutical Co., Ltd.

JEIL PHARMACEUTICAL CO.,LTD.

Product Type Chemical product
Indication 1st indication: Stroke, Stroke (MeSH term)
Target Poly[ADP-ribose] polymerase-1 (PARP-1)
MoA (Mechanism of Action)

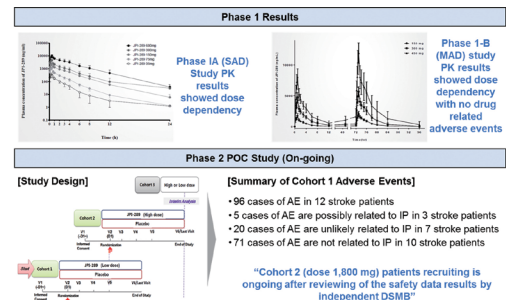
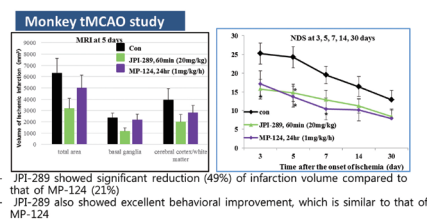


- Differentiation Point**
1. Inhibition of PARP-1 is a significantly distinct mechanism of action and is expected to show high efficacy in clinical trials with ischemic stroke patients through the neuroprotective effects.
 2. In a monkey tMCAO stroke model, JPI-289 showed 49% decrease in infarction volume, which is the best result in the world when compared with that of 21% decrease in infarction volume by MP-124. Therefore, JPI-289 among PARP inhibitors is considered as one of the most promising agents for the treatment of stroke.
 3. Safety of JPI-289 has been confirmed in healthy volunteers and stroke patients because there were no serious adverse events (SAEs) during phase 1 and phase 2A cohort 1 studies.

Current Development Stage Phase II a

Route of Administration Parenteral-Intravenous

Data



Patent Position WO2010056038, WO2013115535

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Pre-Clinical study of new chemical entity for orphan disease: infantile spasms

Bio-Pharm Solutions



Product Type Chemical Product

Indication 1st indication: Infantile Spasms (West Syndrome), Spasms, Infantile (MeSH term)
2nd indication: Status Epilepticus, Status Epilepticus (MeSH term)

Target Metabotropic glutamate receptor family 1 & 3, glycolysis pathway

MoA (Mechanism of Action)

In vitro

- Work as agonist of mGluR 4 & 5 and antagonist of mGluR 1, 5, and 7
- Protects pericytes against tPA-induced cytotoxicity and cell death
- Interacts with phosphoglycerate kinase 1 and pyruvate dehydrogenase E1 alpha in the glycolysis pathway (on-going)

Ex vivo

- Using hippocampal slice on multielectrode array system, JBPOS0101 reduced significantly pilocarpine-induced spontaneous activity and reconfirmed JBPOS0101's agonist for mGluR 4 and antagonist for mGluR 1, and 7

In vivo

- Protects BBB against Li-pilocarpine-induced SE and collagenase-induced intracerebral hemorrhage
- Blocks hippocampal cell death in Li-pilocarpine-induced SE
- JBPOS0101 shows broad spectrum of efficacy in the various conventional nonclinical epilepsy screening results in B-PS & ASP's studies

Bio-Marker

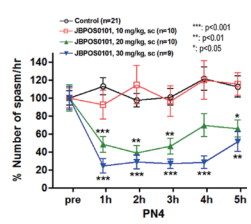
- EEG as a functional Bio-marker to show epilepsy state and efficacy of the stes drug

Differentiation Point Best In Class
The drug is expecting the best efficacy in the pediatric epilepsy.

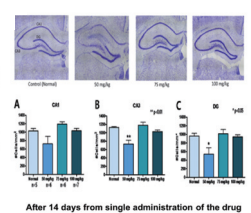
Current Development Stage Phase II

Route of Administration Oral

Data A) The drug shows good suppression of behavioral spasms.



B) The drug protects hippocampal cell death against to Status Epilepticus.



Patent Position 13/175,025 / US, 10-2012-7033745 / KR, 15/190,123 / US

Development of telomerase derived peptide, GV1001 in Alzheimer's disease

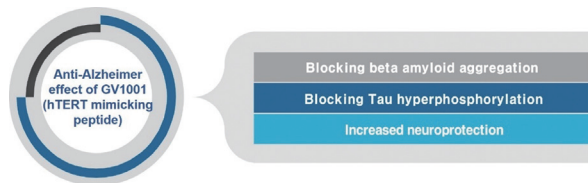
42

KOREA DRUG DEVELOPMENT FUND

GemVax & KAEL Co., Ltd.



Product Type Peptide product
Indication 1st indication: Alzheimer's disease, Alzheimer Disease (MeSH term)
Target Unknown
MoA (Mechanism of Action)

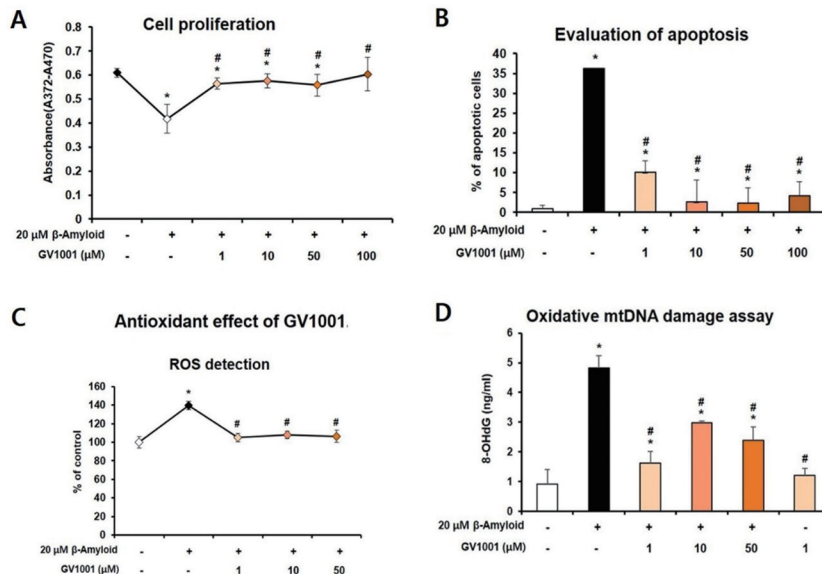


Differentiation Point First In Class
 GV1001 is the first telomerase-derived peptide drug for the treatment of Alzheimer's disease.

Current Development Stage Phase II

Route of Administration Parenteral-Subcutaneous

Data GV1001 effectively blocks β -amyloid toxicity by mimicking the extra-telomeric functions of human telomerase reverse transcriptase, including the induction of cellular proliferation, anti-apoptotic effects, mitochondrial stabilization, and anti-aging and anti-oxidant effects.



*P<0.05 (vs the Control group), #P<0.05 (vs the 20 μ M A β group) N=6

Patent Position Patent No. 10-1685551 / KR, Patent No. 9540419 / US, Patent No. 201380036639.9 / CN

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A Potential Game Changer for partial onset seizures (Cenobamate)

SK biopharmaceuticals

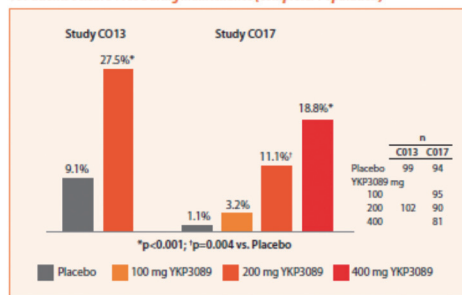


Product Type	Chemical Product
Indication	1st indication: Epilepsy, epilepsy (MeSH term)
Target	Voltage-gated Sodium channel inhibition and GABAergic modulation
MoA (Mechanism of Action)	Dual mechanism having sodium channel blocking and enhancing GABAergic inhibition
Differentiation Point	Best in class - Superior efficacy compared to other commercialized anti-epileptic drugs - Signals of 100% reduction in seizures - unique profile having both neurological and psychological profile
Current Development Stage	Phase III
Route of Administration	Oral
Data	- Cenobamate across all doses (100~400 mg) was highly effective vs. placebo in reducing the frequency of partial-onset seizures with traditional efficacy endpoints (median %seizure reduction; responder rate) - Cenobamate at 200 and 400 mg was highly effective vs. placebo in achieving seizure freedom - Most common AEs were dose-related and included somnolence, dizziness, fatigue, diplopia, and gait disturbances

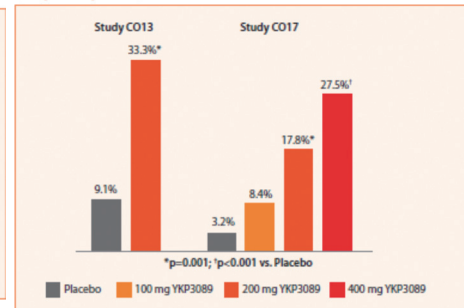
Traditional Efficacy Endpoints: Median % Reduction from Baseline 28-Day Seizure Frequency and Responder Rate (ITT Population)

	C013		C017			
	Placebo	YKP3089	Placebo	YKP3089		
		200		100	200	400
Median % reduction	21.5	55.6	24.0	35.5	55.0	55.0
P value		<0.001		0.007	<0.001	<0.001
Responder rate, %	22.2	50.4	21.7	40.7	56.9	60.4
P value		<0.001		0.003	<0.001	<0.001

% Patients Seizure-Free During Maintenance (Completer Population)



% Patients with ≥90% Reduction from Baseline Seizure Frequency (Completer Population)



Patent Position

US Patent No. US7598279B2

Development of a new lead compound targeting Histamine-releasing factor to treat atopic dermatitis

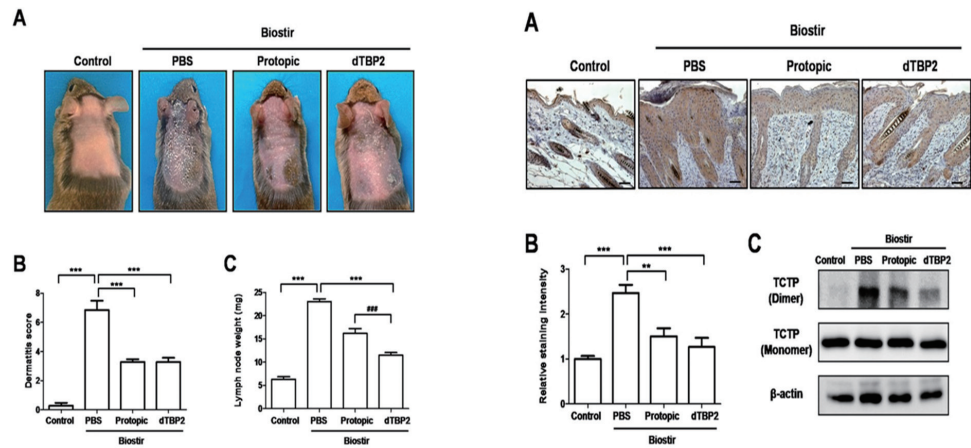
44

KOREA DRUG DEVELOPMENT FUND

Ewha Womans University



Product Type	Peptide Product (7 mer)
Indication	1st indication: atopic dermatitis, Dermatitis, Atopic (MeSH term)
Target	Histamine releasing factor
MoA (Mechanism of Action)	HRF inhibiting peptide(dTBP2)→HRF inhibition→Targeted therapy for AD
Differentiation Point	First In Class Novel target
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Parenteral-Subcutaneous
Data	Dimerized translationally controlled tumor protein-binding peptide 2 (dTBP2) ameliorates house dust mite-induced atopic dermatitis. dTBP2 suppresses level of dTCTP (target) in AD-like skin lesions.

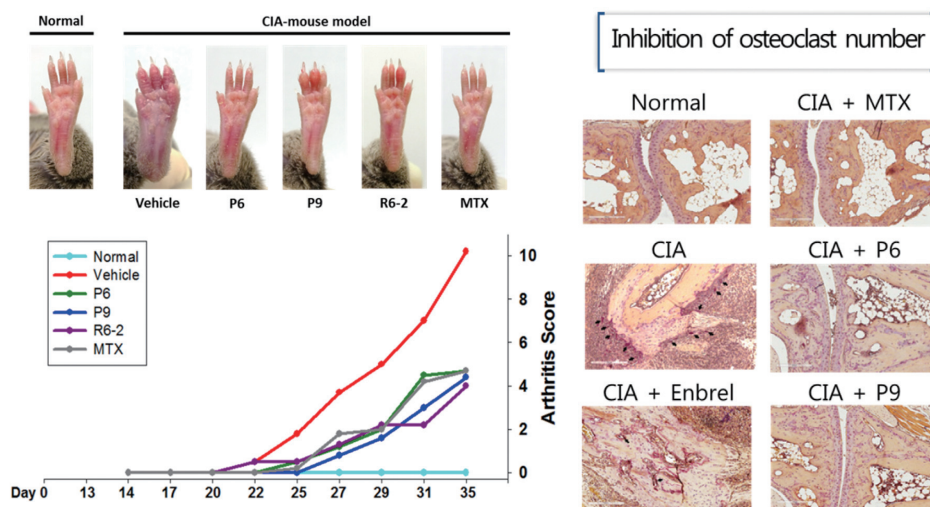


Development of drug candidates for treatment of rheumatoid arthritis by optimization of Erdr1-derived peptides

Sookmyung Women's University



Product Type	Peptide product
Indication	1st indication: Rheumatoid arthritis, Arthritis, Rheumatoid (MeSH term)
Target	Regulatory T-cells (T-reg)
MoA (Mechanism of Action)	Increased Treg cell number and activity → Suppression of rheumatoid arthritis pathogenesis
Differentiation Point	First In Class Novel small peptide from Erdr1 protein Specific target identification for each peptides
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous or Subcutaneous
Data	



YBL-004

Bi-specific Antibody that inhibit dual inflammatory axis

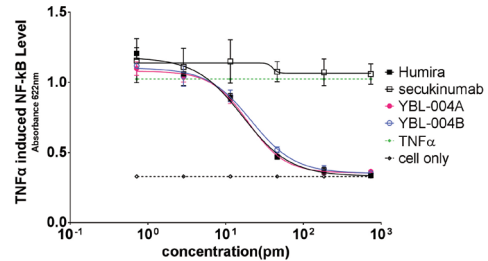
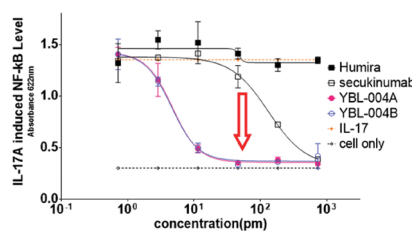
46

KOREA DRUG DEVELOPMENT FUND

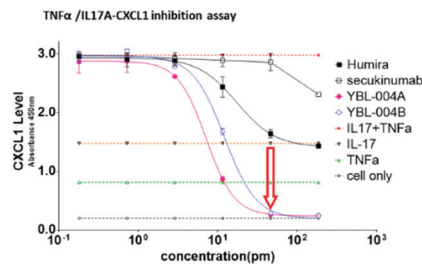
Y-biologics Inc.



Product Type	Immunoglobulin Product (mAb), Bispecific Antibody
Indication	1st indication: Ankylosing Spondylitis, Spondylitis, Ankylosing (MeSH term) 2nd indication: Psoriatic Arthritis, Arthritis, Psoriatic (MeSH term)
Target	Tumor necrosis factor alpha (TNF α) and Interleukin 17A (IL-17A)
MoA (Mechanism of Action)	YBL-004 neutralize TNF α and IL-17A simultaneously.--> Completely Inhibits the expression of pro-inflammatory cytokines than other single antibody.
Differentiation Point	<ul style="list-style-type: none"> • One shot Two kill: response rate & Disease modifying compared to TNF- α, IL-17A blockade • Could be applied to Non-responders to TNF- α blockade as an alternative strategy. • Easy to treatment & low cost compare to combination therapy • May be a choice for TNF- α blockade discontinuation
Current Development Stage	Pre-Clinical
Route of Administration	Parenteral-Subcutaneous
Data	YBL-004 : Synergy in inhibition of inflammatory cytokine production production YBL-004 shows better efficacy to inhibit IL-17 activity compared to Secukinumab & similar efficacy to inhibit TNF α activity compared to Humira.



YBL-004 shows synergy effect to inhibit the production of inflammatory cytokine.



Patent Position 10-2017-0149362 / Korea

2018 R&D Pipeline

A H4R antagonist with anti-pruritic & anti-inflammatory dual effect as a treatment for atopic dermatitis

JW Pharmaceutical



Product Type Chemical Product

Indication 1st indication: Atopic Dermatitis, Dermatitis, Atopic (MeSH term)
2nd indication: Pruritus, Pruritus (MeSH term)

Target Histamine H4 Receptor (H4R)

MoA (Mechanism of Action) ① Inhibition of pruritic signal transduction mediated by H4R
→ Blockade of pruritogen release → anti-pruritic effect
② Inhibition of immune cell migration
→ inhibition of Th2 cytokine production → anti-inflammatory effect

Differentiation Point First In Class
Novel target (FIC), Dual function, Broad safety margin

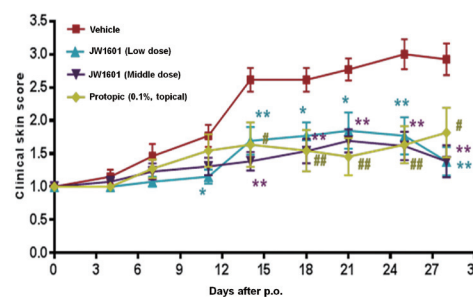
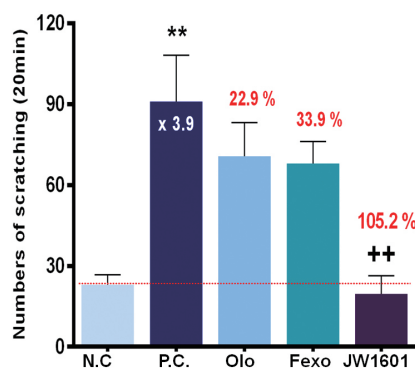
Current Development Stage Pre-Clinical

Route of Administration Oral (QI)

Data

- Histamine-induced itch model
- One-way ANOVA Dunnett's test
- ** p<0.01 vs N.C. and ++ p<0.01 vs P.C.
- Olo: Olopatadine (H1R antagonist)
- Fexo: Fexofenadine (H1R antagonist)

- Spontaneous NC/Nga AD mice model
- JW1601: Steel test, * p<0.05, ** p<0.01 vs Vehicle
- Protopic: Wilcoxon's test, # p<0.05, ## p<0.01 vs. Vehicle



Patent Position WO2013048214

CKD-506, a selective HDAC6 Inhibitor for the Treatment of Autoimmune Disease

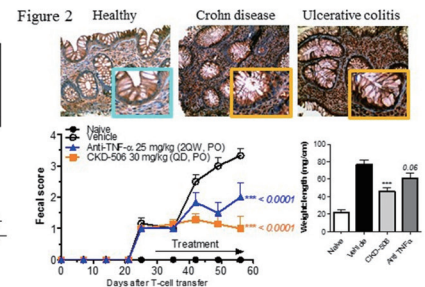
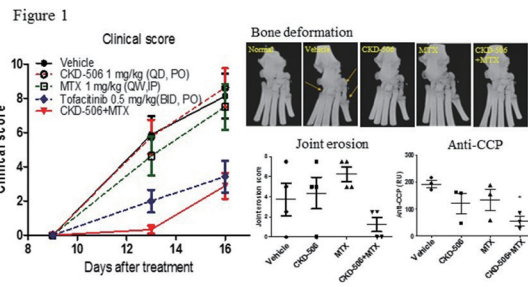
48

KOREA DRUG DEVELOPMENT FUND

Chong Kun Dang Pharmaceutical Corporation



Product Type	Chemical product
Indication	1st indication: Rheumatoid Arthritis, Arthritis, Rheumatoid (MeSH term) 2nd indication: Inflammatory Bowel Disease, Inflammatory Bowel Disease (MeSH term)
Target	Histone Deacetylase 6 (HDAC6)
MoA (Mechanism of Action)	TNF alpha Inhibition and Regulatory T cell Activation
Differentiation Point	First In Class CKD-506 has strong synergistic efficacy with Methotrexate in adjuvant-induced arthritis (AIA) and better efficacy than anti-TNF therapy in animal models of autoimmune disease rheumatoid arthritis and inflammatory bowel disease
Current Development Stage	Phase II a
Route of Administration	Oral
Data	Figure 1. Synergistic efficacy of CKD-506 with Methotrexate in AIA model Figure 2. Better efficacy of CKD-506 than anti-TNF therapy in T cell transferred inflammatory bowel disease animal model



Patent Position 10-2014-0051151/Korea, 14792173.8/Europe, 14/785812/US

2018 R&D Pipeline

Development of SYK inhibitor for Rheumatoid Arthritis

Oscotec



Product Type	Chemical Product
Indication	1st indication: Rheumatoid Arthritis, Arthritis, Rheumatoid (MeSH term) 2nd indication: Autoimmune Thrombocytopenia, Purpura, Thrombocytopenic, Idiopathic (MeSH term)
Target	Spleen tyrosine kinase (SYK)
MoA (Mechanism of Action)	ATP competitive SYK inhibitors
Differentiation Point	Best In Class SKI-O-793 has better efficacy and safety characteristics in animal and human study when compared to existing SYK inhibitors (R788, P505-15).
Current Development Stage	Phase II a
Route of Administration	Oral
Data	<p>1. Phase I clinical study</p> <p>1) Single ascending dose (SAD) study : completed</p> <ul style="list-style-type: none"> - Clinical safety (50 to 800 mg oral qd dosing) : no outstanding issue found at any test dose and no other significant findings, including vital signs, ECG and laboratory tests (hematology, serum chemistry, urinalysis) - Strong PD effect in activated basophil followed by anti-IgE stimulation Estimated EC50 of SKI-O-703, ~350 nM in the % activated basophil <p>2) Multiple ascending dose (MAD) study : completed</p> <ul style="list-style-type: none"> - 200 mg (qd & bid) and 400 mg (qd): completed at Q2, 2017 - Clinical safety : no outstanding issue found at any test dose - Reproducible PD effect in activated basophil followed by anti-IgE stimulation <p>2. Phase II clinical study</p> <p>1) Rheumatoid arthritis : US FDA IND in 2017 Q4</p> <p>2) Immune thrombocytopenic purpura : US FDA IND in 2018 Q2</p>

Novel vaccine lead (Δ SpA-ICW) for methicillin-resistant Staphylococcus aureus (MRSA) infection

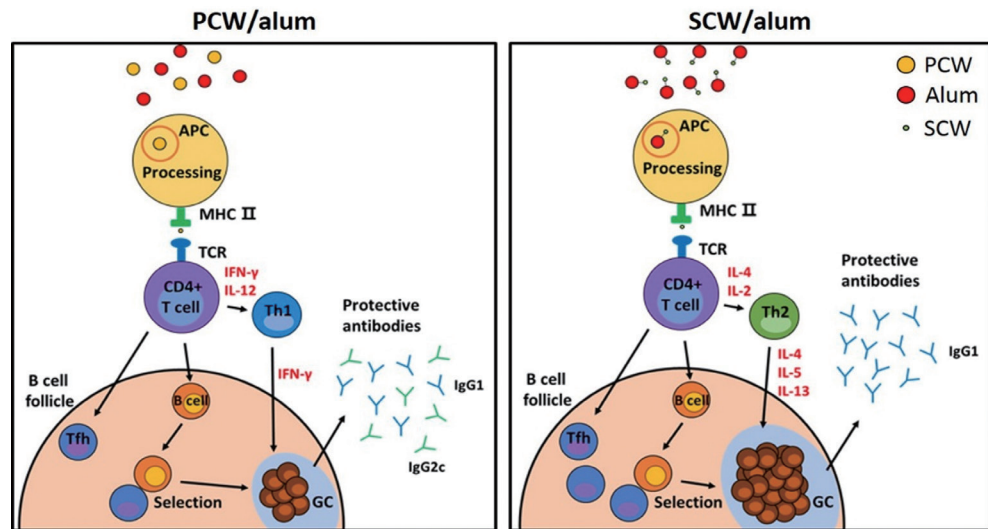
50
KOREA DRUG DEVELOPMENT FUND

Pusan National University



PUSAN NATIONAL UNIVERSITY

Product Type	Vaccine (Subunit Vaccine)
Indication	1st indication: MRSA, Methicillin-Resistant Staphylococcus aureus (MeSH term) 2nd indication: cell wall, cell wall (MeSH term)
Target	Methicillin-Resistant Staphylococcus aureus
MoA (Mechanism of Action)	Vaccination of cell walls → Engulfment by host phagocytes → Phagocytosis → Presentation to T cells & B cells → Memory B cells → MRSA infection → Long-lived Plasma cells → Humoral-immunity mediated clearance
Differentiation Point	First In Class Immunization of purified particulate and soluble staphylococcal cell wall derivatives induce host innate and acquired immune responses, subsequently inducing early stage MRSA clearance, and protecting the host using a memory immune response to a secondary MRSA challenge.
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Undecided
Data	



APC, antigen presenting cell; MHC II, major histocompatibility complex class II; TCE, T cell receptor; Tfh, T follicular helper cell; GC, germinal center
 * PCW : Particulate Cell Wall
 * SCW : Soluble Cell Wall

2018 R&D Pipeline

Novel antibiotic peptide based on toxin-antitoxin system

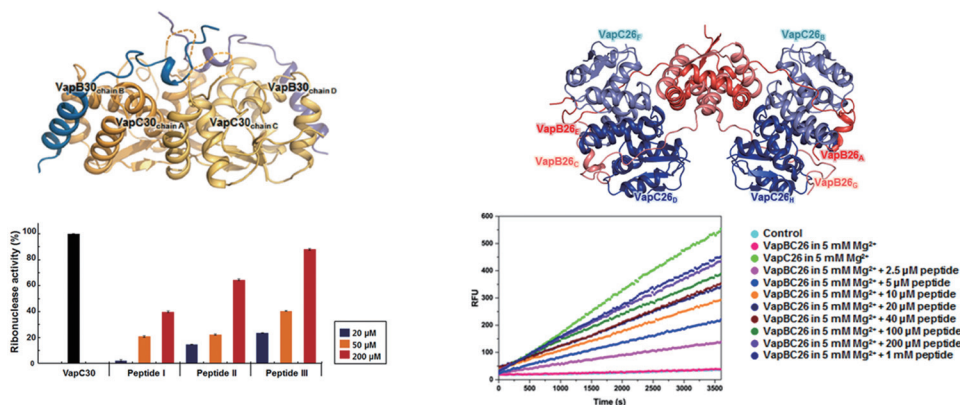
Seoul National University



서울대학교
SEOUL NATIONAL UNIVERSITY

Product Type	Peptide Product (Antimicrobial peptide)
Indication	1st indication: Tuberculosis, Tuberculosis (MeSH term)
Target	Toxin-Antitoxin system
MoA (Mechanism of Action)	Antimicrobial peptides disrupt the Toxin-Antitoxin complex → Free toxin release and cell death
Differentiation Point	First In Class Inhibitory peptides target new mechanism of bacterial physiology
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Parenteral-Intravenous

- Data**
- The toxicity is not detected in The toxin-antitoxin complex where The toxin toxicity is neutralized by The antitoxin.
 - When the complex is collapsed due to the designed drug, the toxin is liberated and toxicity is detected.



Patent Position

10-1746160 / Korea

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Recombinant monomeric and scaffold-based viral antigens

52 Korea University Research and Business Foundation Sejong



KOREA DRUG DEVELOPMENT FUND

Product Type Vaccin (recombinant protein antigens / scaffold-based multiantigens)

Indication 1st indication: influenza vaccines, Influenza Vaccines (MeSH term)

Target Generation of influenza virus hemagglutinin-specific antibodies

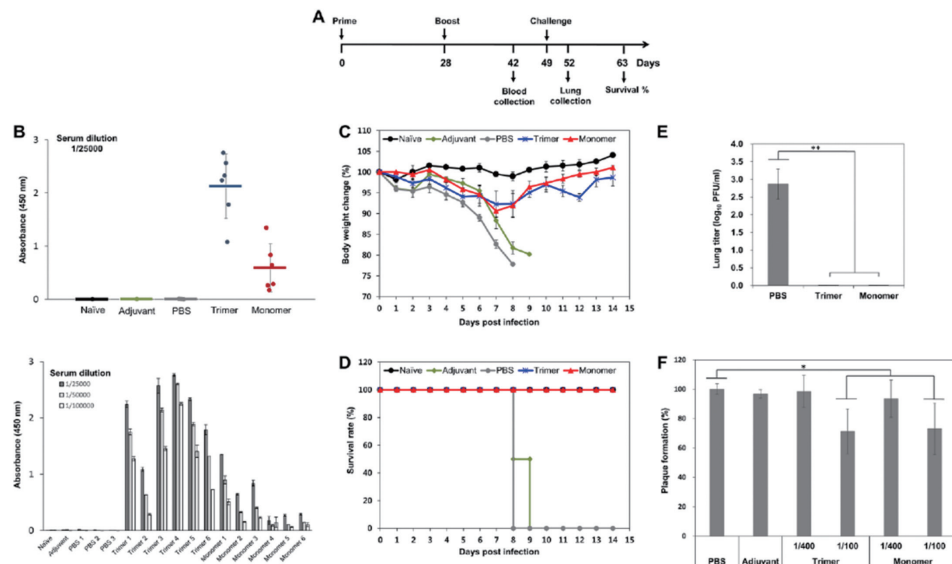
MoA (Mechanism of Action) Virus attachment, fusion, maturation, and budding

Differentiation Point First In Class
Efficacy comparable to the trivalent vaccine marketed by SK Chemicals, first monomeric antigens and scaffold-based multiantigen

Current Development Stage Lead Generation (Hit to Lead)

Route of Administration Intramuscular/Intranasal

Data Protection of mice against virus infection by preliminary recombinant hemagglutinin monomers. A) Schedule of mouse immunization, B) IgG antibody titration, C) Body weight changes and D) survival rate of PR8-challenged groups, and F) Plaque reduction neutralization assay.



Patent Position 10-2016-0151609/Korea

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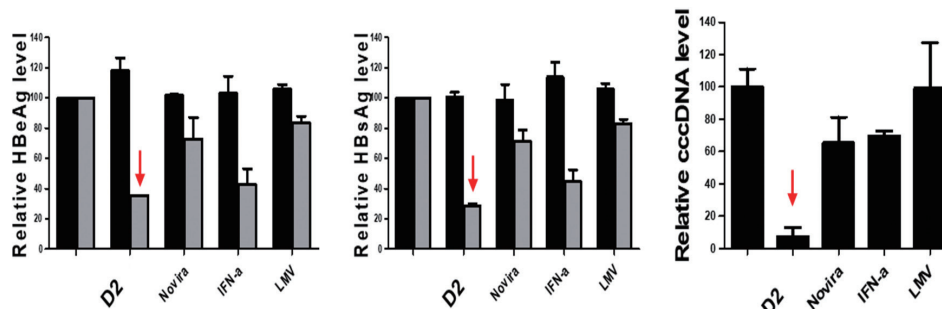
(+82)2-3290-3945

A novel agent targeting covalently closed circular DNA of HBV

Konkuk University



Product Type	Oligonucleotide
Indication	1st indication: Hepatitis B, Hepatitis B (MeSH term)
Target	Covalently closed circular DNA
MoA (Mechanism of Action)	Directly targets HBV covalently closed circular DNA
Differentiation Point	First In Class The drug works on all genotypes of hepatitis B virus and drug-resistant mutant viruses.
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Undecided Subcutaneous / Intravenous
Data	Inhibition of viral replication, antigen expression, and level of covalently closed circular DNA in primary human hepatocytes (PHHs).



Patent Position

PCT/KR2017/014662

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Discovery of novel anti-tubercular agent for the treatment of MDR/XDR TB

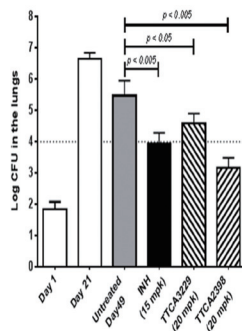
54

KOREA DRUG DEVELOPMENT FUND

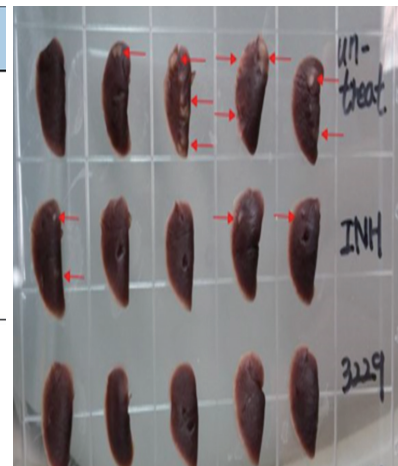
Institut Pasteur Korea



Product Type	Chemical Product
Indication	1st indication: Tuberculosis, Pulmonary, Bacterial Infections and Mycoses (MeSH term)
Target	MDR/XDR TB patients including drug sensitive TB patients
MoA (Mechanism of Action)	Novel (Not identified yet)
Differentiation Point	First In Class Active against latent TB and MDR TB strains / Much more active against TB within macrophage than in liquid broth culture medium / Potent bactericidal effect in vivo / Expected novel MOA related to host or mycobacterial factor within macrophage
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Oral
Data	- Significantly reduced bacterial load (more potent than INH) - Lower number of granulomatous foci in the lung than untreated and INH treated group



Groups	Lung		
	Log ₁₀	SD	n
D_1 Pre-treatment control	1.85	0.24	5
D_21 Pre-treatment control	6.64	0.20	5
After 49 days administration			
Vehicle Un-treat	5.47	0.47	5
INH (positive control)	3.93	0.36	4
TTCA3229_20mg/kg, QD	4.59	0.32	4
TTCA2398_20mg/kg, QD	3.12	0.32	2



Patent Position PCT/EP2015/063982

2018 R&D Pipeline

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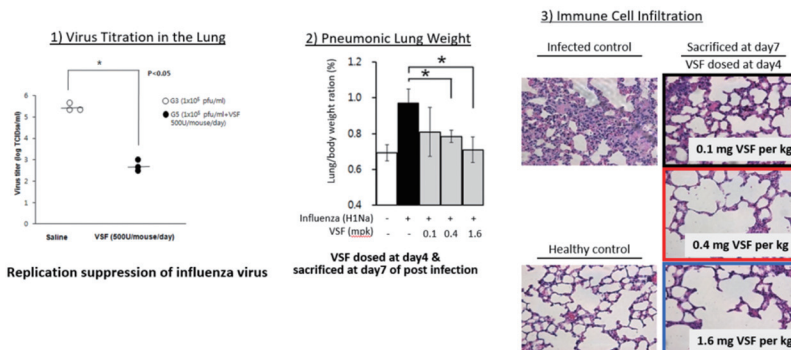
(+82)31-8018-8009

A novel anti-viral therapeutic antibody selectively targeting virus-infected cells

ImmuneMed, Inc.



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: Virus diseases, Virus disease (Mesh term) 2nd indication: Inflammatory skin disease induced by viral infection, Dermatitis (Mesh term)
Target	Virus suppressing factor Receptor (VR): viral infection induced, conformationally changed vimentin expressed on the cell surface
MoA (Mechanism of Action)	Proprietary immunoglobulinG4(IgG4) named as Virus Suppressing Factor (VSF) shows anti-viral, anti-inflammatory activity via association of its receptor (VR) which specifically expressed on the surface of virus-infected cells. The receptor protein is vimentin isoform, 60 kilo-Dalton in size and is conformationally changed form of natural vimentin in the cytoplasm. Anti-viral, anti-inflammatory activity of VSF was shown at association of VSF and its receptor VR followed by internalization of the complex into the lysosome and degradation in it.
Differentiation Point	First in Class
Current Development Stage	Pase I
Route of Administration	Parenteral-Intravenous/Parenteral-Intramuscular/Topical administration
Data	<ul style="list-style-type: none"> • Anti-viral and anti-inflammatory efficacy of VSF in the mice infected with influenza virus: 1) Virus titer was reduced by 1,000 folds, 2) Pneumonic lung was recovered or lung weight was decreased in dose-dependent manner and 3) Infiltration of immune cells was suppressed. • Pre-Clinical tests (in vitro, in vivo) have been done for influenza A, influenza B, Hepatitis B and Hepatitis C. VSF was effective in all the test per formed without cytotoxicity or any side effect.



Patent Position 10-2016-0072697/KR, PCT/KR2016/006215
10-2017-0110924/KR, PCT/KR2017/013706

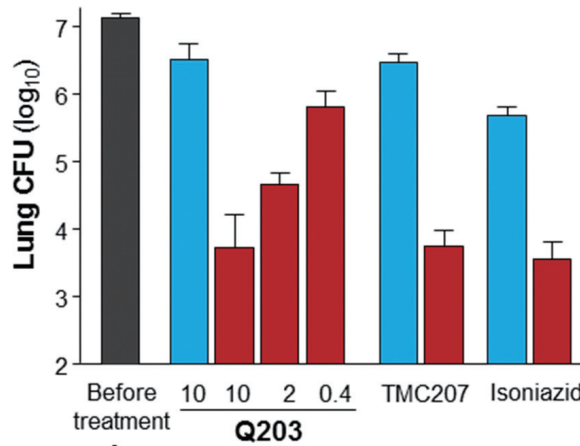
Q203, Anti-tuberculosis drug candidate

56
KOREA DRUG DEVELOPMENT FUND

Qurient Co. Ltd.,



Product Type	Chemical Product
Indication	1st indication: Tuberculosis, Tuberculosis, Pulmonary (MeSH term)
Target	First-in-class compound targeting cytochrome bc1 complex (Complex III) QcrB subunit in TB
MoA (Mechanism of Action)	Blocking of ATP synthase (both aerobic and hypoxic) and decrease in oxygen consumption rate
Differentiation Point	First In Class Novel mode of action (First-in-Class) Potent in vitro/in vivo, activity against drug resistant. Reduce treatment period (latency) QD or less frequent dosing schedule (increase adherence)
Current Development Stage	Phase I b
Route of Administration	Oral
Data	Strong efficacy in an established mouse TB model Strong efficacy against 13 MDR & 15 XDR clinical isolates



2018 R&D Pipeline

Patent Position PCT/EP2011/001345

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Clinical study of new oxazolidinone antimicrobial LCB01-0371

LegoChemBio Sciences



Product Type Chemical Product (Oxazolidinone Antibiotic)

Indication 1st indication: MDR-TB, Tuberculosis, Multidrug-Resistant(MeSH term)
2nd indication: MRSA, Methicillin-Resistant Staphylococcus aureus(MeSH term)

Target Gram+ (MRSA, VRE, S. pneumoniae), MDR-TB

MoA (Mechanism of Action) Inhibition of Mitochondrial Protein Synthesis

Differentiation Point Best in class
Safe and Potent Second Generation Oxazolidinone Antibiotics

Current Development Stage Phase II a

Route of Administration Oral / Parenteral-Intravenous

Data

- Superior In-Vitro & In-Vivo activity compared with Linezolid
- No cross-resistance and Low resistant rate
- Excellent efficacy in TB and NTM (M. abscessus) mouse model

✓ **Activity summary of LCB01-0371 and comparator antimicrobial agents**

Organism	Number	Antimicrobial agent	MIC ₅₀	MIC ₉₀	Range	CLSI %S / %R	EUCAST %S / %R
<i>Staphylococcus aureus</i>	100	LCB-0371	1	4	0.5 – 16	- / -	- / -
		Linezolid	1	8	0.5 – 32	82.0 / 18.0	82.0 / 18.0
		Vancomycin	1	1	0.5 – 2	100.0 / 0.0	100.0 / 0.0
<i>Staphylococcus aureus</i> (Linezolid-resistant)	21	LCB-0371	4	16	1 – 16	- / -	- / -
		Linezolid	8	32	4 – 32	14.3 / 85.7	14.3 / 85.7
		Vancomycin	1	2	0.5 – 2	100.0 / 0.0	100.0 / 0.0
Coagulase-Negative Staphylococci	99	LCB-0371	1	16	0.5 – 32	- / -	- / -
		Linezolid	1	64	0.25 – >64	60.6 / 39.4	60.6 / 39.4
		Vancomycin	2	2	0.5 – 4	100.0 / 0.0	99.0 / 1.0
<i>Enterococcus faecalis</i>	80	LCB-0371	1	1	0.5 – 2	- / -	- / -
		Linezolid	1	1	0.25 – 2	100.0 / 0.0	100.0 / 0.0
		Vancomycin	1	2	1 – 4	100.0 / 0.0	100.0 / 0.0
<i>Enterococcus faecium</i>	99	LCB-0371	1	8	0.5 – 16	- / -	- / -
		Linezolid	1	16	0.5 – 32	68.7 / 23.2	76.8 / 23.2
		Vancomycin	>16	>16	0.5 – >16	24.2 / 75.8	24.2 / 75.8
<i>Enterococcus faecium</i> (Linezolid-resistant)	30	LCB-0371	4	8	2 – 16	- / -	- / -
		Linezolid	8	16	4 – 32	0.0 / 76.7	23.3 / 76.7
		Vancomycin	>16	>16	1 – >16	13.3 / 86.7	13.3 / 86.7
<i>Haemophilus influenzae</i>	51	LCB-0371	4	4	2 – 8	- / -	- / -
		Linezolid	16	16	8 – 32	- / -	- / -

Patent Position

PCT/KR2009/005376, PCT/KR2009/0020525

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Development of New Antibiotics against Gram Negative Pathogens

LegoChemBio Sciences



Product Type	Chemical Product (Siderophore-cephalosporin conjugate)
Indication	1st indication: Pseudomonas aeruginosa, Pseudomonas aeruginosa (MeSH term) 2nd indication: Acinetobacter baumannii, Acinetobacter baumannii (MeSH term)
Target	Penicillin-binding protein
MoA (Mechanism of Action)	Disrupts cell wall synthesis
Differentiation Point	Increased transportation into bacterial cells
Current Development Stage	Pre-Clinical
Route of Administration	Oral
Data	- Excellent anti-pseudomonas activity - Safe and Potent anti-Pseudomonal Antibiotic Compound

	K. pneumoniae (n=198)			P. Aeruginosa (n=209)			A. baumannii (n=200)		
	MIC50	MIC90	Range	MIC50	MIC90	Range	MIC50	MIC90	Range
LCB10-200	0.125	8	0.125 - 64	0.25	1	0.125 - 16	1	16	0.125 - 64
LCB10-200 + BLI	≤0.0625	≤0.0625	≤0.063 - 1	0.25	0.5	≤0.063 - 16	1	4	≤0.063 - 64
Meropenem	0.0625	0.0625	0.063 - 64	0.5	16	0.063 - 128	16	128	0.063 - 128
Aztreonam	0.0625	32	0.063 - 128	8	64	0.25 - 128	64	128	4 - 128
Aztreonam + AVI	0.0625	0.125	0.063 - 1	8	32	0.25 - 128	32	128	4 - 128
Ceftazidime	0.25	128	0.063 - 128	4	128	1 - 128	32	128	1 - 128
CAZ + AVI	0.125	1	0.063 - 128	2	8	1 - 128	16	128	1 - 128
Colistin	0.5	1	0.063 - 128	2	4	0.063 - 64	0.5	1	0.25 - 128

Patent Position PCT/KR2012/002302

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A First-in-class Anti-MRSA Agent

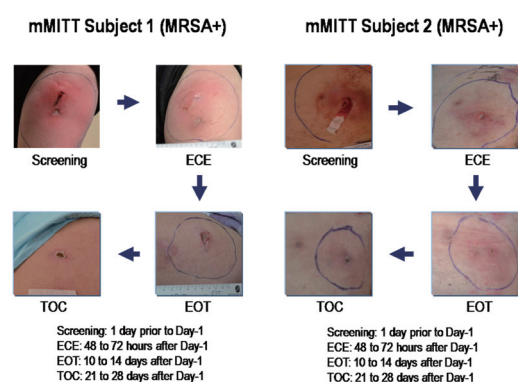
CrystalGenomics



Product Type	Chemical Product
Indication	1st indication: Methicillin-resistant Staphylococcus aureus infections, Bacterial Infections and Mycoses (MeSH term)
Target	Enoyl-acyl carrier protein reductase (FabI)
MoA (Mechanism of Action)	A novel mechanism of blocking synthesis of fatty acid, a key component of bacterial cell membrane formation process.
Differentiation Point	First In Class
Current Development Stage	Phase II
Route of Administration	Oral / Intravenous infusion
Data	<ul style="list-style-type: none"> ① CG400549 demonstrated the lowest MIC values against MRSA and VRSA ② Drastic reduction of redness & edema by ECE and clinical cure by EOT (Phase IIa study in US)

Drug	Methicillin-Susceptible (µg/ml)			Methicillin-Resistant (µg/ml)		
	Range	MIC ₅₀	MIC ₉₀	Range	MIC ₅₀	MIC ₉₀
CG400549*	0.06-1.0	0.25	0.25	0.06-1.0	0.25	0.25
Vancomycin*	1.0-2.0	1.0	2.0	1.0->64.0	1.0	2.0
Linezolid*	0.25-2.0	1.0	2.0	0.25-2.0	1.0	2.0
Quinupristin-dalfopristin*	0.25-2.0	1.0	2.0	0.25-2.0	1.0	2.0
Daptomycin*	0.25-2.0	1.0	1.0	0.25-4.0	0.5	0.5
Clindamycin**	0.25-2.0	0.25	0.25	<0.25->2.0	<0.25	>2.0
Tetracyclines**	<2.0->8.0	<2.0	<2.0	<2.0->8.0	<2.0	<2.0
Tigecycline**	<0.03-1.0	0.12	0.25	<0.03-1.0	0.12	0.25
TMP-SMX**	<0.5->2.0	<0.5	<0.5	<0.5->2.0	<0.5	<0.5
Ceftaroline**	<0.008-1	0.25	0.25	0.12-2.0	1.0	1.0

* Measured in Hershey Hospital, PA, USA
 **Clinical Infectious Diseases 2012, s206.



Patent Position

US7973060B2

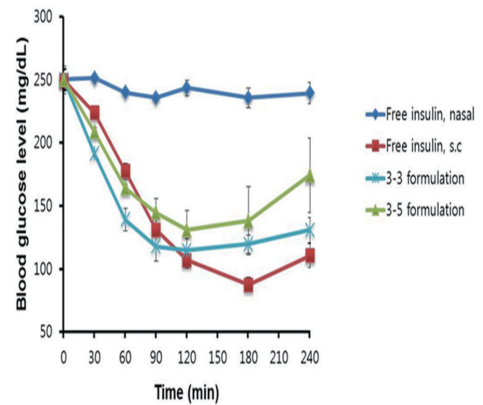
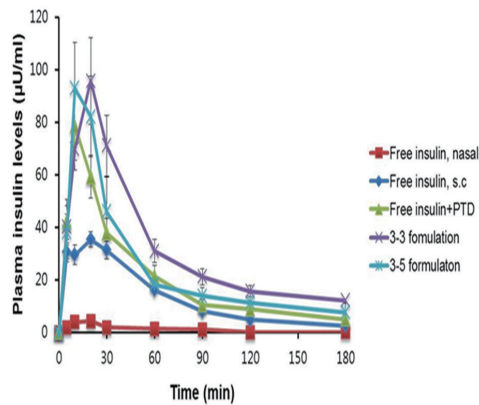
Development of intranasal insulin using TCTP-PTD

60
•
KOREA DRUG DEVELOPMENT FUND

Ewha Womans University



Product Type	Cell penetrating peptide
Indication	1st indication: diabetes, diabetes mellitus (MeSH term)
Target	Insulin resistance
MoA (Mechanism of Action)	Insulin/TCTP-PTD formulation → Nasal delivery of insulin → Change in blood glucose levels
Differentiation Point	First In Class Nasal delivery of insulin using TCTP-PTD may serve the user-friendly medications that are safe and effective for the treatment of diabetes.
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Parenteral-Others (Nasal administration)
Data	Completed studies testing pharmacokinetics of insulin/TCTP-PTD. Changes in blood glucose levels in diabetic rats following nasal administration of insulin/TCTP-PTD



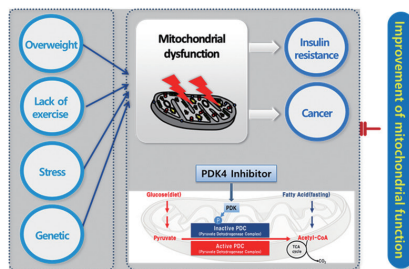
Discovery of PDK4 inhibitor for metabolic disease and cancer

Gwangju Institute of Science and Technology

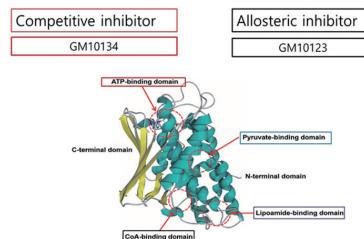


Product Type	Chemical Product
Indication	1st Indication: Diabetes, Diabetes Mellitus (MeSH) 2nd Indication: Breast Cancer, Breast Neoplasms (MeSH)
Target	Pyruvate dehydrogenase kinase (PDK) 4
MoA (Mechanism of Action)	PDK inhibition --> Control PDC (Pyruvate dehydrogenase complex) activity -->Control Glucose metabolism
Differentiation Point	First In class Novel Mitochondrial Targeting Compound
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Oral / Intravenous
Data	

Mitochondrial dysfunction: Insulin resistance and cancer

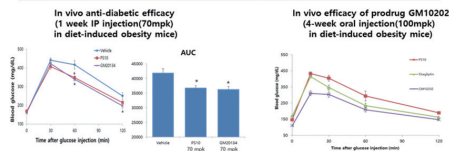


Discovery of PDK4 inhibitors



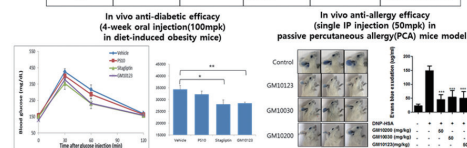
GM10134 (Competitive inhibitor)

code	Inhibitor class	In vitro activity	In vitro (cellular assay)	Selectivity	Liver microsomal stability
GM10134	Competitive inhibitor	81 nM		PDK4-PDK2 81% (I) 81% (II) (30 min incubation)	
	CYP inhibition	Herg inhibition	LogP	In vivo PK	In vivo efficacy
	1A2: 11%, 2C9: 18%, 2C19: 24%, 2D6: 6.8%, 3A4: 39% at 10 uM	2% inhib at 10 uM	3.72	IV (5mg/kg) T1/2: 8.54 h AUC: 1.19 ug/h/ml CL: 3.83 L/h/kg V: 1.87 L/kg BA: 11%	Glucose AUC reduction (OGTT) IP and oral administration



GM10123 (Allosteric inhibitor)

code	Inhibitor class	In vitro activity	In vitro (cellular assay)	Selectivity	Liver microsomal stability
GM10123	Allosteric inhibitor	83 nM		PDK4 + PDK2 inhibition of PDK4a phosphorylation (30 min incubation)	
	CYP inhibition	Permeability	Cell cytotoxicity (IC ₅₀)	In vivo PK	In vivo efficacy (anti-diabetic)
	2C9: 27.4%, 2C19: 52.2%, 2D6: 47.3%, 3A4: 18.5%, inhibition at 10 uM	1.99 (0.31 Papp x 10 ⁻⁶ cm/sec)	VERO: 36.42 uM HFL: 38.54 uM L029: 35.82 uM HEP2: 39.64 uM CHO-K1: 36.02 uM	IV (5mg/kg) Oral (10 T1/2: 1.06 h) AUC: 7.02 ug/h/ml CL: 0.84 L/h/kg V: 5.52 L/kg BA: 64%	Restoration of vascular permeability aggravated by dextranophenylated human serum albumin (DHP-HSA) challenge



Lead optimization of Selective Insulin Receptor Agonist aptamer

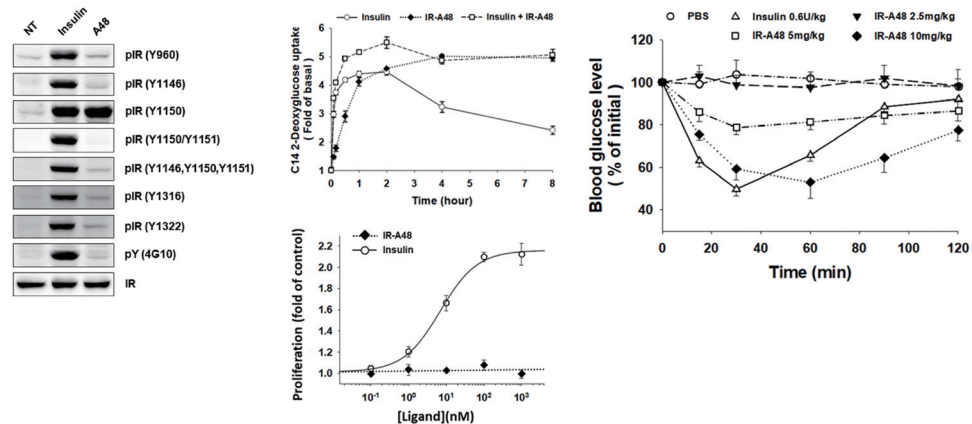
62

KOREA DRUG DEVELOPMENT FUND

Aptamer Sciences Inc.



Product Type	Aptamer
Indication	1st indication: Type 2 Diabetes, Diabetes Mellitus, Type 2 (MeSH term) 2nd indication: Type 1 Diabetes, Diabetes Mellitus, Type 1 (MeSH term)
Target	Insulin Receptor
MoA (Mechanism of Action)	Selective Insulin Receptor Agonist Aptamer binds to Insulin receptor, and induces monophosphorylation at the 1150th tyrosine residue on the kinase domain. → it causes biased activity to glucose uptake without mitogenic activity
Differentiation Point	First In Class
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Subcutaneous
Data	<ul style="list-style-type: none"> - Induces monophosphorylation at Tyr1150 on insulin receptor - Biased signaling activates glucose uptake without mitogenic activity - Dose dependent target activity following systemic injection



Patent Position PCT/KR2016/004665

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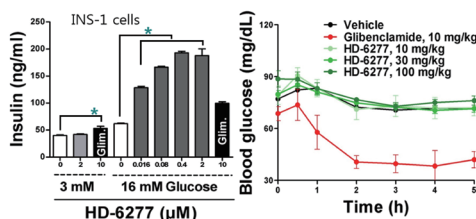
a Novel GPR40 Agonist, HD-6277, on Glycemic Control and Insulin Sensitivity in Type 2 Diabetes

Hyundai Pharmaceuticals

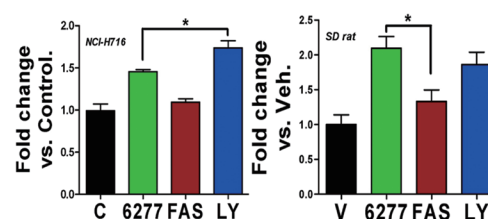


Product Type	Chemical Product
Indication	1st indication: Type 2 Diabetes, Diabetes Mellitus, Type 2 (MeSH term)
Target	G-protein-Coupled Receptor 40
MoA (Mechanism of Action)	Insulin secretagogue, Glucose dependent beta cell function regulation
Differentiation Point	First In Class
Current Development Stage	Phase I a
Route of Administration	Oral

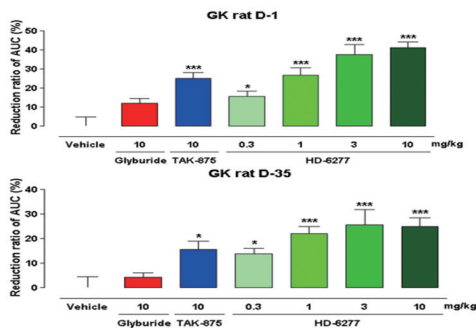
Data (A.) Dose dependent insulin secretion study in in vitro.
No hypoglycemic risk in in vivo.



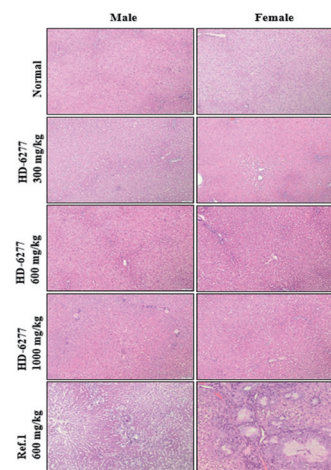
(B.) GLP-1 secretion assay in in vitro and in vivo.



(C.) Dose dependent glucose lowering effects.



(D.) Comparison of liver toxicity vs. Fasiglifam(TAK-875) with 2 weeks DRF study in Beagle dogs.



New drug development study for osteoporosis with novel mechanism to improve the clinical unmet needs

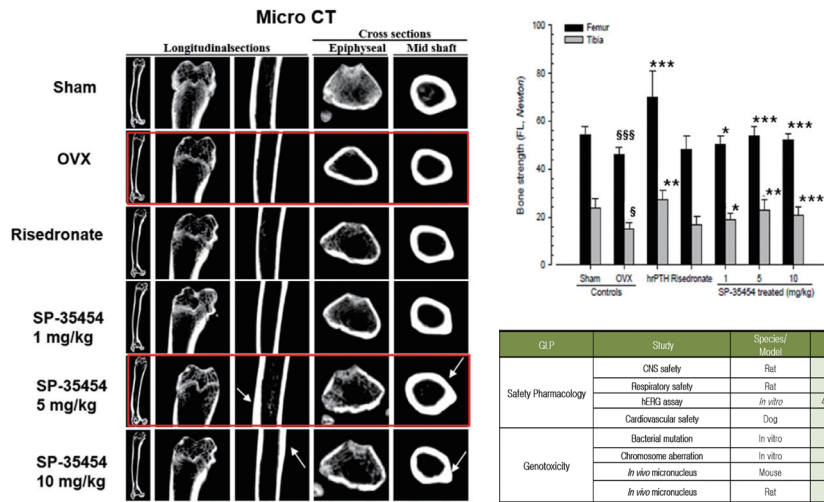
64

KOREA DRUG DEVELOPMENT FUND

Shin Poong Pharm. Co., LTD.



Product Type	Chemical Product
Indication	1st indication: Osteoporosis, Osteoporosis (MeSH term)
Target	Undisclosed
MoA (Mechanism of Action)	Undisclosed
Differentiation Point	Dual action: increase of bone formation and decrease of bone resorption. Oral application with much less side effects.
Current Development Stage	Phase I
Route of Administration	Oral
Data	Increase of the mid-shaft BMD/Bone thickness/Bone strength/ Non-clinical toxicity studies/Clinical Phase I study



GLP	Study	Species/Model	Major Findings
Safety Pharmacology	CNS safety	Rat	NOAEL = 1000 mg/kg
	Respiratory safety	Rat	NOAEL = 1000 mg/kg
	NERG assay	In vitro	45.9% inhibition at top conc.
	Cardiovascular safety	Dog	NOEL 1000 mg/kg
Genotoxicity	Bacterial mutation	In vitro	None
	Chromosome aberration	In vitro	None
	In vivo micronucleus	Mouse	None
	In vivo micronucleus	Rat	None
General toxicity	4 weeks rat toxicity study	Rat	NOAEL = 1000 mg/kg/day
	4 weeks dog toxicity study	Dog	NOAEL = 1000 mg/kg/day
	26 weeks rat toxicity study	Rat	Male: NOAEL = 600 mg/kg Female: NOAEL = 200 mg/kg
	38 weeks dog toxicity study	Dog	Male: NOAEL = 600 mg/kg Female: NOAEL = 70 mg/kg

• **Results: Clinical Phase I study was completed on March 2017 in Europe**

- Volunteers: healthy postmenopausal women
- **No SAE** (AE: fatigue, diarrhea, back pain, myalgia(muscle pain), headache)
- **No issues:** ECGs, vital signs, physical examination
- Pharmacokinetics (Food effects): delayed T_{max} and **dose-proportional increased AUC**
- Pharmacodynamics : Bone biomarker (MAD)

Patent Position PCT/KR2009/006085

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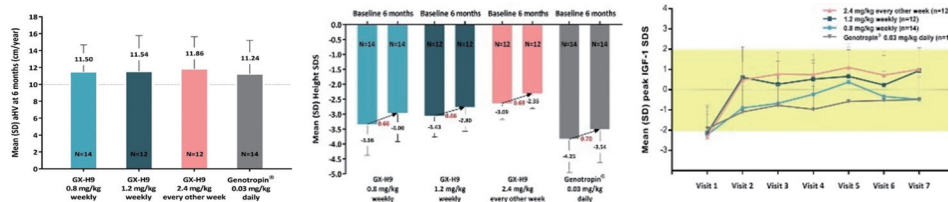
[+82]10-5378-8312

Long-acting Growth Hormone GX-H9 (hGH-hyFc)

Genexine, Inc.



Product Type	Fusion Protein, Hybrid Fc (hyFc), Long-Acting
Indication	1st indication: Pediatric Growth Hormone Deficiency, Pediatric Growth Hormone Deficiency (MeSH term) 2nd indication: Adult Growth Hormone Deficiency, Adult Growth Hormone Deficiency (MeSH term)
Target	Growth Hormone (GH), Insulin-like Growth Factor-1 (IGF-1)
MoA (Mechanism of Action)	Growth hormone stimulates the production of IGF-1 in liver. IGF-1 has growth-stimulating effects on a wide variety of tissues and also has stimulatory effects on osteoblast and chondrocyte activity to promote bone growth. Growth hormone itself also directly stimulates division and multiplication of chondrocytes of cartilage.
Differentiation Point	Best In Class To develop more convenient product versus current daily treatment by reducing dose frequency To address the adherence and treatment outcome issues
Current Development Stage	Phase II
Route of Administration	Parenteral-Subcutaneous
Data	<ul style="list-style-type: none"> - GX-H9 treatment for 6 months was found to be safe and well tolerated in pre-pubertal GH naïve patients with GHD - Annualized height velocity and improvement of height SDS were comparable across GX-H9 treatment groups and with currently available daily rhGH product, Genotropin® - Average peak IGF-1 SDS values were less than +2SDS regardless of GX-H9 dosage regimens



Patent Position

US 8529899 B2, WO2017/142331

Study of Global Clinical Trial and Production for next-generation diabetes drug, HM11260C

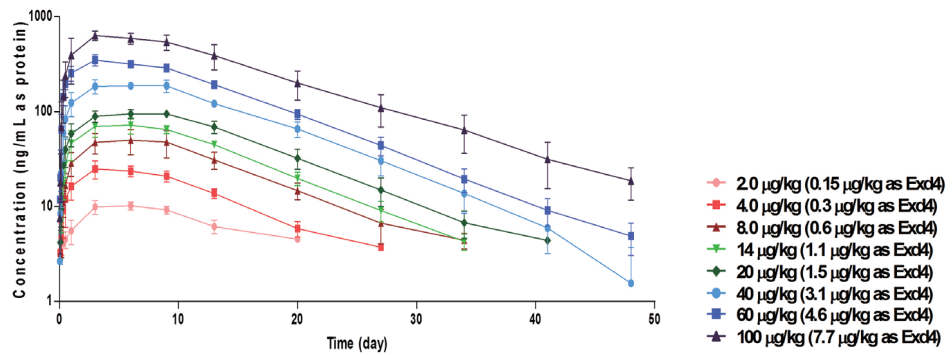
66

KOREA DRUG DEVELOPMENT FUND

Hanmi Pharm. Co., Ltd.



Product Type	Aglycosylated human Fc fragment conjugate
Indication	1st indication: T2DM, Diabetes Mellitus, Type 2(MeSH term) 2nd indication: Obesity, Obesity(Mesh Term)
Target	Glucagon like peptide 1 (GLP-1) receptor
MoA (Mechanism of Action)	GLP-1 receptor agonist
Differentiation Point	Best in Class Epfeglenatide activates GLP-1 receptor with superagonistics character and shows superior blood glucose lowering and body weight loss efficacy
Current Development Stage	Pase III
Route of Administration	Parenteral-Subcutaneous
Data	Completed Phase II study showed potent glucose lowering and body weight loss efficacy with once a week regimen and showed no significant adverse effects related to treatment



Patent Position WO2008082274

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Development of Novel Acute Heart Failure Medicine Targeting Actin-Myosin Cycle

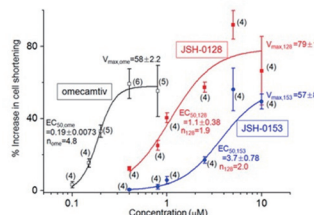
Shin Poong Pharm. Co., LTD.



Product Type	Chemical Product
Indication	1st indication: Heart failure, Heart Failure (Mesh term) 2nd indication: Myocardial Failure , Heart Failure (Mesh term)
Target	Cardiac myosin ATPase
MoA (Mechanism of Action)	Cardiac myosin activator binds S1 sub-domain of myosine, increasing the duration and amount of myocyte contraction.
Differentiation Point	First In Class Establishment of strategic position for world wide market of inotrope through the discovery of novel Ca ²⁺ non-dependent inotrope abolishing the serious side effects of current inotropes.
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous
Data	- In vitro Cardiac myosin ATPase activity and selectivity study

Compound	% ATPase Activity					
	Cardiac		Smooth		Skeletal	
	1 μM	10 μM	10 μM	100 μM	10 μM	100 μM
JSH-0128	ND	48.5	1.4	5.2	0	-4.8
JSH-0153	15.0	47.6	1.4	3.9	11.1	-20.2
Omecamtiv		~ 68.0	0	4.0	-5.5	2.8

- In vitro cardiac ventricular cell contractility study



- Omecamtiv mecabil VS. JSH

	OMM	JSH-0128	JSH-0153
Cardiac ventricular cell contractility, Vmax [in vitro]	58 ± 2.2	79 ± 13	57 ± 8.6
FS% increase [in vivo, normal rat, 16mg/kg/min for 3min]	20.60	31.85	24.07
NOAEL [mg/kg, 7 days repeated toxicity]	1.5	9.0	4.0

G-protein coupled receptor kinase 5 inhibitor to treat heart failure

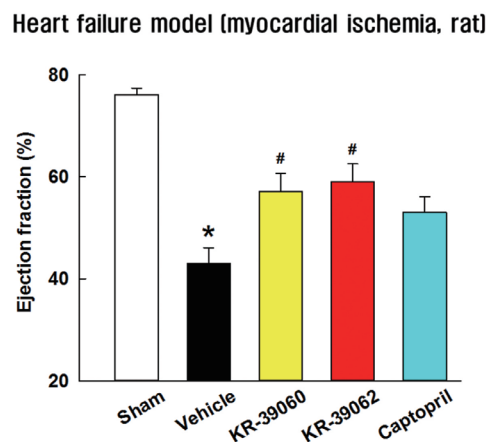
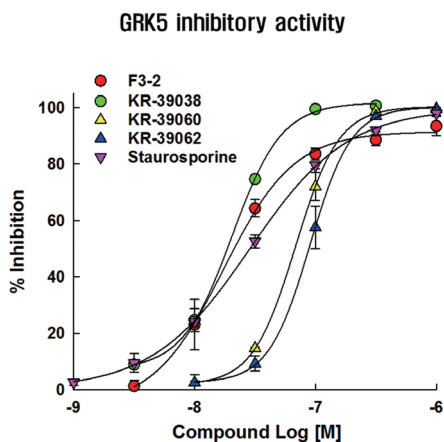
68

KOREA DRUG DEVELOPMENT FUND

Korea Research Institute of Chemical Technology



Product Type	Chemical Product
Indication	1st indication: Heart failure, Heart failure (MeSH term)
Target	G-protein coupled receptor kinase 5 (GRK5)
MoA (Mechanism of Action)	<ul style="list-style-type: none"> - GRK5 inhibitor suppresses the phosphorylation of c-terminal intracellular region of GPCR. - GRK5 inhibitor suppresses the functions as a specific HDAC5 kinase.
Differentiation Point	<p>First In Class</p> <p>The drug is devoid of unwanted side effect- hypotensive effect in rats.</p>
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Oral
Data	<ul style="list-style-type: none"> - Candidate compounds: KR-39038, 39060, 39062 - ATP non-competitive, substrate competitive - Effective both in pressure overload- and ischemia-induced heart failure models - No blood pressure lowering effect

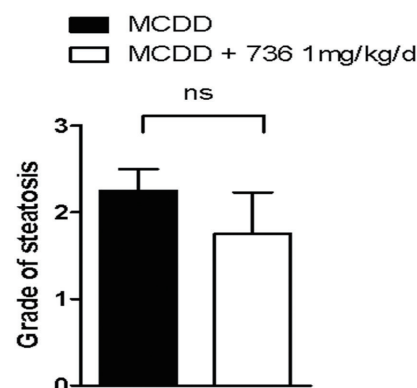
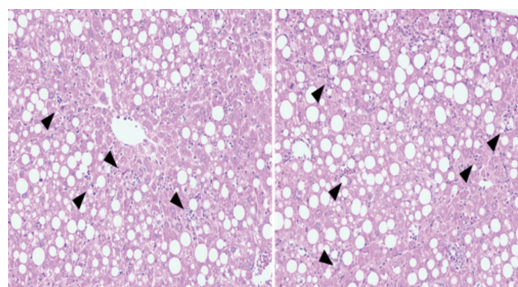


Development of a new drug for non-alcoholic steatohepatitis and liver cirrhosis using myriocin analogues

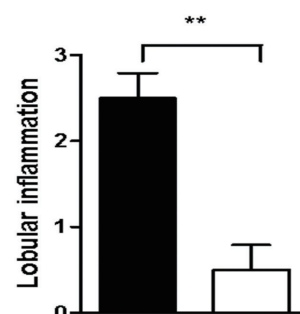
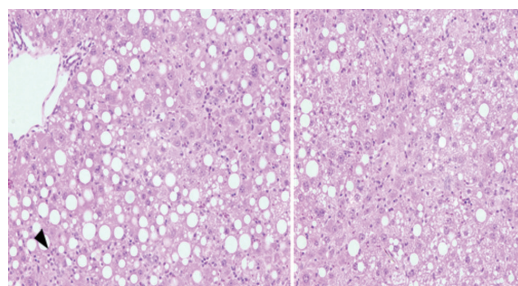
Asan Medical Center



Product Type	Chemical Product
Indication	1st indication: Non-alcoholic steatohepatitis, Non-alcoholic Fatty Liver Disease (Mesh term)
Target	S1PR4
MoA (Mechanism of Action)	Inhibition of inflamamsome activity
Differentiation Point	First In Class Treatment of non-alcoholic steatohepatitis, but not simple hepatic steatosis
Current Development Stage	Lead Generation (Hit to Lead)
Route of Administration	Oral
Data	Non-alcoholic steatohepatitis



SLB 736



Patent Position

PCT-KR-10-2017-0040139

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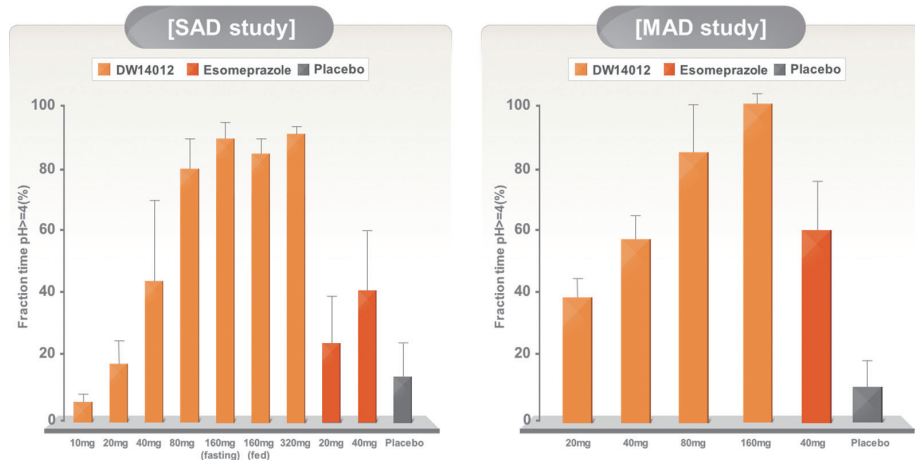
Development of Acid Pump Antagonist for Gastric acid related Disease, DWP14012

Daewoong Pharmaceutical Co., Ltd.



Product Type	Chemical Product
Indication	1st indication: Gastroesophageal Reflux Disease (GERD), Gastroesophageal Reflux (MeSH term) 2nd indication: Peptic Ulcer, Peptic Ulcer (MeSH term)
Target	Acid Pump Antagonist
MoA (Mechanism of Action)	Reversible and Potassium-Competitive Inhibition of Acid Secretion
Differentiation Point	Best In Class DWP14012 inhibits acid secretion in a reversible and potassium-competitive manner with fast onset of action. Phase I clinical trial shows that DWP14012 exhibited dose-dependent acid suppression in human, rapid onset time, excellent safety & tolerability, and favorable pharmacokinetic profiles with a once a day regimen.
Current Development Stage	Phase II
Route of Administration	Oral
Data	Phase I Clinical Study: Pharmacodynamics : DWP14012 showed dose-dependent acid suppression and clear exposure-response relationship. Compared to esomeprazole, DWP14012 demonstrated more favorable or higher extent of 24-hour pH(≥4) holding time.

Intragastric pH fraction time pH≥4(%)



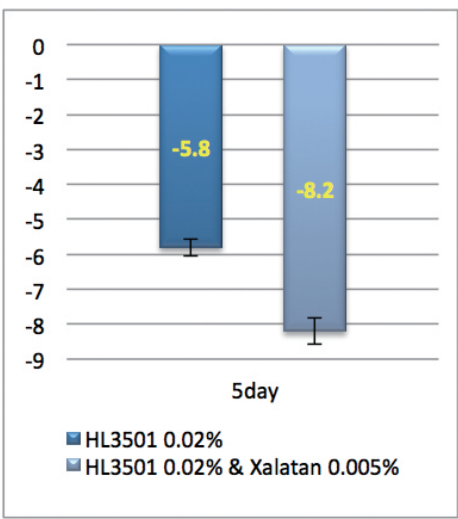
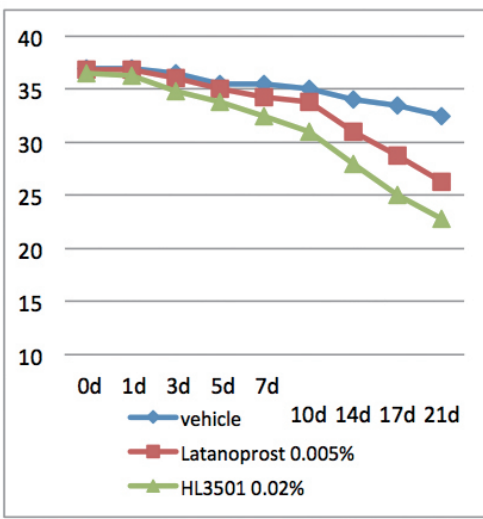
Patent Position PCT/KR2016/004411

Development of Anti-Glaucoma Drug with a Novel Mechanism of Action

Handok Inc.



Product Type	Chemical Product
Indication	1st indication: Primary open angle glaucoma, Glaucoma, Open-Angle (MeSH term) 2nd indication: Ocular hypertension, Ocular Hypertension (MeSH term)
Target	A3 adenosine receptor
MoA (Mechanism of Action)	A3 adenosine receptor antagonism effect induced lowering intraocular pressure.
Differentiation Point	Favorable safety profile from single tox(topical ocular treatment)
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Eye Drop
Data	<ul style="list-style-type: none"> HL3501 was superior to Latanoprost in laser model blocked main TM outflow of AH. The treatment of combination elicited faster IOP lowering effect than mono in HL3501 by 5day



Patent Position 15/406,556(US), PCT/KR2017/000492

Development of the dry AMD treatment

72

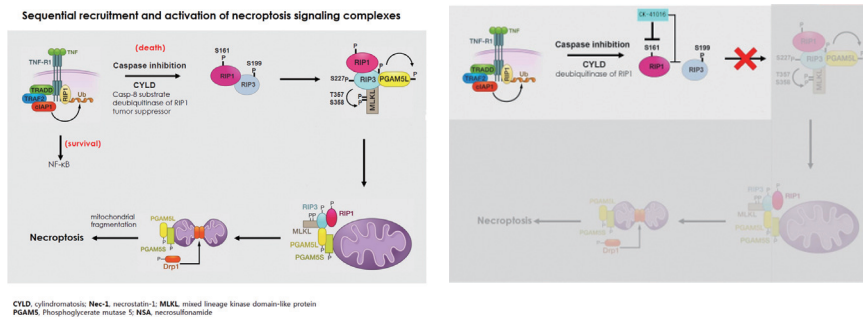
KOREA DRUG DEVELOPMENT FUND

Chungnam National University

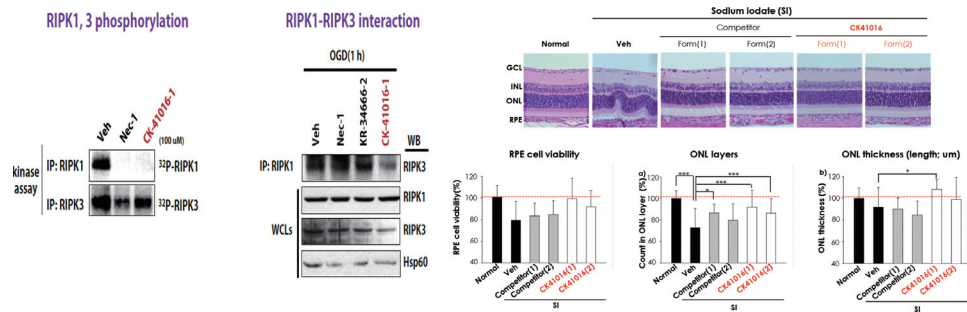
*KUKJE Pharm licensed KDDF-201202-10 and is developing this project.



Product Type	Chemical Product
Indication	1st indication: Blindness, Eye diseases (MeSH term) 2nd indication: Macular degeneration, Retina (MeSH term)
Target	Age-Related Macular Degeneration
MoA (Mechanism of Action)	RIP-dependent necroptosis signal pathway



Differentiation Point	First it Class - New molecular entity(NME) with a new structure - There are currently no available treatments in market
Current Development Stage	Pre-Clinical
Route of Administration	Eye drop
Data	- Inhibition of RIPK1 phosphorylation/RIPK1-RIPK3 interaction of CK41016 - Following eye drops administration in animals model of dry AMD, in vivo protective effects of CK41016 were superior to those of competitors



Patent Position KR10-1551313/KR10-1515985, PCT/KR2015/007562

2018 R&D Pipeline

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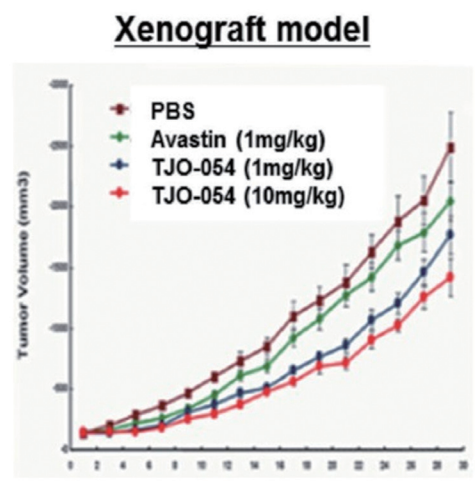
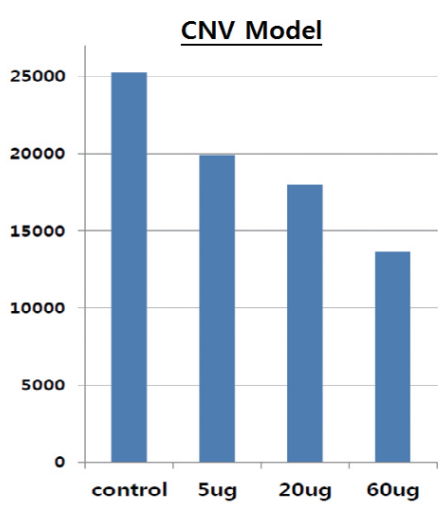
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Nonclinical studies of novel antibody TJO-054 for treatment of wet AMD

Taejoon Pharmaceutical Co., Ltd.



Product Type	Immunoglobulin Product (mAb)
Indication	1st indication: wet AMD, Eye Diseases(MeSH term)
Target	VEGFR2
MoA (Mechanism of Action)	VEGFR2-specific binding →blocking not only VEGF-A, but as well as VEGF-C and VEGF-D
Differentiation Point	First In Class TJO-054 is a fully human monoclonal antibody to bind VEGFR-2/KDR on abnormal vasculature related to new angiogenesis diseases including wet AMD.
Current Development Stage	Pre-Clinical
Route of Administration	Intravitreal injection
Data	Pre-Clinical Pharmacology



A Long-acting coagulation factor VIIa

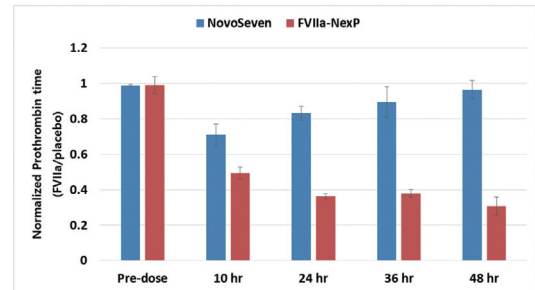
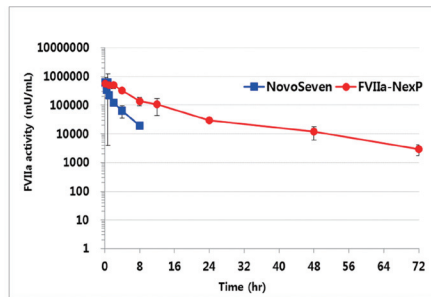
74

KOREA DRUG DEVELOPMENT FUND

Alteogen, Inc.



Product Type	Protein Product
Indication	1st indication: Hemophilia, Hemophilia A (MeSH term)
Target	Coagulation factor VIIa (FVIIa)
MoA (Mechanism of Action)	Recombinant FVIIa + NexPTM fusion → Long-acting factor VIIa (FVIIa-NexPTM)
Differentiation Point	Best In Class Prolonged half-life of factor VIIa activity among those of competitors (best-in-class potential)
Current Development Stage	Lead Optimization (Lead to Candidate)
Route of Administration	Parenteral-Intravenous - The possibility of subcutaneous administration is also being explored.
Data	- Significantly enhanced pharmacokinetics of factor VIIa as a NexPTM fusion in hemophilia mice - Prolonged coagulation activity of factor VIIa compared to NovoSeven in rats



Patent Position PCT/KR2010/002520, PCT/KR2012/006441

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Management of Chemotherapy-induced Neutropenia in Advanced Breast Cancer Patients

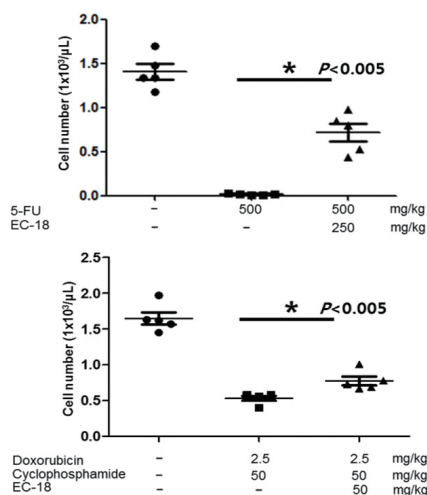
Enzychem Lifesciences Corporation

ENZYCHEM LIFESCIENCES

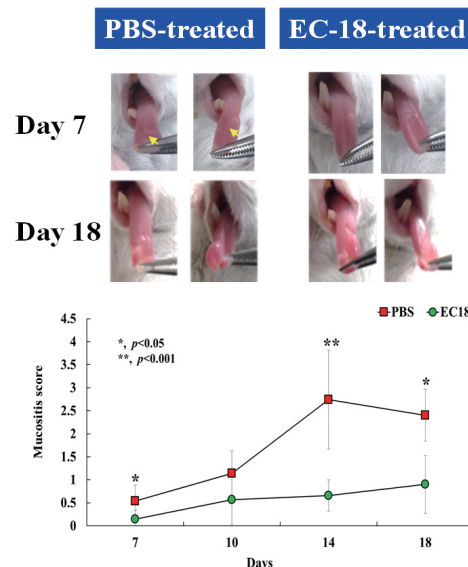
Product Type	Chemical Product (Diacylglycerol)
Indication	1st indication: Chemotherapy-induced Neutropenia (CIN), Neutropenia (MeSH term) 2nd indication: ChemoRadiation-induced Oral mucositis (CRIOM), Stomatitis (MeSH term)
Target	Undisclosed
MoA (Mechanism of Action)	Suppression of STAT3 → Down-regulation of chemokines/Decrease of DAMP → regulation of neutrophil migration/induction of bacterial clearance
Differentiation Point	First in Class-The drug can maintain blood levels of neutrophils with no direct impact on bone marrow functions, especially in the condition that homeostasis imbalance happens
Current Development Stage	Phase II
Route of Administration	Oral (Softgel Capsule)

Data

EC-18 inhibits blood neutrophils extravasation in CIN mouse model



EC-18 ameliorates oral mucositis in CRIOM-induced Model



Patent Position

10-2016-7035244/Korea, PA17-67/Japan

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Pre-Clinical Toxicology Study & Clinical IND approval for the development of idiopathic pulmonary fibrosis using SAMiRNA

76

KOREA DRUG DEVELOPMENT FUND

Bioneer Coporation

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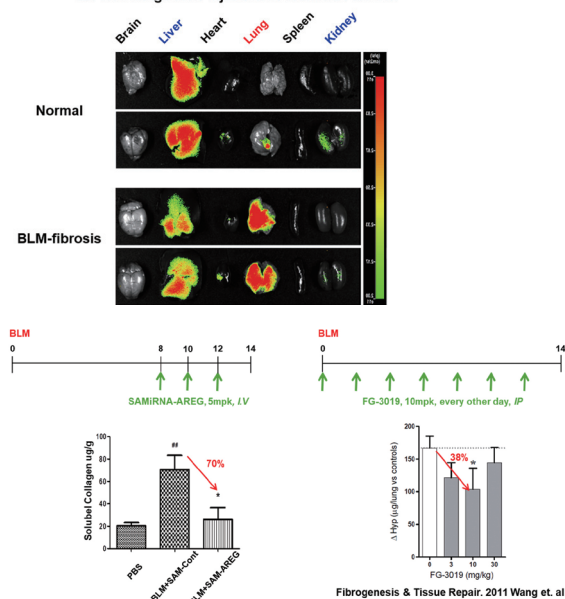
Product Type	RNAi Nanoparticles
Indication	1st indication: Idiopathic Pulmonary Fibrosis, Respiratory Tract Diseases (MeSH term)
Target	Amphiregulin
MoA (Mechanism of Action)	Amphiregulin: highly expressed downstream genes by TGF- β signaling in lung Fibroblast cells SAMiRNA-AREG: inhibit the expression of AREG \rightarrow block the differentiation of fibroblast to myofibroblast \rightarrow inhibit proloferation, secretion of ECM in the lung tissues
Differentiation Point	Esbriet® (Pirfenidone) and Ofev® (Nintedanib), which are approved by FDA in 2014, is expensive and exhibits limited efficacy. Urgent need for the development of new effective drug which can actively reverse the process of fibrosis. in vivo efficacy test: The efficacy of the SAMiRNA-IPF drug has been evaluated with TGF- β transgenic & Bleomycin induced mouse models of pulmonary fibrosis. Based on these comprehensive evaluations, the candidates show the more significant therapeutic potential for IPF treatment than Pirfenidone of InterMune, Inc and FG-3019 (CTGF mAb) of Fibrogen, Inc.

Current Development Stage Pre-Clinical

Route of Administration Parenteral-Intravenous

Data In vivo biodistribution of SAMiRNA by I.V inj. in BLM-induced IPF models

Ex-vivo image after injection of SAMiRNA for 48h



SAMiRNA by I.V injection shows collagen clearance efficacy compare to FG-3019 targeting proteins with much less doses and frequency

Patent Position

PCT/KR2010/003039, PCT/KR2014/006031, PCT/KR2014/006033

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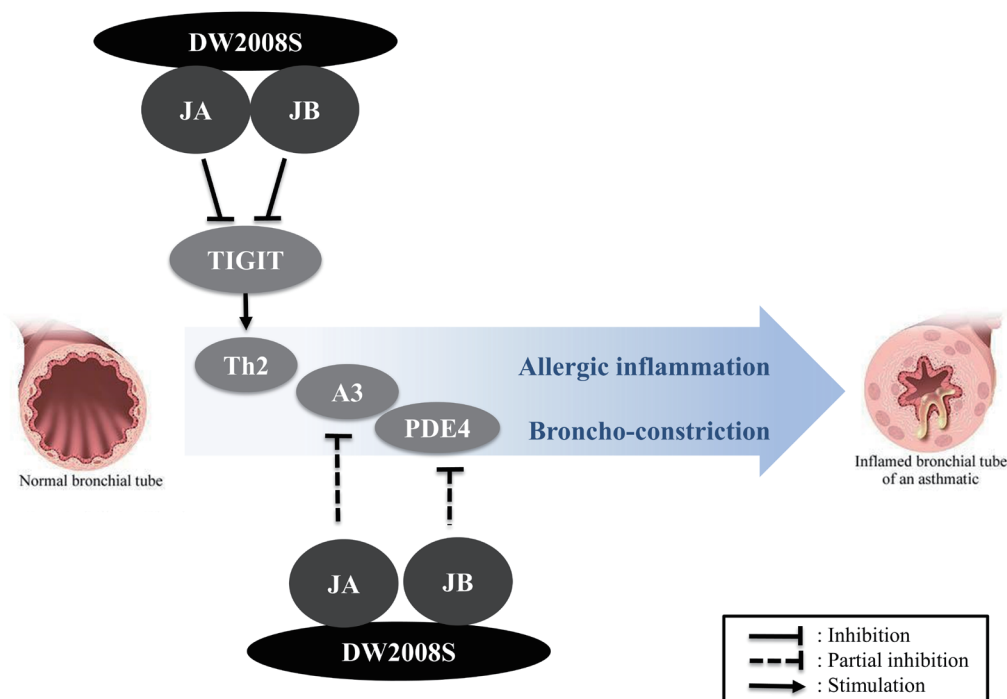
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Development of Anti-Allergic asthma agents

Dong-Wha Pharmaceutical



Product Type	Botanical drug
Indication	1st indication: Asthma, Respiratory Tract Diseases (MeSH term)
Target	TIGIT, PDE4 (partial), Adenosine Receptor 3 (A3AR, partial)
MoA (Mechanism of Action)	Th2 selective inhibition & Bronchodilation
Differentiation Point	First In Class
Current Development Stage	Phase I
Route of Administration	Oral
Data	1. Safety: NOAEL - SD Rat: 2,000 mg/kg, Beagle: 250 mg/kg → Safety Margin (H.E.D) : approximately 80 fold 2. Efficacy (Animal) : Superior to Montelukast



Patent Position 10-1747139/Korea

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Development of anti-scarring therapeutics OLX101 (previously BMT101), self-delivering RNAi molecule

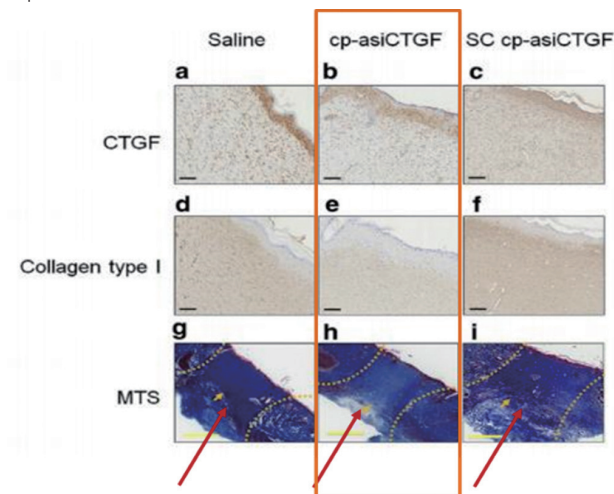
78

KOREA DRUG DEVELOPMENT FUND

OliX Pharmaceuticals, Inc.



Product Type	Cell-penetrating asymmetric siRNA
Indication	1st indication: Hypertrophic Scar, Cicatrix, Hypertrophic (MeSH term)
Target	Connective tissue growth factor (CTGF)
MoA (Mechanism of Action)	RNAi mediated repression of target gene CTGF which is a key factor in pathogenesis of fibrotic diseases including hypertrophic scra
Differentiation Point	First In Class
Current Development Stage	Phase I
Route of Administration	Local (Intradermal injection)
Data	Therapeutic Efficacy in rat skin excision wound model - Reduced expression of both target gene (CTGF) and collagen by cp-asiCTGF (OLX10010) treatment. - Masson's trichrome staining (MTS) data also revealed fibrosis repressing effect by cp-asiCTGF treatment.



Major outcomes in Phase I study

- Completion of dosing and follow-up (N=24/8; OLX10010/placebo)
- No SAE (serious adverse event) until 10X of expected therapeutic dose
- Very limited systemic exposure up to cohort 2 (\leftarrow LLOQ (2 ng/ml), waiting for results in cohorts 3 and 4)
- Mild local erythema in OLX10010 treated subjects

Patent Position

PCT/KR2008/007530, PCT/KR2011/006632, PCT/KR2013/004463

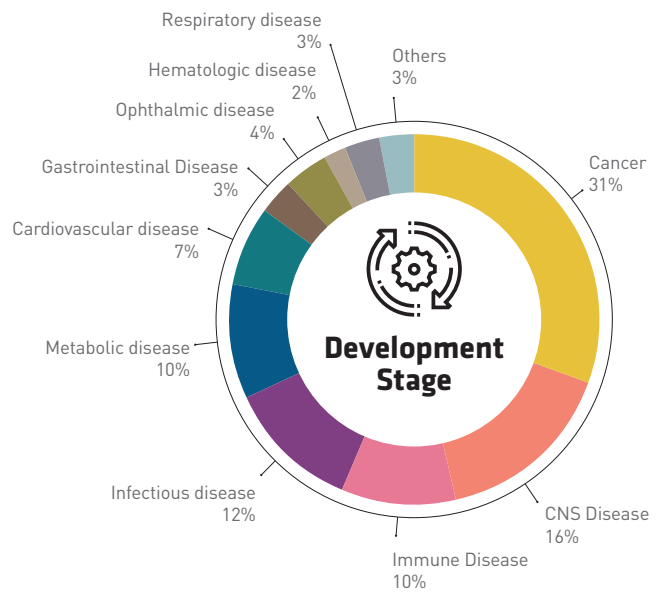
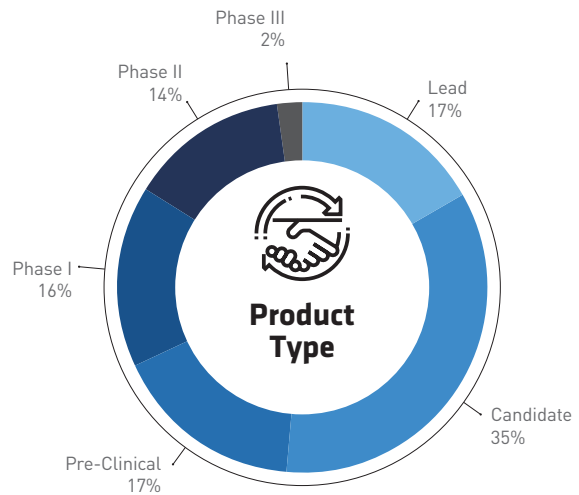
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R&D PIPELINE OVERVIEW

KDDF covers broad range of drug development fields.





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