

Inclusion

- Male or female patients aged ≥ 18 years (follow local regulatory requirement if legal age of consent > 18 years)
- Histologically or cytologically documented locally advanced or metastatic nonsquamous NSCLC not amenable to curative surgery or radiation
- Documentation of radiographic disease progression while on/after receiving a third-generation EGFR TKI for metastatic or locally advanced disease
- Documentation of an EGFR-activating mutation detected from tumor tissue or blood sample: exon 19 deletion or L858R
- ≥ 1 measurable lesion confirmed by investigator assessment as per RECIST v1.1
- 1 or 2 prior line(s) of an approved EGFR TKI treatment in the metastatic or locally advanced setting, which must include a third-generation EGFR TKI (eg, osimertinib, lazertinib, aumolertinib, alflutinib, and others in consultation with Medical Monitor)
 - i) If a patient has received 2 prior lines of EGRF TKI therapy, administration of the third-generation EGRF TKI must have been in the most recent line and in the setting of NSCLC with a demonstrated T790M substitution
 - ii) Enrollment of patients receiving third-generation EGFR TKIs other than osimertinib will be a maximum of $\approx 20\%$ of the enrolled population in each treatment arm
- Has adequate bone marrow reserve and organ function, within 14 days prior to randomization:
- Platelet count: $\geq 100\,000/\text{mm}^3$ or $\geq 100 \times 10^9/\text{L}$ (platelet transfusions are not allowed within 14 days
 - prior to randomization to meet eligibility)
- Hemoglobin: ≥ 9.0 g/dL (transfusion and/or CSF support not allowed within 14 days prior to randomization)
- Absolute neutrophil count: $\geq 1500/\text{mm}^3$ or $\geq 1.5 \times 10^9/\text{L}$ (CSF stimulating support not allowed within 14 days prior to randomization)
- Creatinine Clearance: CrCl ≥ 45 ml/min calculated by using the Cockcroft-Gault equation or measured CrCl.
- AST/ALT: $\leq 3 \times \text{ULN}$ (if liver metastases are present, $\leq 5.0 \times \text{ULN}$)

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- TBL: $\leq 1.5 \times \text{ULN}$ if no liver metastases ($< 3.0 \times \text{ULN}$ in the presence of documented Gilbert syndrome [unconjugated hyperbilirubinemia] or liver metastases)
- Serum albumin : ≥ 2.5 g/dL
- PT or PT-INR and aPTT/PTT: $\leq 1.5 \times \text{ULN}$, except for subjects on coumarin-derivative anticoagulants or other similar anticoagulant therapy, who must have PT-INR within therapeutic range as deemed appropriate by the investigator