



2024 World Conference  
on Lung Cancer

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# Novel Immunotherapies

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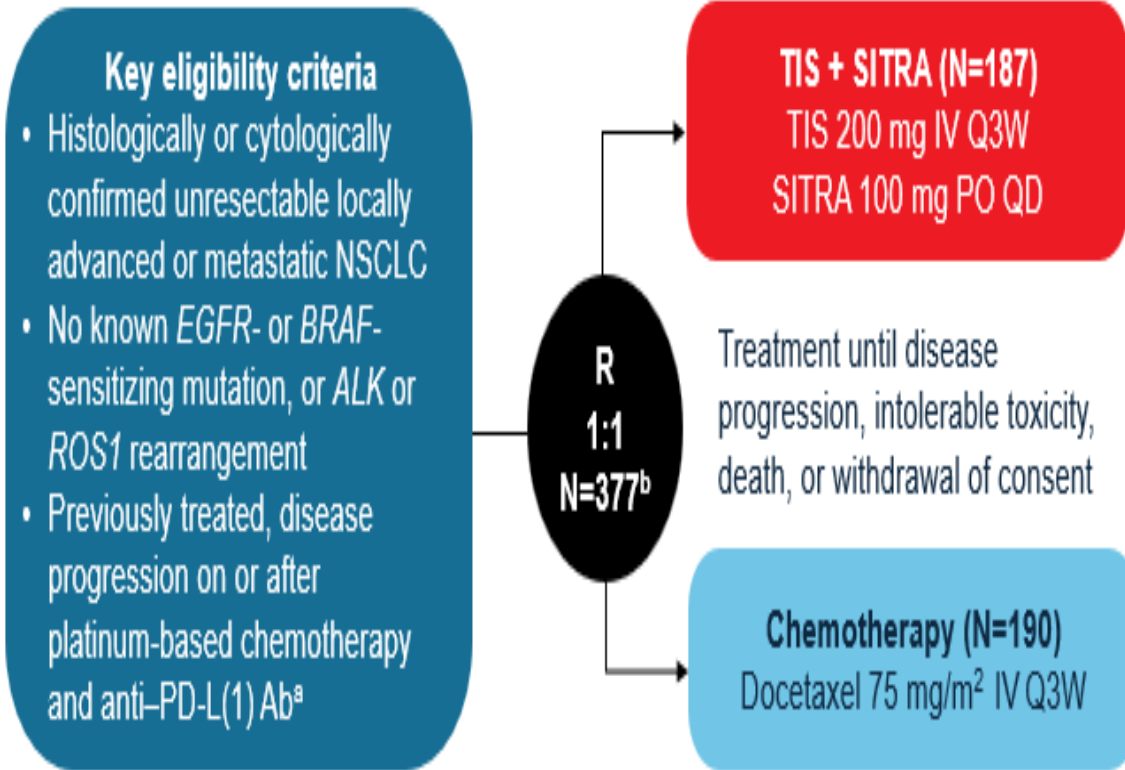


*Author*

## Abstracts covered:

- OA06.03 Q. Zhou et al. SAFFRON-301: Tislelizumab Plus Sitravatinib in Advanced/Metastatic NSCLC Progressing on/after Chemotherapy and Anti-PD-(L)1
- OA06.04 M Marmarelis. Phase II Study of Pembrolizumab and Itacitinib for Patients with Metastatic NSCLC Expressing PD-L1
- OA06.05 J Wrangle. IL15 Superagonist N-803 + Checkpoint Inhibitor (CPI) Prolongs OS in 2nd Line or Greater NSCLC Patients Failing CPI
- OA11.03 J Hiltermann. Efficacy and Safety of Rilvegostomig, an Anti-PD-1/TIGIT Bispecific, for CPI-naïve Metastatic NSCLC with PD-L1 1–49% or  $\geq 50\%$
- OA11.04: D. Spigel. Volrustomig + platinum doublet chemotherapy in first-line non-small-cell lung cancer (NSCLC): Phase 1b trial update





## Background

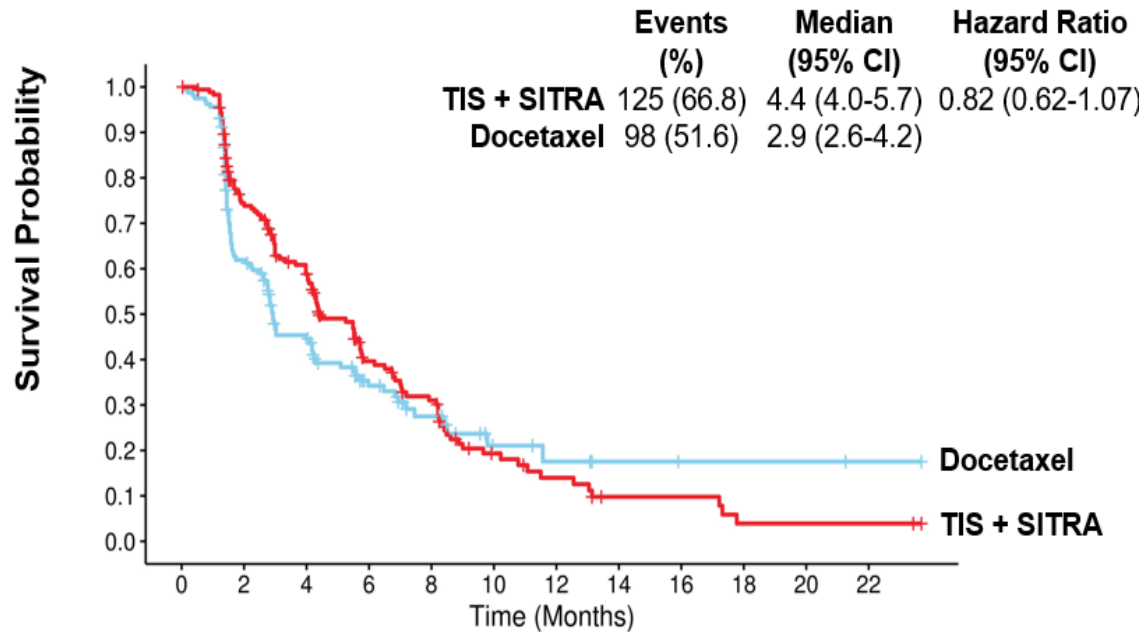
- Tislelizumab is anti-PD-1 immunotherapy
- Sitravatinib is multitargeted TKI against VEGF and TAM

## Clinical implications

- No PFS or OS benefit to this combination in the 2L setting
- These data are consistent with other trials showing no benefit of TKI and continuation immunotherapy



### PFS by IRC



Number at risk

ARM=Docetaxel	190	84	53	30	17	7	5	3	2	2	2	1
ARM=TIS + SITRA	187	118	87	47	35	15	10	5	5	2	2	2

### Background

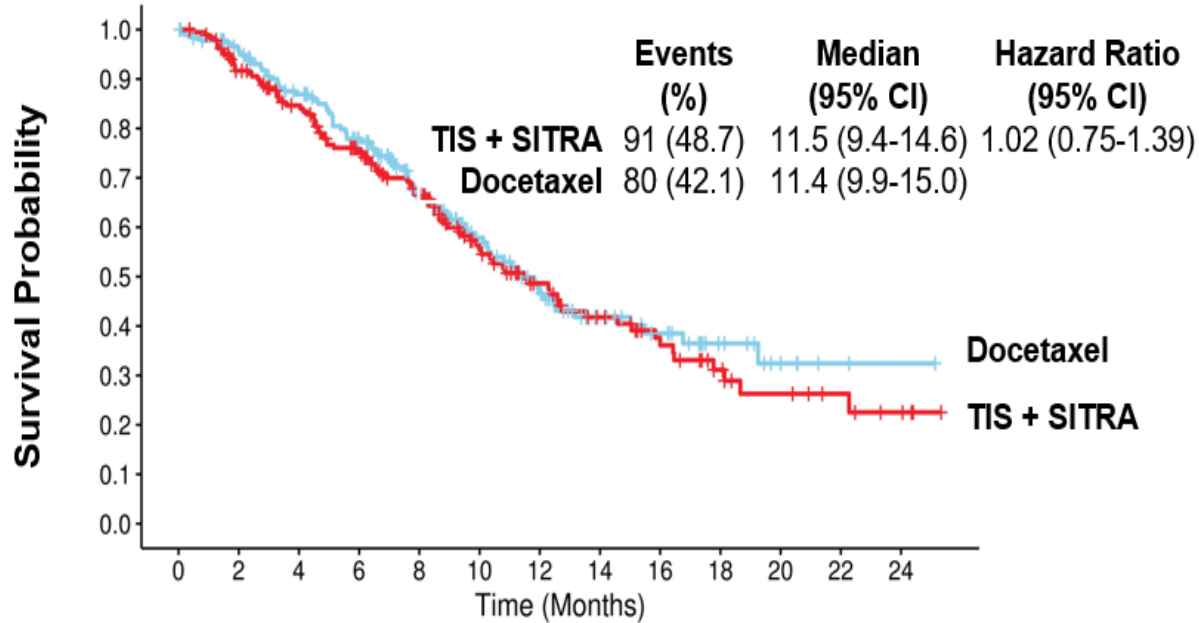
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**OS**



	0	2	4	6	8	10	12	14	16	18	20	22	24
ARM=Docetaxel	190	162	140	117	82	60	43	29	22	12	6	2	1
ARM=TIS + SITRA	187	161	140	116	91	60	44	33	25	15	10	7	4

**Background**

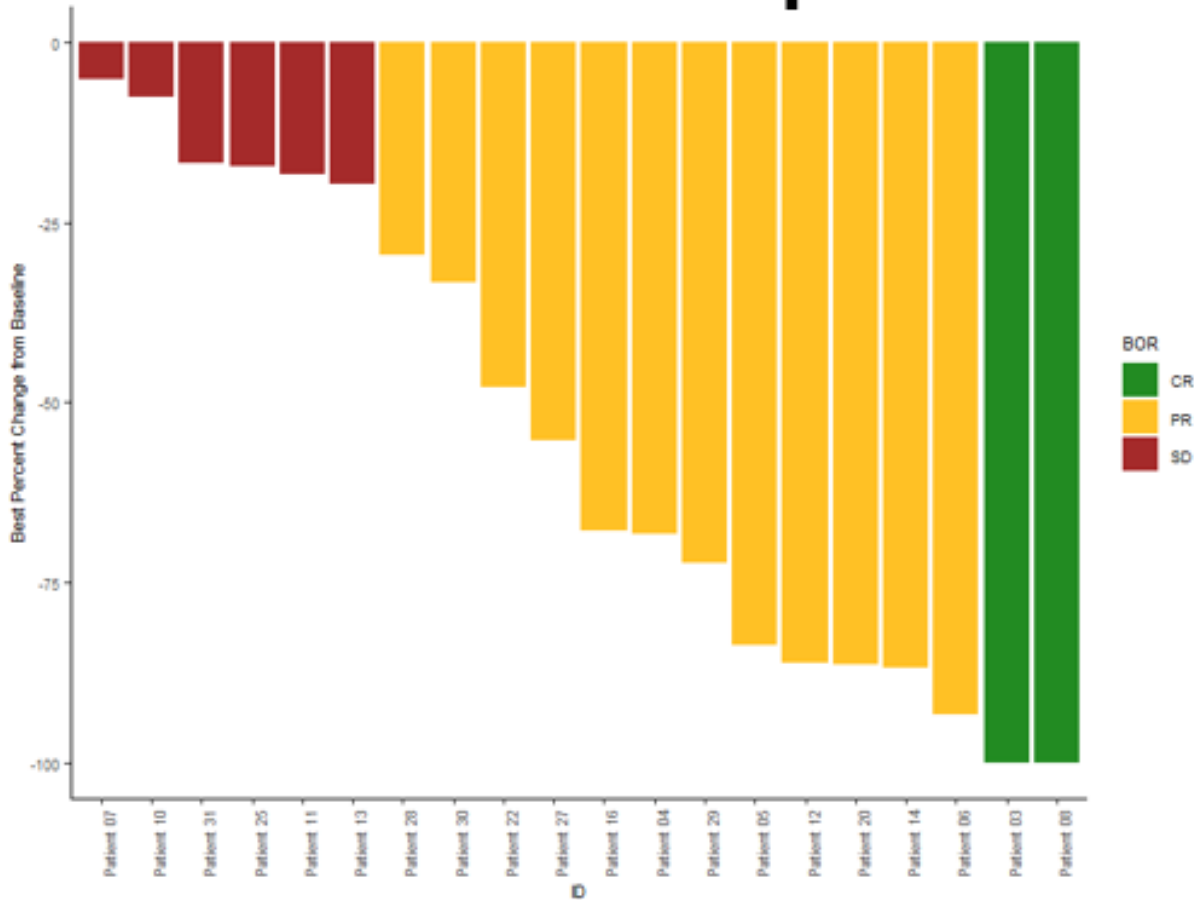
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## Best Overall Response



## Background

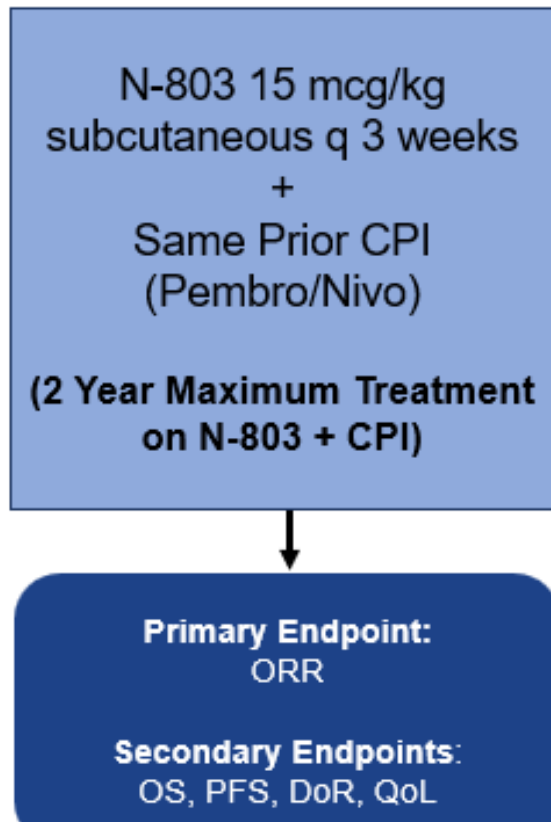
- Pembrolizumab is anti-PD-1 immunotherapy
- Itacitinib is JAK inhibitor
- This is 1L trial for PD-L1 50%

## Clinical implications

- ORR 66%; mPFS 15.6mo; mOS 53.4mo
- Delayed JAK inhibitors are feasible



## QUILT-3.055 Study Design 2nd and 3rd Line NSCLC Progressing on CPI



## Background

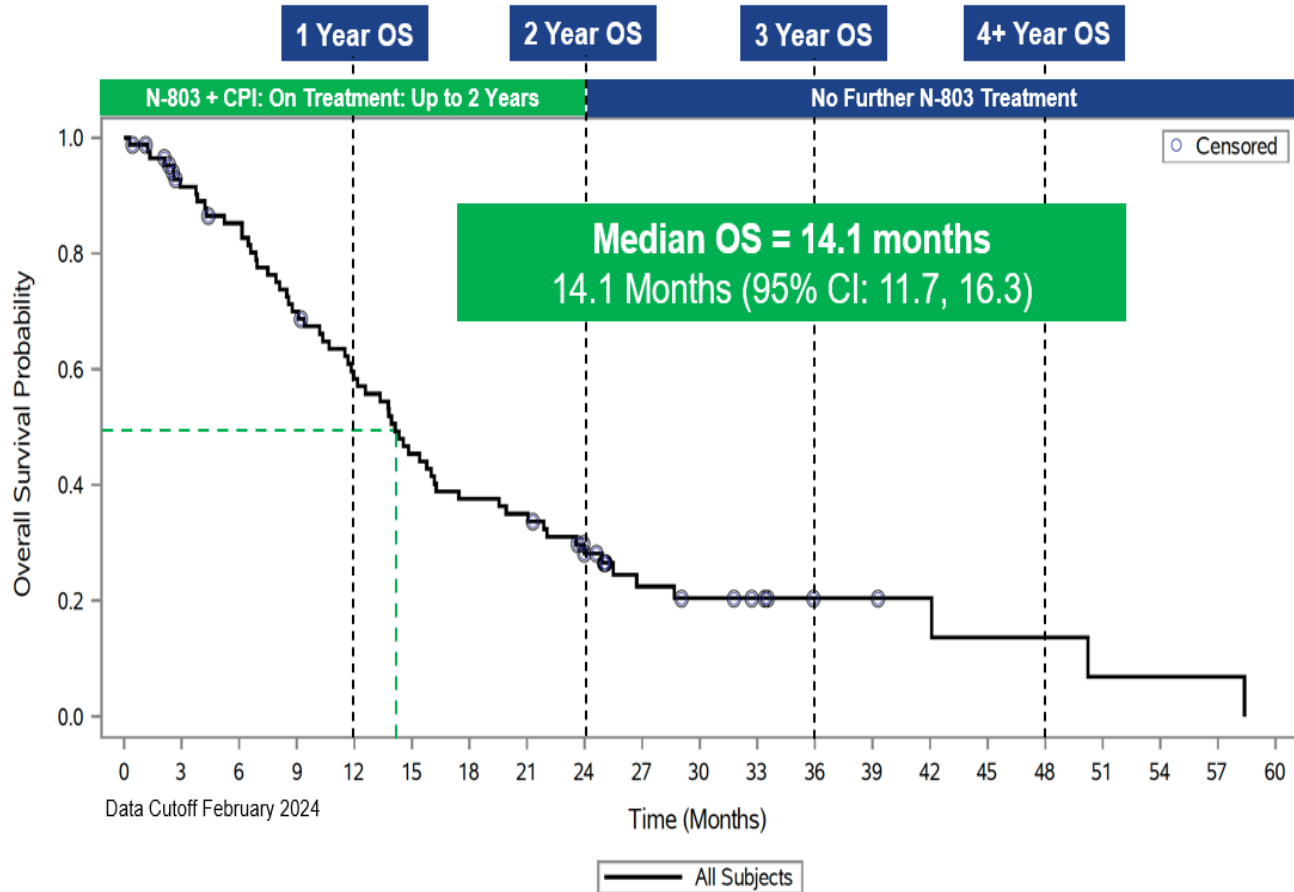
- N-803 is IL15 agonist
- - Activates and proliferates CD8+ killer cells and NK cells

## Clinical implications

- 2L and 3L NSCLC with prior CPI and chemo
- Approved in localized bladder cancer, being developed in NSCLC



**OA06.05 J Wrangle. IL15 Superagonist N-803 + Checkpoint Inhibitor (CPI) Prolongs OS in 2nd Line or Greater NSCLC Patients Failing CPI**



**Background**

- N-803 is IL15 agonist
- - Activates and proliferates CD8+ killer cells and NK cells

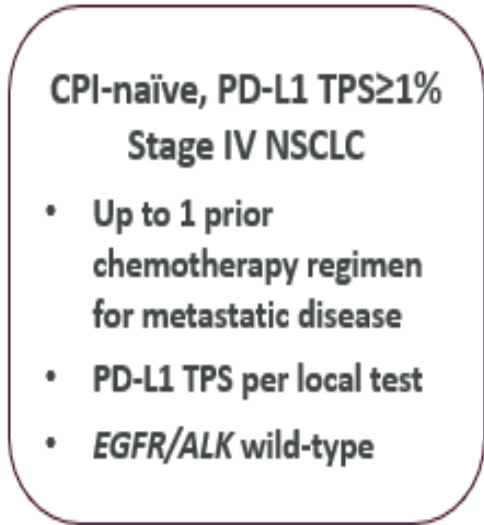
**Clinical implications**

- 2L and 3L NSCLC with prior CPI and chemo – response rate 5-16%
- Approved in noninvasive bladder cancer, being developed in NSCLC

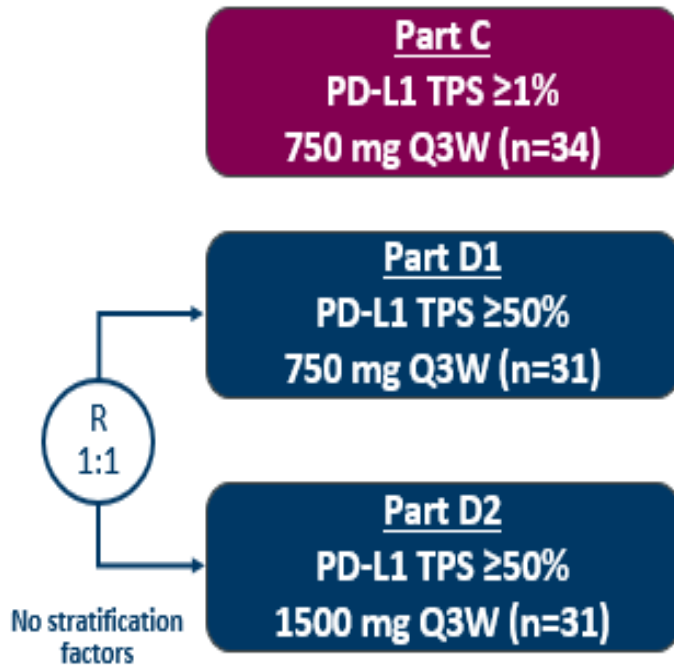




## ARTEMIDE-01



Part A (dose escalation) and Part B (dose expansion) in CPI resistant NSCLC not pictured



## Background

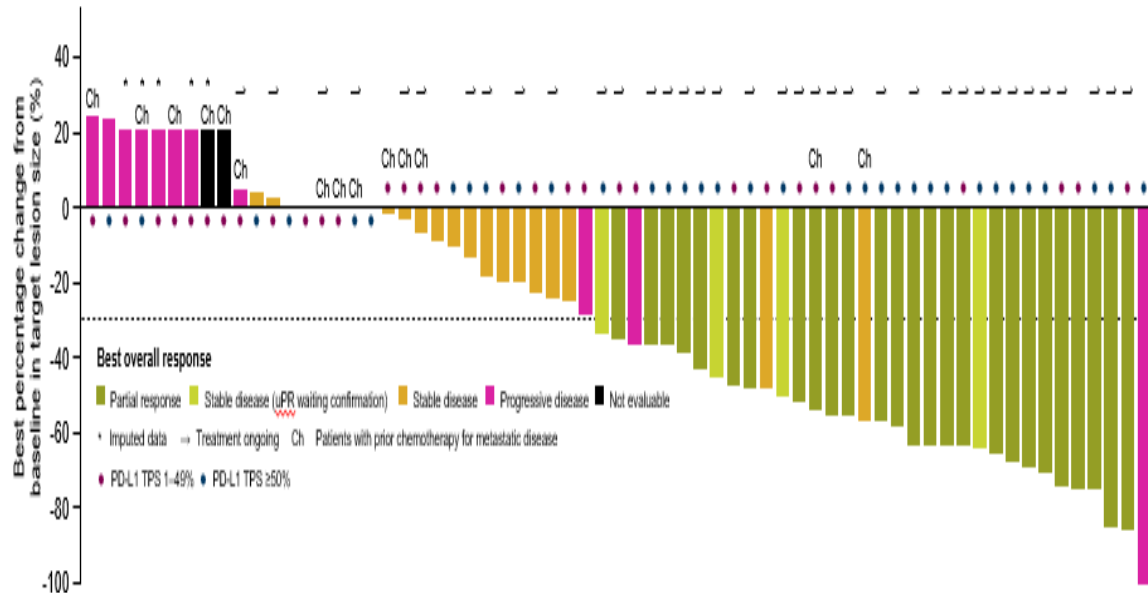
- Rilvegostomig is anti-PD-1/TIGIT bispecific antibody

## Clinical implications

- 1L NSCLC – response rate 29-62%
- Unclear whether this is “better” than pembrolizumab alone in this selected population



## ARTEMIDE-01



n (%)	PD-L1 1–49% TPS (n=31)	PD-L1 ≥50% TPS (n=34)
ORR, confirmed + pending [95%CI]	9 (29%) [14.2, 48.0]	21 (61.8%) [43.6, 77.8]
Continuing treatment	12 (38.7%)	24 (70.6%)

4 patients with PD-L1 ≥50% pending confirmation

## Background

- Rilvegostomig is anti-PD-1/TIGIT bispecific antibody

## Clinical implications

- 1LNSCLC – response rate 29-62%
- Unclear whether this is “better” than pembrolizumab alone in this selected population



## Background

- Rilvegostomig is anti-PD-1/TIGIT bispecific antibody
- Chemo is chemo

## Clinical implications

- RR 43% for non-squam, 65% for squam
- Unclear if better than otherwise expected

### Key inclusion criteria

1L recurrent  
or metastatic NSCLC

EGFR/ALK wild-type

Cohort 1: nonsquamous (N=120)

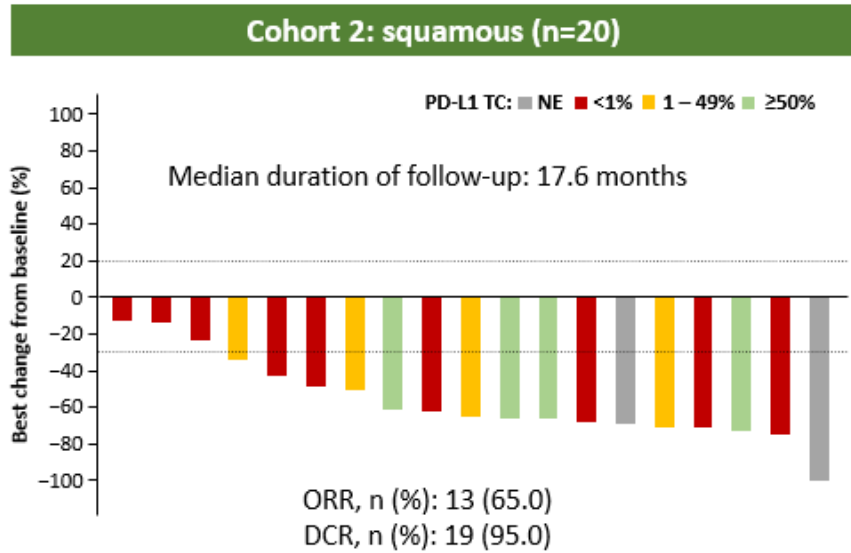
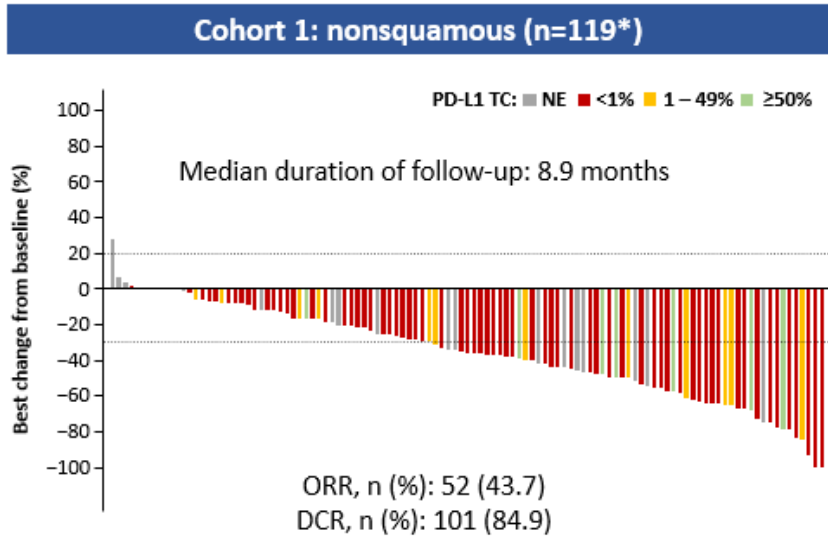
Volrustomig 750 mg IV Q3W + CTx\*

Cohort 2: squamous (N=20)

Volrustomig 750 mg IV Q3W + CTx\*



# OA11.04: D. Spigel. Volrustomig + platinum doublet chemotherapy in first-line non-small-cell lung cancer (NSCLC): Phase 1b trial update



## Background

- Rilfevostomig is anti-PD-1/TIGIT bispecific antibody
- Chemo is chemo

## Clinical implications

- RR 43% for non-squam, 65% for squam
- Unclear if better than otherwise expected



## Conclusions

- Second line continuation immunotherapy together with VEGF/TAM TKIs does not beat docetaxel (cabozantinib, lenvatinib, and sitravatinib have negative trials). JAK inhibitors are TBD.
- IL-15 agonists are interesting addition to immunotherapy refractory setting – trials are ongoing
- Unclear if PD-1/TIGIT bispecific approach better than first line immunotherapy alone – trials are ongoing

**THANK YOU!**

