FDA Approves Braftovi Plus Mektovi for Advanced Melanoma With BRAF Mutations

Encorafenib and binimetinib combination doubled progression-free survival in the Phase III COLUMBUS trial.

June 27, 2018 By Food and Drug Administration (FDA)

On June 27, 2018, the Food and Drug Administration approved encorafenib and binimetinib (Braftovi and Mektovi, Array BioPharma Inc.) in combination for patients with unresectable or metastatic melanoma with a BRAF V600E or V600K mutation, as detected by an FDA-approved test.

Approval was based on a randomized, active-controlled, open-label, multicenter trial (COLUMBUS; NCT01909453) in 577 patients with BRAF V600E or V600K mutation-positive unresectable or metastatic melanoma. Patients were randomized (1:1:1) to receive binimetinib 45 mg twice daily plus encorafenib 450 mg once daily, encorafenib 300 mg once daily, or vemurafenib 960 mg twice daily. Treatment continued until disease progression or unacceptable toxicity.

The major efficacy measure was progression-free survival (PFS) using RECIST 1.1 response criteria and assessed by blinded independent central review. The median PFS was 14.9 months for patients receiving binimetinib plus encorafenib, and 7.3 months for the vemurafenib monotherapy arm (hazard ratio 0.54, 95% CI: 0.41, 0.71, $p<0.0001$). Overall response rates assessed by central review were 63% and 40%, respectively. Median response duration was 16.6 months vs. 12.3 months, respectively.

The most common (≥25%) adverse reactions in patients receiving the combination were fatigue, nausea, diarrhea, vomiting, abdominal pain, and arthralgia. Discontinuation of therapy due to adverse reactions occurred in 5% of patients receiving the combination; the most common reasons were hemorrhage and headache.

FDA today also granted approval of the THxID BRAF Kit (bioMérieux) as a companion diagnostic for these therapeutics.

The recommended doses are binimetinib 45 mg orally twice daily and encorafenib 450 mg orally once daily.

View full prescribing information for BRAFTOVI.

View full prescribing information for MEKTOVI.

Healthcare professionals should report all serious adverse events suspected to be associated with the use of any medicine and device to FDA's MedWatch Reporting System or by calling 1-800-FDA-1088.

This announcement was originally published on the Food and Drug Administration website on June 27, 2018.
