

# Chi-Med and AstraZeneca's Savolitinib Shows Encouraging Clinical Activity in Second-Line EGFR Mutation-Positive Lung Cancer with *MET*-Amplification

– Data for savolitinib in combination with Tagrisso<sup>®</sup> or Iressa<sup>®</sup> presented at World Conference on Lung Cancer –

### – New data give insights into disease progression and potential next-generation treatment strategies in patients with EGFR-mutated NSCLC with MET amplification –

London: Tuesday, October 17, 2017: Hutchison China MediTech Limited ("Chi-Med") (AIM/Nasdaq: HCM) and AstraZeneca PLC ("AstraZeneca") today presented preliminary safety and clinical activity data of savolitinib when given in combination with either Tagrisso<sup>®</sup> (osimertinib) or Iressa<sup>®</sup> (gefitinib) in two Phase Ib/II proof-of-concept trials conducted in patients with epidermal growth factor receptor ("EGFR") mutation-positive ("EGFRm") non-small cell lung cancer ("NSCLC") with *MET*-amplification who had progressed following first-line treatment with an EGFR inhibitor. In both trials, the addition of savolitinib (600mg, once daily), an investigational selective inhibitor of mesenchymal epithelial transition factor ("c-MET") receptor tyrosine kinase, to Tagrisso<sup>®</sup> (80mg, once daily) or Iressa<sup>®</sup> (250mg, once daily) demonstrated preliminary anti-tumor activity. The data were shared in two oral presentations at the International Association for the Study of Lung Cancer 18<sup>th</sup> World Conference on Lung Cancer ("WCLC") in Yokohama, Japan, October 15 to 18, 2017.

Dr. Myung-Ju Ahn, Department of Hematology & Oncology, Samsung Medical Center, Seoul, South Korea, said, "Secondary resistance mechanisms often emerge during treatment with mutation-targeted medicines, leading to disease progression. The data presented at WCLC demonstrate the potential of utilizing savolitinib in c-MET-driven lung cancers to address resistance challenges."

Susan Galbraith, Head of Oncology, AstraZeneca Research and Early Development, said, "We are committed to developing innovative medicines to overcome the key drivers of cancer mechanisms of resistance and are strategically focused on developing effective combinations. The latest results for savolitinib in combination with osimertinib and gefitinib support our approach in collaboration with Chi-Med."

# Preliminary Results for Savolitinib in Combination with Tagrisso<sup>®[1]</sup>

In the Phase Ib/II proof-of-concept TATTON trial in patients with EGFRm advanced NSCLC with *MET*amplification confirmed locally or centrally, early data on safety and anti-tumor activity for savolitinib in combination with Tagrisso<sup>®</sup> were presented. In 66 patients treated, the most common all-causality adverse events ("AEs") were nausea (44%), vomiting (35%), fatigue (30%), and decreased appetite (30%), which were consistent with the known safety profiles of savolitinib and Tagrisso<sup>®</sup>.

Preliminary data showed partial response according to Response Evaluation Criteria in Solid Tumors ("RECIST") 1.1 criteria in 33% of patients previously treated with third-generation T790M-directed EGFR inhibitors, including Tagrisso<sup>®</sup> (n=30). In patients who had progressed after prior treatment with a first- or second-generation EGFR inhibitor, 61% of T790M mutation negative patients (n=23) had a partial response, while 55% of T790M mutation positive patients (n=11) had a partial response.

In those patients where *MET*-positive status was determined centrally, preliminary data showed partial response in 28% of patients previously treated with T790M-directed EGFR inhibitors (n=25). In patients who had progressed after prior treatment with a first- or second-generation EGFR inhibitor, 53% of T790M mutation negative patients (n=15) had a partial response, while 57% of T790M mutation positive patients (n=7) had a partial response.

The presentation is available at www.chi-med.com/ph2-savo-plus-tagrisso-nsclc/.

# Preliminary data for Savolitinib in Combination with Iressa®[2]

Data from the Phase Ib/II proof-of-concept trial assessing savolitinib in combination with Iressa<sup>®</sup> in patients in China with EGFRm advanced NSCLC with centrally confirmed *MET*-amplification who had progressed following EGFR inhibitor therapy were also reported. The most common AEs independent of causality in 51 patients treated were vomiting (39%), increased alanine aminotransferase (ALT) (37%), increased aspartate

aminotransferase (AST) (35%), nausea (35%), and rash (35%). These results were consistent with the known safety profiles of savolitinib and Iressa<sup>®</sup>.

Preliminary results showed that 31% of patients had a partial response according to RECIST 1.1 criteria, of which 52% of T790M negative patients (n=23) and 9% of T790M positive patients (n=23) achieved a partial response.

The presentation is available at www.chi-med.com/ph2-savolitinib-plus-iressa-nsclc/.

Mr. Christian Hogg, Chief Executive Officer of Chi-Med, said, "*MET*-amplification impacts a meaningful proportion of patients with EGFRm NSCLC who experience disease progression following treatment with a tyrosine kinase inhibitor in the first or second-line setting. Among patients with this difficult-to-treat resistance mechanism, there is a clear unmet medical need."

## About Savolitinib

Savolitinib (AZD6094/HMPL-504) is a potential first-in-class selective inhibitor of c-MET (also known as mesenchymal epithelial transition factor) receptor tyrosine kinase, an enzyme which has been shown to function abnormally in many types of solid tumors. It was developed as a potent and highly selective oral inhibitor specifically designed to address issues observed in the clinic with other selective c-MET inhibitors, such as renal toxicity.

Savolitinib was discovered by Chi-Med and is being developed in collaboration with AstraZeneca. Savolitinib is currently being studied in multiple tumor types worldwide including kidney, lung and gastric cancers, both as a monotherapy and in combination with other targeted and immunotherapy agents.

## About Chi-Med

Chi-Med is an innovative biopharmaceutical company which researches, develops, manufactures and sells pharmaceuticals and healthcare products. Its Innovation Platform, Hutchison MediPharma Limited, focuses on discovering and developing innovative therapeutics in oncology and autoimmune diseases for the global market. Its Commercial Platform manufactures, markets, and distributes prescription drugs and consumer health products in China.

Chi-Med is majority owned by the multinational conglomerate CK Hutchison Holdings Limited (SEHK: 0001). For more information, please visit: <u>www.chi-med.com</u>.

#### About AstraZeneca

AstraZeneca is a global, science-led biopharmaceutical company that focuses on the discovery, development and commercialization of prescription medicines, primarily for the treatment of diseases in three main therapy areas - Oncology, Cardiovascular & Metabolic Diseases and Respiratory. The Company also is selectively active in the areas of autoimmunity, neuroscience and infection. AstraZeneca operates in over 100 countries and its innovative medicines are used by millions of patients worldwide.

For more information, please visit <u>www.astrazeneca.com</u> and on Twitter @AstraZeneca.

Tagrisso<sup>®</sup> and Iressa<sup>®</sup> are trademarks of the AstraZeneca PLC group of companies.

#### **Forward-Looking Statements**

This announcement contains forward-looking statements within the meaning of the "safe harbor" provisions of the U.S. Private Securities Litigation Reform Act of 1995. These forward-looking statements reflect Chi-Med's current expectations regarding future events, including its expectations for the clinical development of savolitinib, plans to initiate clinical studies for savolitinib, its expectations as to whether such studies would meet their primary or secondary endpoints, and its expectations as to the timing of the completion and the release of results from such studies. Forward-looking statements involve risks and uncertainties. Such risks and uncertainties include, among other things, assumptions regarding enrollment rates, timing and availability of subjects meeting a study's inclusion and exclusion criteria, changes to clinical protocols or regulatory requirements, unexpected adverse events or safety issues, the ability of the drug candidate savolitinib to meet

the primary or secondary endpoint of a study, to obtain regulatory approval in different jurisdictions, to gain commercial acceptance after obtaining regulatory approval, the potential market of savolitinib for a targeted indication and the sufficiency of funding. In addition, as certain studies rely on the use of Tagrisso<sup>®</sup> or Iressa<sup>®</sup> as a combination therapeutic with savolitinib, such risks and uncertainties include assumptions regarding the safety, efficacy, supply and continued regulatory approval of Iressa<sup>®</sup> and Tagrisso<sup>®</sup>. Existing and prospective investors are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof. For further discussion of these and other risks, see Chi-Med's filings with the U.S. Securities and Exchange Commission and on AIM. Chi-Med undertakes no obligation to update or revise the information contained in this announcement, whether as a result of new information, future events or circumstances or otherwise.

### **Inside Information**

This announcement contains inside information for the purposes of Article 7 of Regulation (EU) No 596/2014.

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#### References

<sup>1</sup> Ahn M-J, *et al.* TATTON Phase Ib Expansion Cohort: Osimertinib Plus Savolitinib for Patients with EGFR-mutant MET-amplified NSCLC After Progression on Prior EGFR-TKI. Abstract #8985. Presented at the World Lung Cancer Congress 2017, Yokohama, Japan, 15-18 October 2017.

<sup>2</sup> Yang J-J, *et al.* A Phase Ib Trial of Savolitinib Plus Gefitinib for Patients with EGFR-mutant MET-amplified Advanced NSCLC. Abstract #8995. Presented at the World Lung Cancer Congress 2017, Yokohama, Japan, 15-18 October 2017.