TREATMENT OF METASTATIC NSCLC WITH A TARGETABLE TUMOR MUTATION

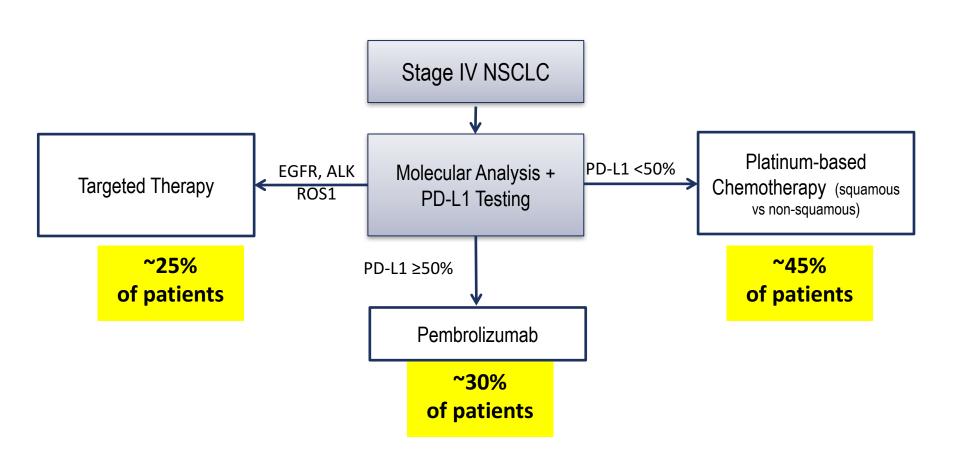
Gregory J. Riely

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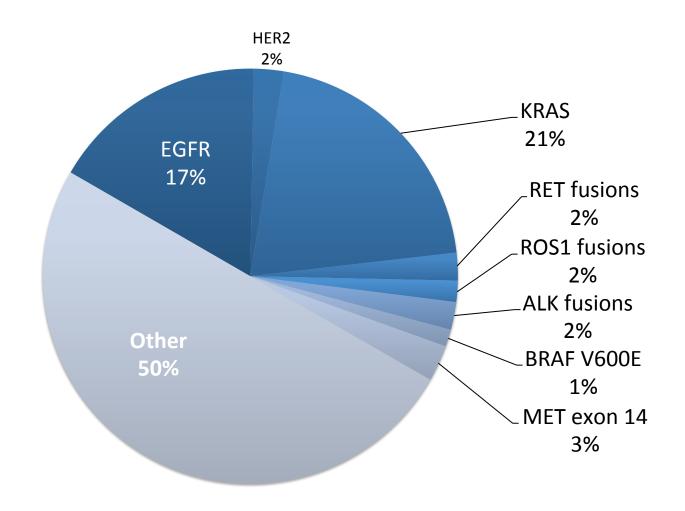
Disclosures

Consulting Agreement	Genentech BioOncology
Contracted Research	Ariad Pharmaceuticals Inc, Astellas Pharma Global Development Inc, Novartis, Pfizer Inc

The Current Approach to First-Line Treatment of Patients with Advanced NSCLC

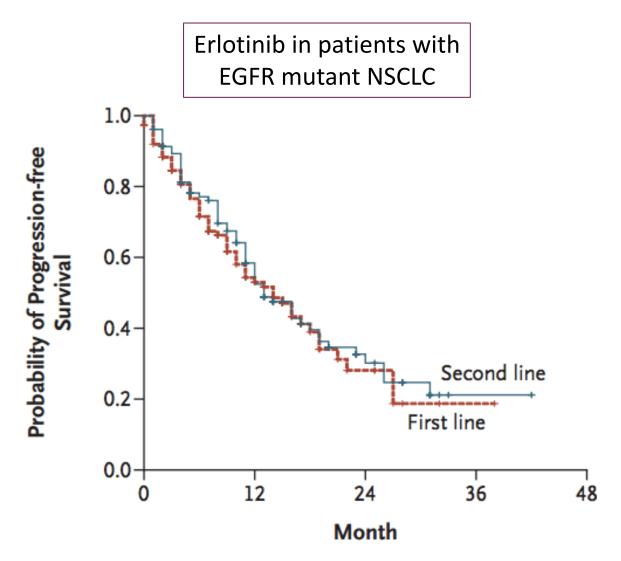


Subtypes of NSCLC can be defined by genotypes

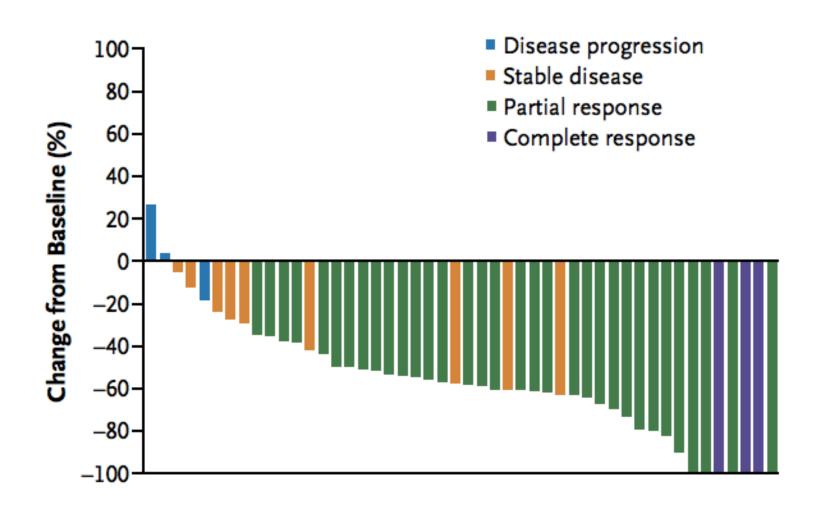


MSK-IMPACT data, n~2000

For molecularly targeted therapies, line of therapy may not matter



Crizotinib in patients with *ROS1* positive lung cancers

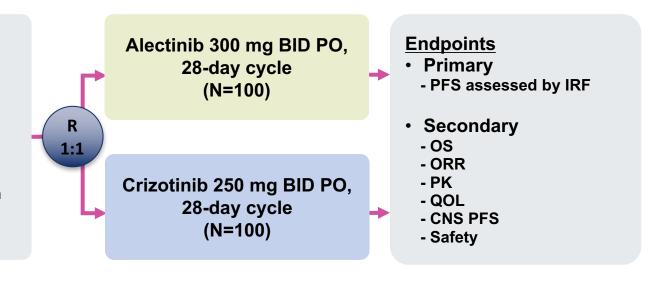


Alectinib versus crizotinib in patients with ALK-positive non-small-cell lung cancer (J-ALEX): an open-label, randomised phase 3 trial

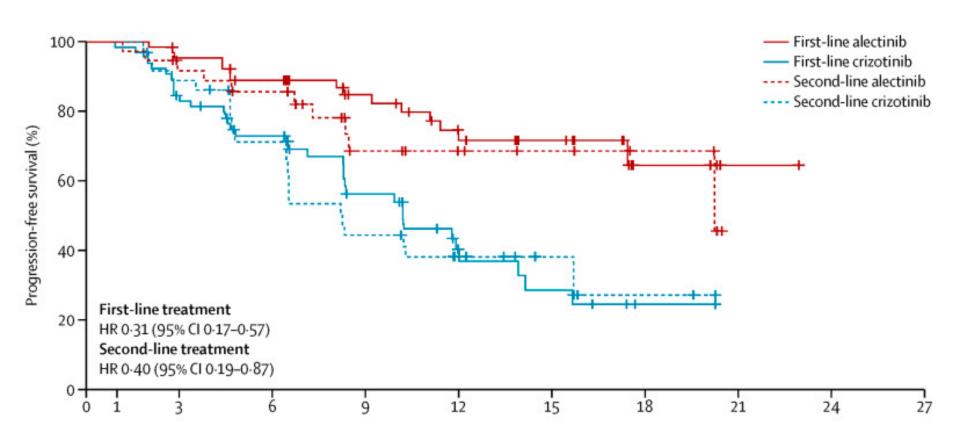
Toyoaki Hida, Hiroshi Nokihara, Masashi Kondo, Young Hak Kim, Koichi Azuma, Takashi Seto, Yuichi Takiguchi, Makoto Nishio, Hiroshige Yoshioka, Fumio Imamura, Katsuyuki Hotta, Satoshi Watanabe, Koichi Goto, Miyako Satouchi, Toshiyuki Kozuki, Takehito Shukuya, Kazuhiko Nakagawa, Tetsuya Mitsudomi, Nobuyuki Yamamoto, Takashi Asakawa, Ryoichi Asabe, Tomohiro Tanaka, Tomohide Tamura

Key Entry Criteria

- Stage IIIB/IV or recurrent ALK-positive NSCLC
- ALK centralized testing (IHC and FISH or RT-PCR)
- ECOG PS 0-2
- ≥1 measurable lesion assessed by investigator
- Treated/asymptomatic brain metastases allowed
- ≤1 prior chemotherapy

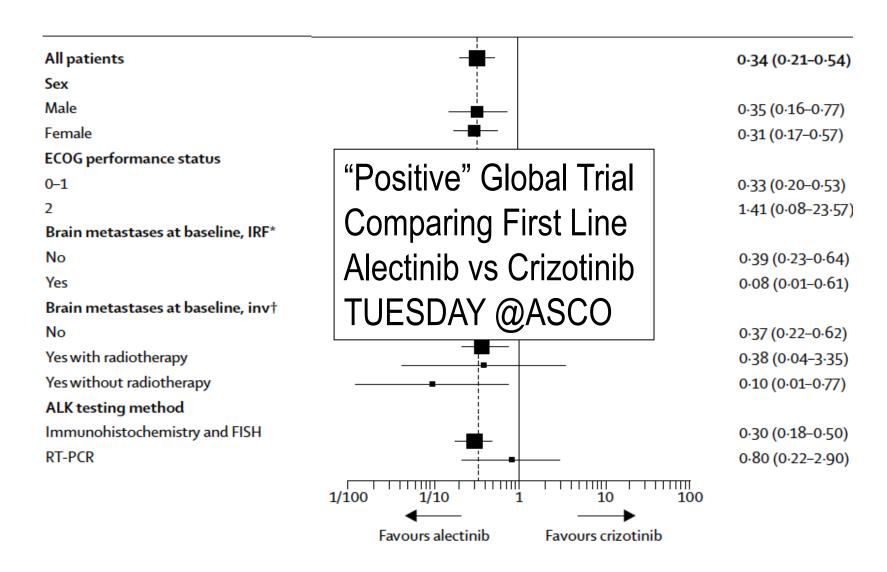


ALECTINIB VS CRIZOTINIB AS 1ST LINE ALK INHIBITOR



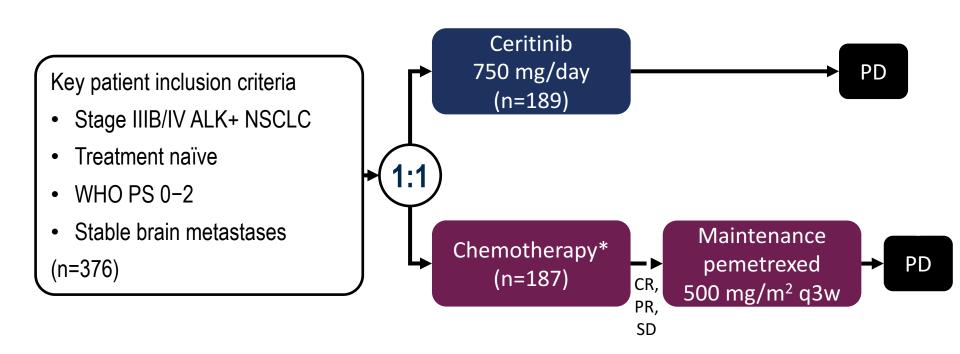
ALECTINIB VS CRIZOTINIB AS 1ST LINE ALK INHIBITOR

HR (95% CI)



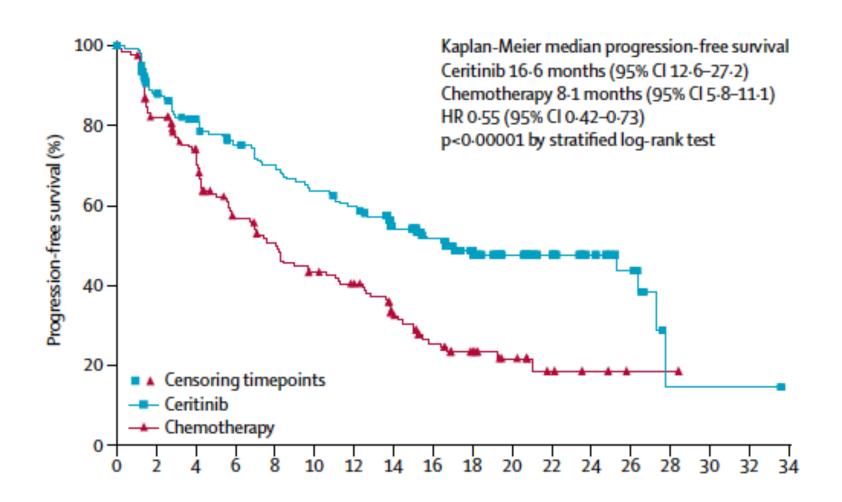
First-line ceritinib versus platinum-based chemotherapy in advanced ALK-rearranged non-small-cell lung cancer (ASCEND-4): a randomised, open-label, phase 3 study

Jean-Charles Soria, Daniel SW Tan, Rita Chiari, Yi-Long Wu, Luis Paz-Ares, Juergen Wolf, Sarayut L Geater, Sergey Orlov, Diego Cortinovis, Chong-Jen Yu, Maximillian Hochmair, Alexis B Cortot, Chun-Ming Tsai, Denis Moro-Sibilot, Rosario G Campelo, Tracey McCulloch, Paramita Sen, Margaret Duqan, Serafino Pantano, Fabrice Branle, Cristian Massacesi, Gilberto de Castro Jr



^{*} Cisplatin 75 mg/m², or carboplatin (target AUC 5-6)/pemetrexed 500 mg/m² q3w

1ST LINE CERITINIB IS BETTER THAN CHEMOTHERAPY



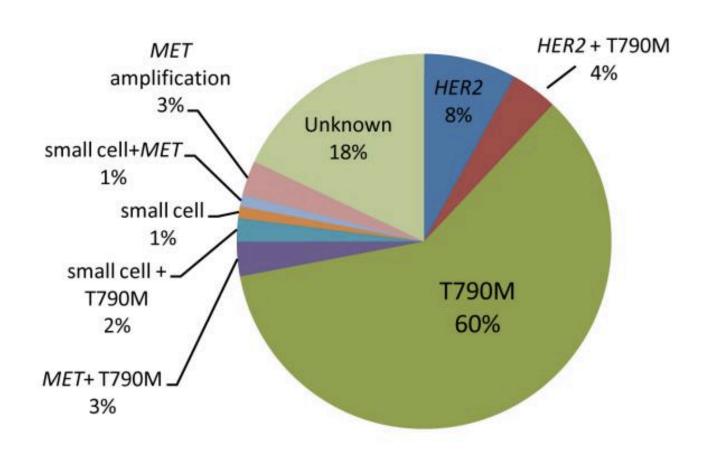
PRESS RELEASE: FDA APPROVAL FOR EXPANDED USE OF CERITINIB

May 26, 2017

The US Food and Drug Administration (FDA) approved the expanded use of ceritinib to include the first-line treatment of patients with metastatic non-small cell lung cancer (NSCLC) whose tumors are anaplastic lymphoma kinase (ALK)-positive, as detected by an FDA-approved test ...

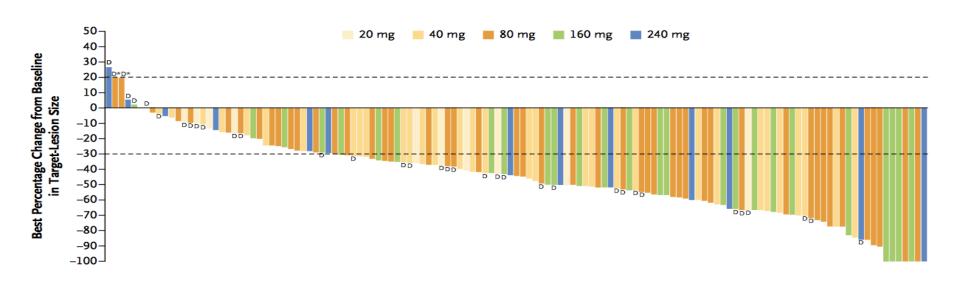
... based on results from an open-label, randomized, multicenter, global, Phase III trial, ASCEND-4.

Mechanisms of Acquired Resistance to EGFR TKI



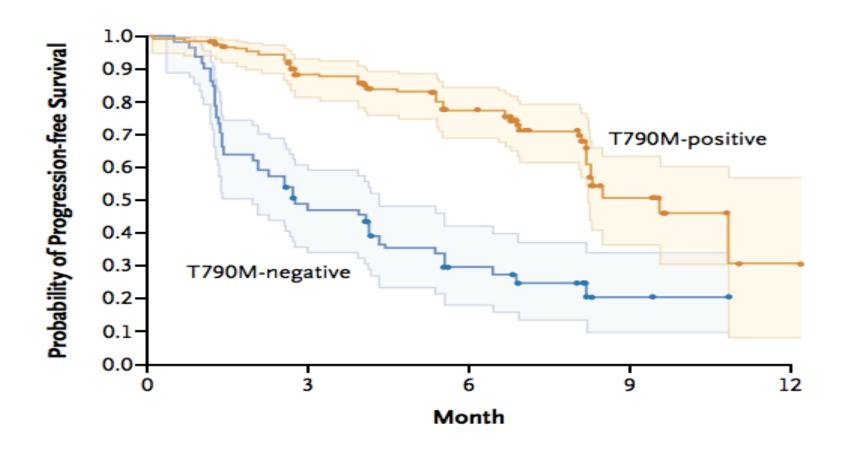
Osimertinib in Patients With Acquired Resistance to EGFR TKI and *EGFR* T790M

EGFR T790M-Positive

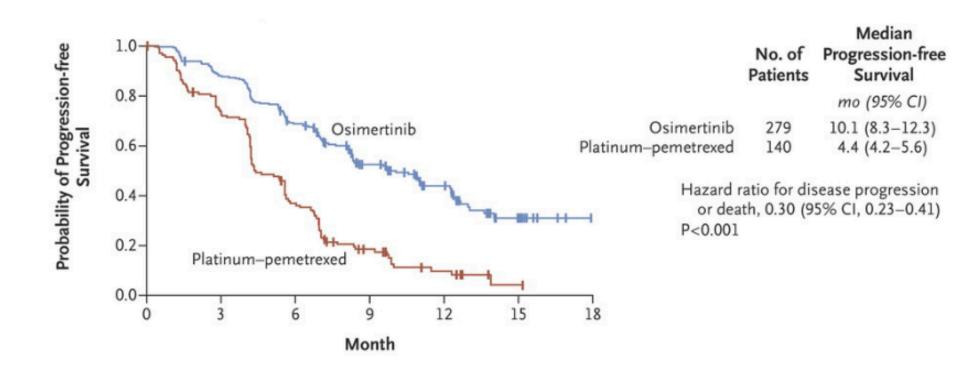


Objective Response Rate = 61%

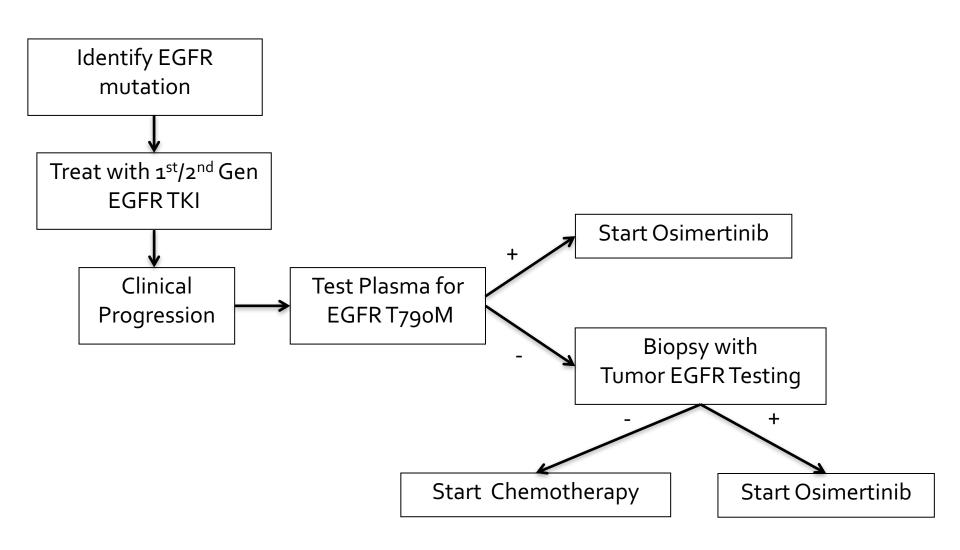
Osimertinib Progression-Free Survival by T790M



Osimertinib as Second Line Therapy



How I Treat Patients with EGFR mutant NSCLC



The Method of Molecular Analysis Matters

