PRESS RELEASE

This press release is for U.S. audiences only

Blue Earth Diagnostics, a Bracco Company, Announces Results Evaluating Diagnostic Performance of POSLUMA® (Flotufolastat F 18) and Clinical Trial Enrollment in African American Men with Prostate Cancer

Post-hoc analysis from Phase 3 SPOTLIGHT trial demonstrated consistent performance between African Americans and other patient groups for FDA-approved POSLUMA in men with recurrent prostate cancer[1]

Study achieved 17% African American enrollment, more than twice the average for oncology clinical trials[2]

MONROE TOWNSHIP, N.J., and OXFORD, UK, July 30, 2024 – Blue Earth Diagnostics, a Bracco company and recognized leader in the development and commercialization of innovative PET radiopharmaceuticals, today announced results of a post-hoc analysis of data from the Phase 3 SPOTLIGHT study (NCT04186845), which assessed FDA-approved POSLUMA® (flotufolastat F 18) injection in patients with recurrent prostate cancer.[3,4] Based on the high mortality and prevalence of prostate cancer in African American men, a sub-analysis was conducted to evaluate the performance of POSLUMA and the rate of enrollment for African American men in the trial. Results showed that the detection rate was high among African American patients, with 93% found to have a positive POSLUMA scan, consistent with the 87% detection rate for all other patients in the study.[1] The 17% participation of African American men in the SPOTLIGHT study was twice the enrollment typically reported in other types of oncology clinical trials (8.5%).[1,2] The manuscript, "^{18}F-Flotufolastat Positron Emission Tomography in African American Patients with Suspected Prostate Cancer Recurrence: Findings from the Phase 3 SPOTLIGHT Study" has been published online in the journal *Advances in Radiation Oncology* (https://doi.org/10.1016/j.adro.2024.101571), and will appear in an upcoming print issue.

POSLUMA is approved in the United States for positron emission tomography (PET) of prostate-specific membrane antigen (PSMA) positive lesions in men with prostate cancer with suspected metastasis who are candidates for initial definitive therapy or with suspected recurrence based on elevated serum prostate-specific antigen (PSA) level.

"Prostate cancer is the most commonly diagnosed cancer among African American men, who are twice more likely to die from the disease than White men,"[5-7] said Soroush Rais-Bahrami, MD, Department of Urology, University of Alabama at Birmingham Heersink School of Medicine, Birmingham, Ala. "It is encouraging that African American enrollment in the SPOTLIGHT trial closely aligns with their 14% representation of the U.S. population[8], because results from oncology trials with low diversity populations are less useful for clinical decision-making and can contribute to racial disparities in cancer outcomes."

The U.S. Food and Drug Administration (FDA) has issued draft guidance on clinical trial diversity and organizations such as the American Society of Clinical Oncology (ASCO) have called out the need for enriched diversity in oncology clinical trial participation.[9,10]

"Beyond validating the diagnostic performance of POSLUMA in African American men, findings from the SPOTLIGHT study provide useful considerations for planning future clinical trials, to facilitate patient enrollment and achieve clinical trial diversity," said Marco Campione, Chief Executive Officer, Blue Earth Diagnostics. "The results achieved in African American men were derived from clinical sites across the United States, and further support the broad applicability of POSLUMA for its indicated use across the U.S. population as a whole."

FDA-approved POSLUMA represents a newer class of high-affinity PSMA-targeted PET radiopharmaceuticals based on novel radiohybrid technology and leverages the high image quality of ^{18}F-labeled PSMA PET imaging to help facilitate detection of prostate cancer. It can provide clinically valuable information to guide patient management based on its high-affinity PSMA binding, low urinary uptake and positive performance at very low PSA levels. POSLUMA is included in nationally recognized clinical oncology guidelines for prostate cancer, covered by the vast majority of insurance plans and is readily available for patients across the country through the network of Blue Earth Diagnostics' commercial U.S. manufacturer and distributor, PETNET Solutions Inc, A Siemens Healthineers Company.

Indication and Important Safety Information About POSLUMA

INDICATION

POSLUMA® (flotufolastat F 18) injection is indicated for positron emission tomography (PET) of prostate-specific membrane antigen (PSMA) positive lesions in men with prostate cancer

- with suspected metastasis who are candidates for initial definitive therapy
- with suspected recurrence based on elevated serum prostate-specific antigen (PSA) level

IMPORTANT SAFETY INFORMATION

- Image interpretation errors can occur with POSLUMA PET. A negative image does not rule out the presence of prostate cancer and a positive image does not confirm the presence of prostate cancer. The performance of POSLUMA for imaging metastatic pelvic lymph nodes in patients prior to initial definitive therapy seems to be affected by serum PSA levels and risk grouping. The performance of POSLUMA for imaging patients with biochemical evidence of recurrence of prostate cancer seems to be affected by serum PSA levels. Flotufolastat F 18 uptake is not specific for prostate cancer and may occur in other types of cancer, in non-malignant processes, and in normal tissues. Clinical correlation, which may include histopathological evaluation, is recommended.
- Risk of Image Misinterpretation in Patients with Suspected Prostate Cancer Recurrence: The interpretation of POSLUMA PET may differ depending on imaging readers, particularly in the prostate/prostate bed region. Because of the associated risk of false positive interpretation, consider multidisciplinary consultation and histopathological confirmation when clinical decision-making hinges on flotufolastat F 18 uptake only in the prostate/prostate bed region or only on uptake interpreted as borderline.
- POSLUMA use contributes to a patient's overall long-term cumulative radiation exposure. Long-term cumulative radiation exposure is associated with an increased risk for cancer. Advise

patients to hydrate before and after administration and to void frequently after administration. Ensure safe handling to minimize radiation exposure to the patient and health care providers.
- The adverse reactions reported in ≥0.4% of patients in clinical studies were diarrhea, blood pressure increase and injection site pain.
- Drug Interactions: androgen deprivation therapy (ADT) and other therapies targeting the androgen pathway, such as androgen receptor antagonists, may result in changes in uptake of flotufolastat F 18 in prostate cancer. The effect of these therapies on performance of POSLUMA PET has not been established.

To report suspected adverse reactions to POSLUMA, call 1-844-POSLUMA (1-844-767-5862) or contact FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

Full POSLUMA prescribing information is available at www.posluma.com/prescribing-information.pdf.

About Blue Earth Diagnostics

Blue Earth Diagnostics, an indirect subsidiary of Bracco Imaging S.p.A., is a growing international molecular imaging company focused on delivering innovative, well-differentiated diagnostic solutions that inform patient care. Formed in 2014, the Company's success is driven by its management expertise and supported by a demonstrated track record of rapid development and commercialization of positron emission tomography (PET) radiopharmaceuticals. Blue Earth Diagnostics' expanding oncology portfolio encompasses a variety of disease states, including prostate cancer and neuro-oncology. Blue Earth Diagnostics is committed to the timely development and commercialization of precision radiopharmaceuticals for potential use in imaging and therapy. For more information, please visit: www.blueearthdiagnostics.com.

About Bracco Imaging

Bracco Imaging S.p.A., part of the Bracco Group, is a world-leading diagnostic imaging provider. Headquartered in Milan, Italy, Bracco Imaging develops, manufactures and markets diagnostic imaging agents and solutions. It offers a product and solution portfolio for all key diagnostic imaging modalities: X-ray imaging (including Computed Tomography-CT, Interventional Radiology, and Cardiac Catheterization), Magnetic Resonance Imaging (MRI), Contrast Enhanced Ultrasound (CEUS), and Nuclear Medicine through radioactive tracers and novel PET imaging agents to inform clinical management and guide care for cancer patients in areas of unmet medical need. Our continually evolving portfolio is completed by a range of medical devices, advanced administration systems and dose-management software. In 2019 Bracco Imaging enriched its product portfolio by expanding the range of oncology nuclear imaging solutions in the urology segment and other specialties with the acquisition of Blue Earth Diagnostics. In 2021, Bracco Imaging established Blue Earth Therapeutics as a separate, cutting-edge biotechnology vehicle to develop radiopharmaceutical therapies. Visit: www.braccoimaging.com.

References

1. Rais-Bahrami S, Fleming M, Gartrell B, et al. ^{18}F-Flotufolastat Positron Emission Tomography in African American Patients with Suspected Prostate Cancer Recurrence: Findings from the Phase 3 SPOTLIGHT Study. *Adv Clin Onc* 2024; doi 10.1016/j.adro.2024.101571
2. Bebi T, Horovitz R, Blum K, et al. How granularity of data matters in understanding and accelerating racial diversity in U.S. clinical trials. *J Clin Oncol* 2022;40 (Suppl 28):88.
3. Jani AB, Ravizzini G, Gartrell BA, et al. Diagnostic performance and safety of ^{18}F-rhPSMA-7.3 PET in men with suspected prostate cancer recurrence: Results from a phase 3, prospective, multicenter study (SPOTLIGHT) *J Urol* 2023;210:299-311.
4. FDA. Highlights of prescribing information: Posluma (flotufolastat f 18) injection: https://www.accessdata.fda.gov/drugsatfda_docs/label/2023/216023s000lbl.pdf; 2023.
5. Giaquinto AN, Miller KD, Tossas KY, et al. Cancer statistics for African American/Black people 2022. *CA Cancer J Clin* 2022;72:202-229.

6. Frego N, Labban M, Stone BV, et al. Effect of type of definitive treatment on race- based differences in prostate cancer-specific survival. *Prostate* 2023;83:1099-1111.
7. Rais-Bahrami S, Zhu Y. Disparities in prostate cancer diagnosis and management: Recognizing that disparities exist at all junctures along the prostate cancer journey. *Prostate cancer and prostatic diseases* 2023;26:441-442.
8. Facts About the U.S. Black Population. Pew Research Center. Fact Sheet. January 14, 2024. https://www.pewresearch.org/social-trends/fact-sheet/facts-about-the-us-black-population/
9. FDA. Diversity Action Plans to Improve Enrollment of Participants from Underrepresented Populations in Clinical Studies *https://www.fda.gov/media/179593/download*; 2024.
10. Oyer RA, Hurley P, Boehmer L, et al. Increasing racial and ethnic diversity in cancer clinical trials: An American Society of Clinical Oncology and Association of Community Cancer Centers joint research statement. *Journal of Clinical Oncology* 2022;40:2163-2171.

Contact:
For Blue Earth Diagnostics (U.S.)
Priscilla Harlan
Vice President, Corporate Communications
(M) (781) 799-7917
priscilla.harlan@blueearthdx.com

For Blue Earth Diagnostics (UK)
Clare Gidley
Associate Director Marketing and Communications
Tel: +44 (0)1865 784186
clare.gidley@blueearthdx.com

Media
Sam Brown Inc.
Mike Beyer
(M) (312) 961-2502
mikebeyer@sambrown.com

# # #

PP-rh-US-0523