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Press release

Basilea reports derazantinib/PD-L1 checkpoint inhibitor combination results from dose-finding part of FIDES-02 study in patients with solid tumors at ASCO GU symposium

Basel, Switzerland, February 12, 2021

Basilea Pharmaceutica Ltd. (SIX: BSLN) announced today that data on the safety, tolerability and preliminary efficacy of the fibroblast growth factor receptor (FGFR) inhibitor derazantinib in combination with the PD-L1 checkpoint inhibitor atezolizumab, in patients with advanced solid tumors from the phase 1b dose-finding cohort in the FIDES-02 study, were presented at the virtual American Society of Clinical Oncology Genitourinary Cancers Symposium (ASCO GU 2021), held from February 11 to 13, 2021.[1]

The derazantinib-atezolizumab combination was well tolerated and no dose-limiting toxicities were observed. The most commonly reported adverse events were fatigue/asthenia (weakness), nausea and diarrhea. The study demonstrated that derazantinib and atezolizumab can be safely combined at doses of 300 mg of daily oral derazantinib, which is the derazantinib monotherapy dose used in the phase 2 study FIDES-01, and 1200 mg atezolizumab, administered intravenously once every three weeks which was the maximum dose pre-specified in the protocol.[2]

Several patients are still receiving treatment with derazantinib and atezolizumab, including one patient with a bile duct cancer and FGFR2 gene fusion, who is ongoing in the study for more than nine months and was reported with a partial response with continued tumor shrinkage. The combination of derazantinib and atezolizumab is supported by preclinical data demonstrating derazantinib's strong inhibition of colony-stimulating-factor-1-receptor kinase (CSF1R). CSF1R inhibition may revert tumor-induced immunosuppression and thereby enhance the response to immune-checkpoint inhibition.

Dr. Marc Engelhardt, Chief Medical Officer, said: "The recently announced clinical proof of concept for derazantinib as monotherapy in the FIDES-01 study in patients with FGFR2 gene fusion-positive intrahepatic cholangiocarcinoma and the efficacy signal obtained in the group of patients with advanced solid tumors in the FIDES-02 study are encouraging. Considering the manageable safety and tolerability profile and the rapidly evolving treatment landscape in urothelial cancer in patients both with and without FGFR genetic aberrations, we additionally plan to amend the FIDES-02 study protocol to explore a higher dose of derazantinib in two cohorts of this study. These cohorts would explore whether dosing derazantinib at its previously

established maximum tolerated daily dose, which is about 30% higher than the current phase 2 dose, may provide additional benefits in monotherapy and combination to patients with FGFR-positive urothelial cancer."

The FIDES-02 study (**F**ibroblast growth factor **I**nhibition with **DE**razantinib in **S**olid tumors) is an ongoing multi-center, open-label phase 1/2 study evaluating once-daily orally administered derazantinib and atezolizumab in patients with inoperable or metastatic urothelial (bladder) cancer and confirmed FGFR genetic aberrations.

The following e-poster was presented at ASCO GU 2021:

Abstract #	Title
437	Derazantinib (DZB) in combination with atezolizumab (AZB) in patients with solid tumors: Results from the dose-finding phase Ib substudy of FIDES-02.
	Authors: Raghad Karim, Arvind Chaudhry, Anna Patrikidou, Alejandro Falcon Gonzalez, Fabricio Racca, Yohann Loriot, Damien Pouessel, Jean-Laurent Deville, Shinkyo Yoon, Hyo Jin Lee, Frederique Cantero, Michalina Marszewska, Mikael Saulay, Stephan Braun, Rodryg Ramlau

For further information, please visit https://meetings.asco.org/gu/virtual-program

About derazantinib

Derazantinib is an investigational orally administered small-molecule FGFR inhibitor with strong activity against FGFR1, 2, and 3.[3] FGFR kinases are key drivers of cell proliferation, differentiation and migration. FGFR genetic aberrations, e.g. gene fusions, mutations or amplifications, have been identified as potentially important therapeutic targets for various cancers, including intrahepatic cholangiocarcinoma (iCCA), urothelial, breast, gastric and lung cancers.[4] In these cancers, FGFR genetic aberrations are found in a range of 5% to 30%.[5] Derazantinib also inhibits the colony-stimulating-factor-1-receptor kinase (CSF1R).[3, 6] CSF1R-mediated signaling is important for the maintenance of tumor-promoting macrophages and therefore has been identified as a potential target for anti-cancer drugs.[7] Pre-clinical data has shown that tumor macrophage depletion through CSF1R blockade renders tumors more responsive to T-cell checkpoint immunotherapy, including approaches targeting PD-L1/PD-1.[8, 9] Derazantinib has demonstrated initial antitumor activity and a manageable safety profile in a previous biomarker-driven phase 1/2 study in iCCA patients,[10] and has received U.S. and EU orphan drug designation for iCCA. Basilea is currently conducting three clinical studies with derazantinib. The first study, FIDES-01, is a registrational phase 2 study in patients with inoperable or advanced iCCA. It comprises one cohort of patients with FGFR2 gene fusions and another cohort of patients with mutations or amplifications.[2] Positive topline results for the FIDES-01 cohort with FGFR2 gene-fusion positive iCCA patients have just been reported. The second study, FIDES-02, is a phase 1/2 study evaluating derazantinib alone and in combination with Roche's PD-L1 checkpoint inhibitor, atezolizumab, in patients with advanced urothelial cancer, including metastatic, or recurrent surgically unresectable disease, expressing FGFR

genetic aberrations.[1] The third study, FIDES-03, is a phase 1/2 study evaluating derazantinib alone and in combination with Lilly's anti-VEGFR2 antibody ramucirumab and paclitaxel or with Roche's PD-L1 checkpoint inhibitor atezolizumab in patients with advanced gastric cancer with FGFR genetic aberrations.[11] Basilea in-licensed derazantinib from ArQule Inc., a wholly-owned subsidiary of Merck & Co., Inc., Kenilworth, N.J., U.S.A.

About Basilea

Basilea Pharmaceutica Ltd. is a commercial-stage biopharmaceutical company, focused on the development of products that address the medical challenges in the therapeutic areas of oncology and infectious diseases. With two commercialized drugs, the company is committed to discovering, developing and commercializing innovative pharmaceutical products to meet the medical needs of patients with serious and life-threatening conditions. Basilea Pharmaceutica Ltd. is headquartered in Basel, Switzerland and listed on the SIX Swiss Exchange (SIX: BSLN). Additional information can be found at Basilea's website www.basilea.com.

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This press release can be downloaded from www.basilea.com.

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