



National Securities Market Commission
Markets Directorate General
c/ Edison núm. 4
28006 Madrid

Colmenar Viejo (Madrid), June 12th, 2018

Pursuant to article 228 of the consolidated text of the Spanish Securities Market Act, we hereby inform you of the following **SIGNIFICANT EVENT**:

“Pharma Mar announced today that based on recent receipt of OS (overall survival) data from lurbinectedin Phase II small-cell lung cancer studies, including the monotherapy trial presented at ASCO on June 3rd that saw an OS of 11.8 months, a protocol amendment was submitted to FDA and other competent authorities to change the primary endpoint of the ATLANTIS Phase III trial from PFS (progression free survival) to OS. The changes will begin when the competent authorities with responsibility for review and approval of the study approve of the changes, which in the US is expected to happen in the next few weeks. The safety of the patients and the integrity of this study is not compromised by these changes. PharmaMar continues to expect completion of recruitment in this trial in the third quarter of 2018. This means PharmaMar expects the top line data to readout in the second half of 2019. In this regard please find attached press release that Pharma Mar, S.A. will distribute to the media today”



PharmaMar requests the modification from primary endpoint to OS for the ATLANTIS trial

Madrid, 12th of June 2018. - PharmaMar (MSE:PHM) announced today that based on recent receipt of OS (overall survival) data from lurbinectedin Phase II small-cell lung cancer studies, including the monotherapy trial presented at ASCO on June 3rd that saw an OS of 11.8 months, a protocol amendment was submitted to FDA and other competent authorities to change the primary endpoint of the ATLANTIS Phase III trial from PFS (progression free survival) to OS. The changes will begin when the competent authorities with responsibility for review and approval of the study approve of the changes, which in the US we expect to happen in the next few weeks. The safety of the patients and the integrity of this study is not compromised by these changes. PharmaMar remains blinded to the data, and continues to expect completion of recruitment in the third quarter of 2018. This means PharmaMar expects the top line data, which is event driven, to readout in the second half of 2019.

According to Luis Mora, Managing Director of PharmaMar´s Oncology Business Unit, *"we feel that this change in endpoint to OS given what we have seen in the recent data, including those presented at ASCO, offers us a better chance for success, especially as we know regulators prefer OS data over a surrogate endpoint subject to interpretation in this type of disease setting."*

About Zepsyre®

Zepsyre® (lurbinectedin, PM1183) is a compound under clinical investigation. It is an inhibitor of RNA polymerase II. This enzyme is essential for the transcription process that is over-activated in tumors with transcription addiction.

About small-cell lung cancer

SCLC is a very aggressive cancer that usually presents with distant metastases and has already spread at the time of diagnosis, thus limiting the role of traditional approaches and posing a worse prognosis compared to other lung cancer types. The 5-year survival rate is about 5%ⁱ. About 18% of all the lung cancer cases diagnosed are SCLC, and only in the US more than 34,000 new cases are recorded every year. This tumor is strongly associated with tobacco smoking, posing an important public health problemⁱⁱ. After failure to treatment with a platinum-based therapy in first line, the therapeutic alternatives are very limited, and the approval of the last drug for this disease took place 20 years ago.

About PharmaMar

Headquartered in Madrid, PharmaMar is a world-leading biopharmaceutical company in the discovery and development of innovative marine-derived anticancer drugs. The company has a pipeline of drug

candidates and a robust R&D oncology program. PharmaMar develops and commercializes YONDELIS® in Europe and has other clinical-stage programs under development for several types of solid and hematological cancers, Zepsyre® (PM1183), plitidepsin, PM184 and PM14. PharmaMar is a global biopharmaceutical company with subsidiaries in Germany, Italy, France, Switzerland, United Kingdom, Belgium, Austria and the United States. PharmaMar fully owns other companies: GENOMICA, a leading molecular diagnostics company; Sylentis, dedicated to researching therapeutic applications of gene silencing (RNAi); and two other chemical enterprises, Zelnova Zeltia and Xylazel. To learn more about PharmaMar, please visit us at www.pharmamar.com.

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ⁱ <http://www.cancer.gov/types/lung/hp/small-cell-lung-treatment-pdq>

ⁱⁱ <http://www.jnccn.org/content/11/1/78.full.pdf>