

SmartPA Criteria Proposal

Drug/Drug Class:	Alpelisib Clinical Edit
First Implementation Date:	January 12, 2023
Proposed Date:	October 17, 2023
Prepared for:	MO HealthNet
Prepared by:	MO HealthNet/Conduent
Criteria Status:	<input checked="" type="checkbox"/> Existing Criteria <input type="checkbox"/> Revision of Existing Criteria <input type="checkbox"/> New Criteria

Executive Summary

Purpose: Ensure appropriate utilization and control of alpelisib agents.

Why Issue Selected: Alpelisib is a kinase inhibitor targeting phosphatidylinositol-3-kinase (PI3K) with inhibitory activity predominantly against PI3K α . Pathogenic variants in the gene encoding the catalytic α -subunit of PI3K lead to activation of PI3K α and Akt-signaling, cellular transformation and the generation of tumors in in vitro and in vivo models. Piqray[®] (alpelisib) was FDA-approved in 2019 in combination with fulvestrant for the treatment of postmenopausal women, and men, with hormone receptor (HR)-positive, human epidermal growth factor receptor 2 (HER2)-negative, PIK3CA-mutated, advanced or metastatic breast cancer. Vioice[®] (alpelisib) was FDA-approved on April 5, 2022, for the treatment of adult and pediatric patients 2 years of age and older with severe manifestations of *PIK3CA*-Related Overgrowth Spectrum (PROS) who require systemic therapy. PROS is a spectrum of rare disorders involving variants in the *PIK3CA* gene causing overgrowth in various parts of the body. These variants result in an abnormally active phosphatidylinositol-3-kinase (PI3K) enzyme, causing affected cells to grow and divide more than they should. The pathogenic variants in PROS are mosaic and are only present in certain body cells that affect certain areas. Manifestations include abnormal bone, soft tissue, and blood vessel growth in these affected areas. PROS has an estimated prevalence of 14 people per million. Treatment of PROS includes symptom management and treatment of complications such as bleeding, clotting, pain, and functional impairment. Vioice is the first FDA-approved medication for PROS. Piqray and Vioice both contain alpelisib as the active ingredient and differ solely in how they are supplied.

Due to the high cost, possible adverse events, and specific approved indications, MO HealthNet will impose clinical criteria to ensure appropriate utilization of alpelisib agents.

Program-Specific Information:

Date Range FFS 07/01/2022 to 06/30/2023			
Drug	Claims	Spend	Average Spend per Claim
PIQRAY 200 MG DOSE PACK	5	\$75,036.38	\$16,258.23
PIQRAY 250 MG DOSE PACK	6	\$111,496.08	
PIQRAY 300 MG DOSE PACK	11	\$167,032.13	
VIJOICE 50 MG TABLET	5	\$161,899.96	\$32,381.24
VIJOICE 125 MG TABLET	1	\$32,382.49	
VIJOICE 250 MG BLISTER PACK	0	-	

Type of Criteria: ☐ Increased risk of ADE
☒ Appropriate Indications

☐ Preferred Drug List
☒ Clinical Edit

Data Sources: ☐ Only Administrative Databases

☒ Databases + Prescriber-Supplied

Setting & Population

- Drug class for review: Alpelisib agents
- Age range: All appropriate MO HealthNet participants aged 2 years and older

Approval Criteria

Initial Therapy:

- Claim is for Piqray **OR**
- For Vioice tablets:
 - Prescribed by or in consultation with an appropriate specialist in the treated disease state **AND**
 - Participant is aged ≥ 2 years **AND**
 - Participants aged ≥ 18 years: therapeutic reason why Piqray 250 mg daily dose pack cannot be utilized **AND**
 - Documented diagnosis of PROS as verified by:
 - Genetic testing **OR**
 - National Institutes of Health (NIH) Workshop Diagnostic Guidelines **AND**
 - Documentation of at least one target lesion identified on imaging with baseline measurement of target lesion volume **OR**
 - Participants who are unable to complete baseline imaging must have at least one quantifiable target lesion able to be measured, requests will be subject to clinical review **AND**
 - Initial approval for 6 months

Continuation of Therapy:

- For Vioice:
 - Documentation of benefit of therapy by one of the following from baseline:
 - $\geq 20\%$ reduction in measurement of target lesion volume
 - Reduction in sum of lesion volume
 - Clinically meaningful improvement in signs and symptoms of disease (i.e., vascular malformation, functional improvement, limb asymmetry, pain)
 - Continued approval for 1 year

Denial Criteria

- Therapy will be denied if all approval criteria are not met
- For Vioice: participant is currently pregnant

Required Documentation

Laboratory Results: ☐
MedWatch Form: ☐

Progress Notes: ☐
Other: ☒

Disposition of Edit

Denial: Exception code "0682" (Clinical Edit)

SmartPA Clinical Proposal Form

© 2023 Conduent Business Services, LLC. All rights reserved. Conduent™ and Conduent Design™ are trademarks of Conduent Business Services, LLC in the United States and/or other countries.

Other company trademarks are also acknowledged.

Default Approval Period

6 months

References

- Vijoce® (alpelisib) [package insert]. East Hanover, New Jersey. Novartis Pharmaceuticals Corporation. Nov 2022.
- Piqray® (alpelisib) [package insert]. East Hanover, New Jersey. Novartis Pharmaceuticals Corporation. May 2022.
- IPD Analytics. Rx Insights: Endocrinology. Vijoce for *PIK3CA*-Related Overgrowth Spectrum (PROS). April 2022.
- National Organization for Rare Disorders. *PIK3CA*-Related Overgrowth Spectrum. Updated 2022. [PIK3CA-Related Overgrowth Spectrum - NORD \(National Organization for Rare Disorders\) \(rarediseases.org\)](https://rarediseases.org/rare-diseases/piqray/). Accessed May 11, 2022.
- Keppler-Noreuil KM, Rios JJ, Parker VE, et al. *PIK3CA*-related overgrowth spectrum (PROS): diagnostic and testing eligibility criteria, differential diagnosis, and evaluation. *Am J Med Genet A*. 2015;167A(2):287-295. doi:10.1002/ajmg.a.36836. [PIK3CA-Related Overgrowth Spectrum \(PROS\): Diagnostic and Testing Eligibility Criteria, Differential Diagnosis, and Evaluation - PMC \(nih.gov\)](https://pubmed.ncbi.nlm.nih.gov/26081111/)
- Pagliuzzi A, Oranges T, Traficante G, et al. *PIK3CA*-related overgrowth spectrum from diagnosis to targeted therapy: a case of CLOVES syndrome treated with alpelisib. *Front. Pediatr.*, 09 September 2021 | <https://doi.org/10.3389/fped.2021.732836>.
- Novartis Pharmaceuticals Corporation. *PIK3CA*-Related Overgrowth Spectrum (PROS). Updated April 2021. [PIK3CA-Related Overgrowth Spectrum \(PROS\) \(prospectum.com\)](https://prospectum.com/). Accessed May 11, 2022.
- Novartis Pharmaceuticals Corporation. VIOICE® (alpelisib) tablets for PROS Treatment. Updated March 2022. [VIOICE® \(alpelisib\) tablets for PROS Treatment | HCP \(novartis.com\)](https://www.novartis.com/usa/clinical-trials/phase-3/vioice-tablets-for-pros-treatment). Accessed May 11, 2022.
- Canaud G, Lopez Gutierrez JC, Irvine A, et al. EPIK-P1: Retrospective chart review study of patients with *PIK3CA*-related overgrowth spectrum who have received alpelisib as part of a compassionate use programme. Presented at the 2021 ESMO Virtual Congress; September 17–21, 2021. [ESMO Congress 2021 Speakers Template \(novartisoncology.com\)](https://www.esmo.org/2021/abstracts/2021-09-17/ESMO-Congress-2021-Speakers-Template). Accessed May 11, 2022.
- A Plain-Language Summary About the EPIK-P1 Study. [Canaud PlainLanguageSummay_LBA23.pdf \(novartisoncology.com\)](https://www.novartisoncology.com/press-releases/2021-09-17/ESMO-Congress-2021-Speakers-Template). Accessed May 11, 2022.
- NIH: U.S National Library of Medicine. Retrospective Chart Review Study of Patients With *PIK3CA*-Related Overgrowth Spectrum Who Have Received Alpelisib (EPIK-P1). [Retrospective Chart Review Study of Patients With PIK3CA-Related Overgrowth Spectrum Who Have Received Alpelisib - No Study Results Posted - ClinicalTrials.gov](https://www.clinicaltrials.gov/study/NCT04301111/). Accessed May 11, 2022.
- NIH: U.S National Library of Medicine. Study Assessing the Efficacy, Safety and PK of Alpelisib (BYL719) in Pediatric and Adult Patients With *PIK3CA*-related Overgrowth Spectrum. [Study Assessing the Efficacy, Safety and PK of Alpelisib \(BYL719\) in Pediatric and Adult Patients With PIK3CA-related Overgrowth Spectrum - Full Text View - ClinicalTrials.gov](https://www.clinicaltrials.gov/study/NCT04301111/). Accessed May 11, 2022.
- NIH: U.S National Library of Medicine. Study Assessing Long-term Safety and Efficacy of Alpelisib in Patients With *PIK3CA*-Related Overgrowth Spectrum (PROS) Who Previously Participated in Study CBYL719F12002 (EPIK-P1) (EPIK-P3). [Study Assessing Long-term Safety and Efficacy of Alpelisib in Patients With PIK3CA-Related Overgrowth Spectrum \(PROS\) Who Previously Participated in Study CBYL719F12002 \(EPIK-P1\) - Full Text View - ClinicalTrials.gov](https://www.clinicaltrials.gov/study/NCT04301111/). Accessed May 11, 2022.