





DISTRIBUTION AND PROGNOSTIC IMPACT OF EGFR AND KRAS MUTATIONS ACCORDING TO HISTOLOGICAL SUBTYPE AND TUMOR INVASION STATUS IN pTIS-3N0M0 LUNG ADENOCARCINOMA

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Objectives The impact of EGFR mutation on recurrence in resected lung adenocarcinoma is still controversial. KRAS G12C inhibitor became clinically available and KRAS mutations can guide the post-recurrence treatment. We evaluated prognostic impact of EGFR and KRAS mutations clinicopathological recurrence risks in resected by considering pTis-3N0M0 adenocarcinoma. Methods Clinicopathological features including EGFR and KRAS status were estimated in 877 resected cases. RFS and cumulative recurrence rate (CRR) were compared. Uni- and multivariate analyses for RFS were performed after excluding cases with little or no recurrence risks. Results EGFR mutation was more harbored in female, never-smoker, or patients with > 5% lepidic component. KRAS mutation was more harbored in patients with smoking history, IASLC histology grade 3, or with lymphovascular invasion. In IASLC grade 2 and 3, EGFR or KRAS mutation patients had significantly worse 5-year RFS than wild type patients (76.9% vs. 85.0%, HR = 1.55, 95% CI = 1.62-6.41, P < 0.001). KRAS mutant cases had higher 5-year CRR than EGFR mutant cases (16.7% vs. 21.4%, HR = 1.62, 95% CI = 0.96-7.19, P = 0.061). Multivariate analysis revealed positive EGFR/KRAS mutation status was related to worse RFS (HR = 2.007, 95% CI = 1.265-3.183, P = 0.003). Conclusions KRAS mutations were more confirmed in cases with increased risk of recurrence compared to EGFR. Positive EGFR and KRAS mutation statuses were risk for recurrence in resected IASLC grade 2 and 3 patients. KRAS mutation statuses should be evaluated simultaneously when assessing recurrence risk of EGFR.