



*This educational activity is supported by an independent  
medical education grant from Genmab*

*An Industry Supported Symposium during the IGCS 2025 Global Annual Meeting*

# **Global Perspective in Advance, Recurrent Cervical Cancer: Resources, Opportunities and Access to Care**

*This session is not included in the main event CME/CPD credit.*

**Cape Town, South Africa**

**Thursday, November 6, 2025**

**18:15 - 19:15 GMT+2**



# Welcome and Introductions

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**Leslie Randall, MD**

Inova Health

Fairfax, Virginia, USA



# MODERATOR

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**Leslie Randall, MD, MAS**

Inova Health  
Fairfax, Virginia, USA

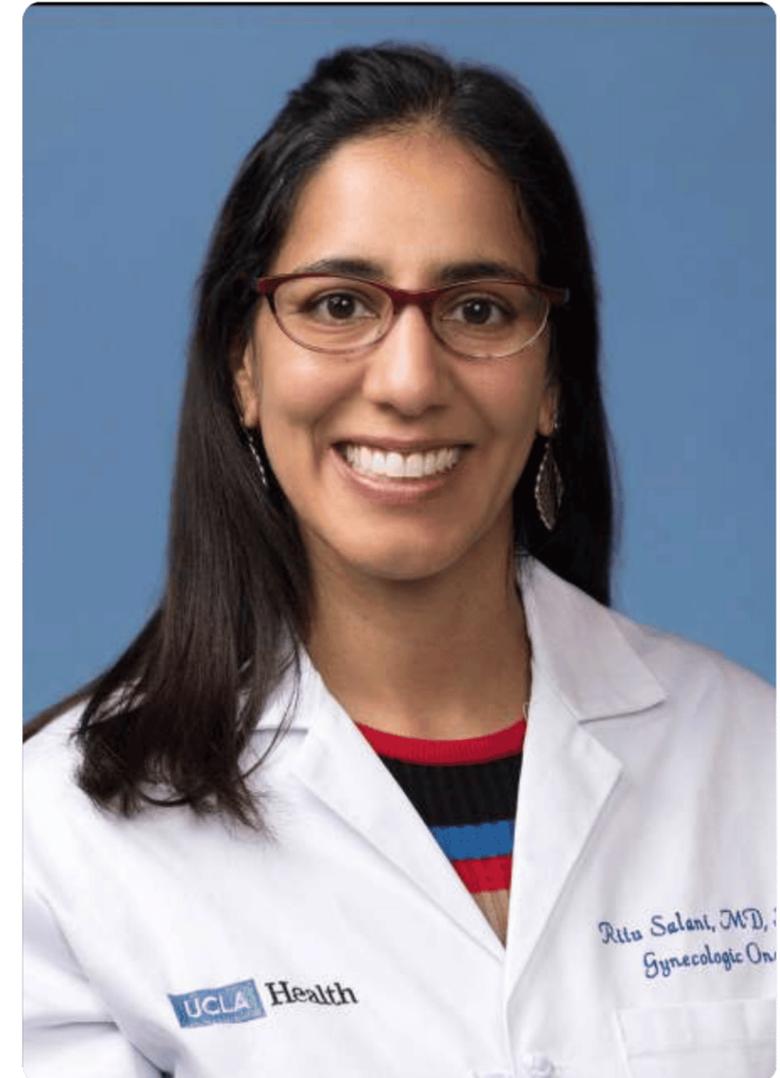
# FACULTY

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**Kosei Hasegawa, MD, PhD**

Saitama Medical University  
Kawagoe, Japan



**Ritu Salani, MD, MBA**

UCLA Health  
Los Angeles, California, USA



# Disclosures

## Faculty

**Leslie Randall, MD, MAS**

Consultant (AstraZeneca; Genmab; Pfizer; GSK, Eisai; Merck; Abbvie; GOG Foundation, Corcept, Regeneron)  
Research Funding (Merck; AbbVie; GOG Foundation, Regeneron, Genmab, Seagen/Pfizer)

**Ritu Salani, MD, MBA**

Advisory board: Genmab, Pfizer, Abbvie, Daiichi, Merck, Eisai, Karyopharm  
Royalties: Up to Date.

**Kosei Hasegawa, MD, PhD**

# Learning Objectives

**Upon completion of this activity, learners will:**

- Review the current landscape for the treatment of recurrent cervix cancer based on pivotal clinical trial data and current insights.
- Understand the role of ADC in cervix cancer, and how to educate patients on this novel treatment option.
- Review and understand prevalence of disease, prevention and available resources.



# Agenda

Topic	Presenter
<b>Welcome and Introduction</b>	<b>Dr. Leslie Randall</b> Inova Health, Washington, DC, USA
<b>Current Landscape of Recurrent Cervical Cancer</b>	<b>Dr. Kosei Hasegawa</b> Saitama Medical University, Saitama, Japan
<b>Antibody Drug Conjugates (ADCs) in Cervical Cancer</b>	<b>Dr. Ritu Salani</b> UCLA Health, Los Angeles, CA, USA
<b>Global Epidemiology &amp; Prevention Strategies</b>	<b>Dr. Leslie Randall</b> Inova Health, Washington, DC, USA
<b>Panel Discussion, Q&amp;A and Closing Comments</b>	<b>All Faculty</b>



# Current Landscape of Recurrent Cervical Cancer

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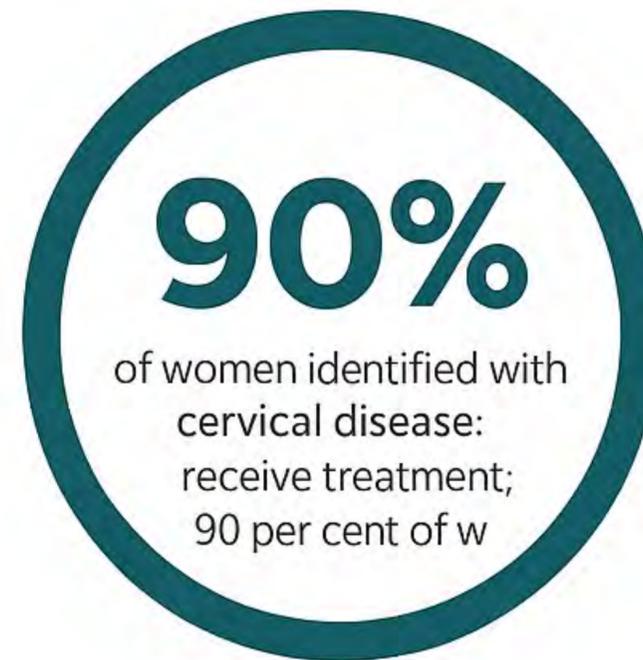
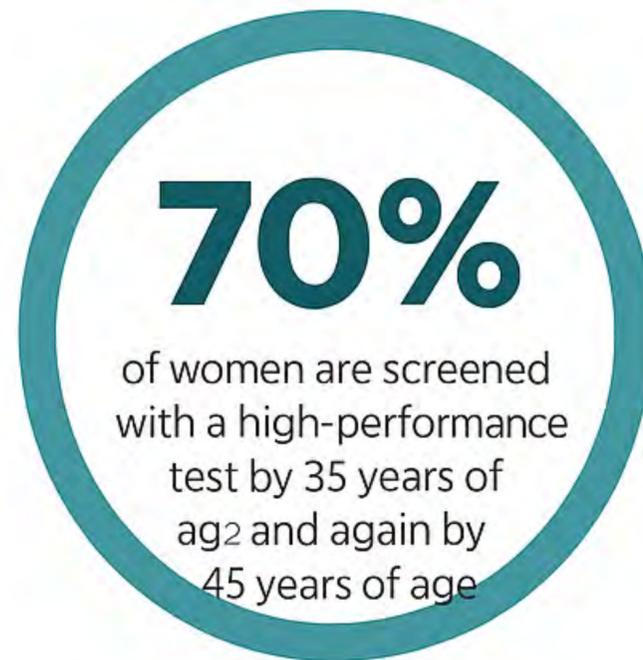
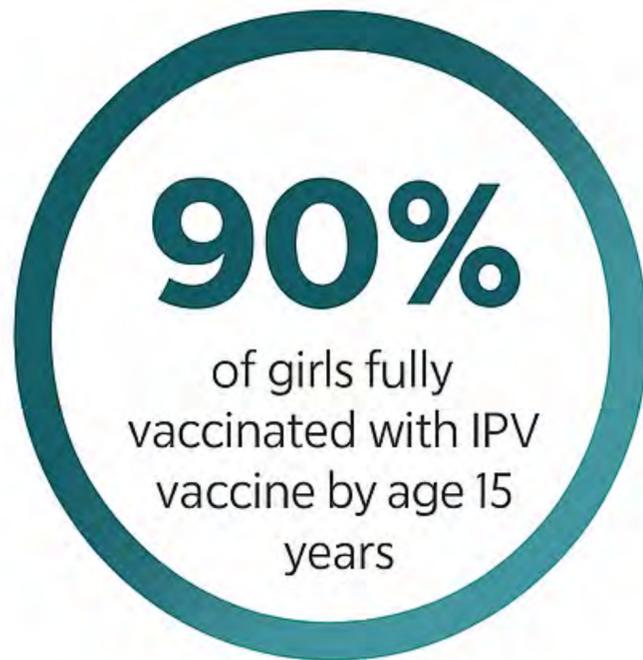
**Kosei Hasegawa, MD, PhD**

Saitama Medical University International Medical Center  
Saitama, Japan



- Cervical cancer: ~600,000 new cases, ~340,000 deaths annually (GLOBOCAN 2024).
- 9 of 10 deaths occur in low- and middle-income countries.
- But for the first time, *elimination* is possible.

Each country should meet the 90-70-90 targets by 2030 to get on the path to eliminate cervical cancer within the next century.



# Cervical Cancer in Japan and US

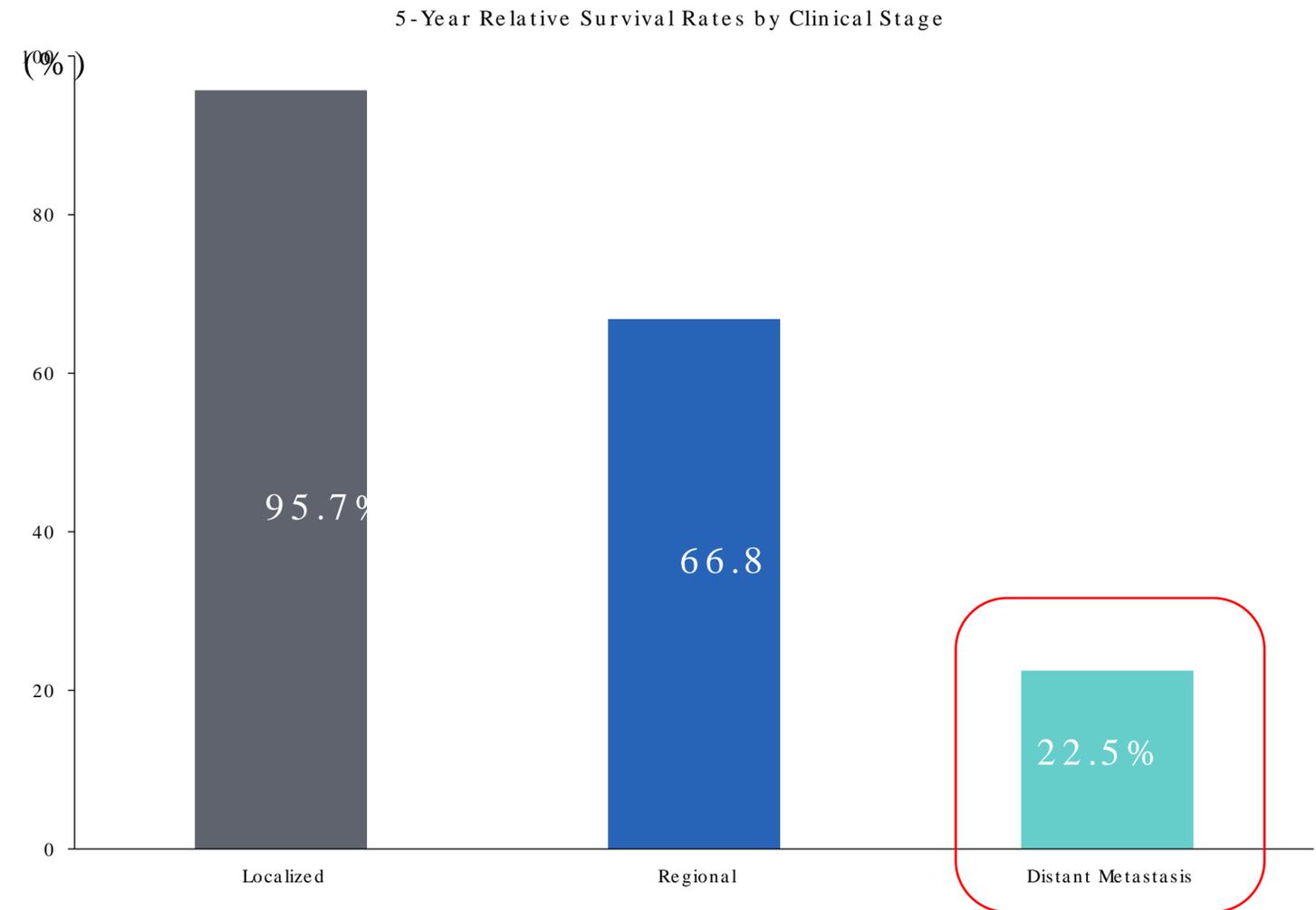
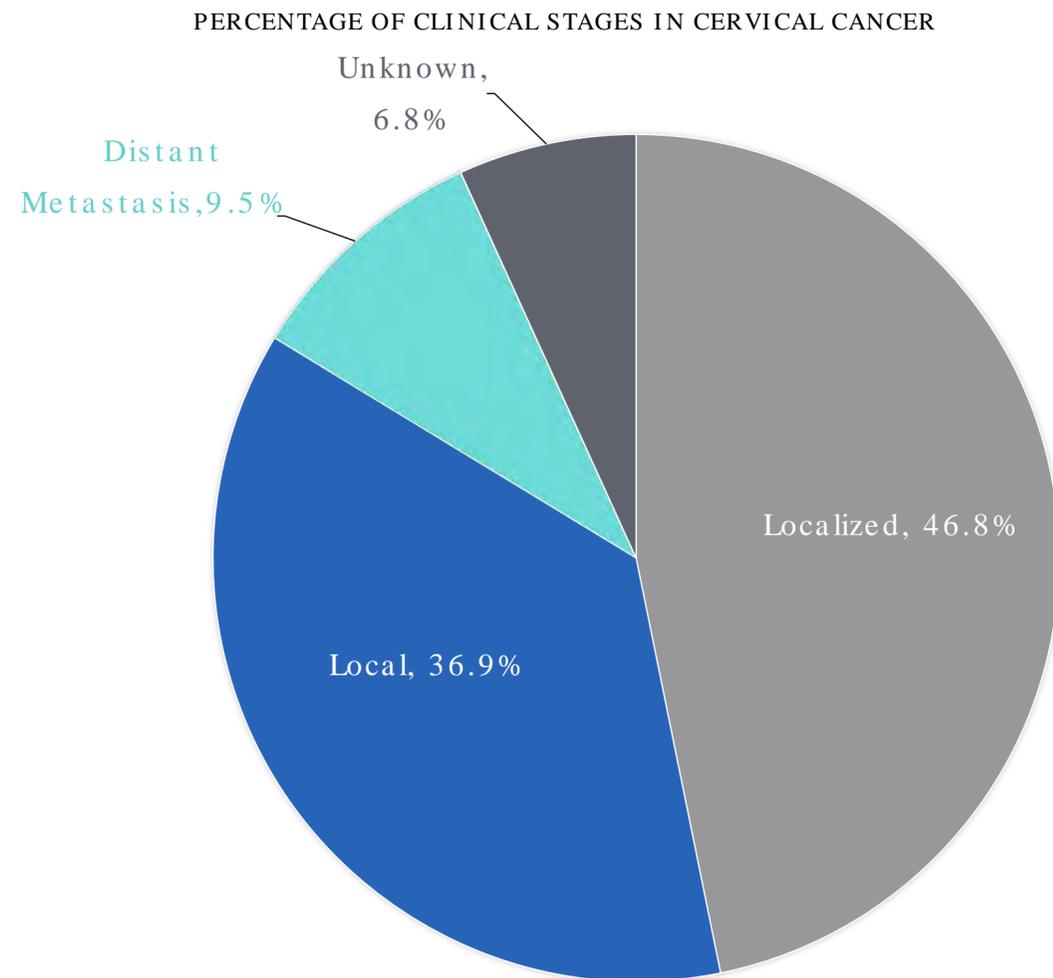
Cancer types	Number of Diagnosis (2018)	Number of Death (2019)	5 year relative survival rate (2009-11)
Japan	10,978	2,921	76.5 %
US	12,967	4,152	-

- HPV vaccination coverage by age15, females: Japan 6%, US 44% in 2015
- Cervical cancer screening: Japan 43.7%, US 72.6% in 2019



# Epidemiology of Advanced or Recurrent Cervical Cancer

In Japan, 9.5% of cervical cancer patients are diagnosed with distant metastasis, and the 5-year relative survival rate for patients with distant metastasis is 22.5%, indicating this is a group with a poor prognosis.



Localized: Confined to the primary organ

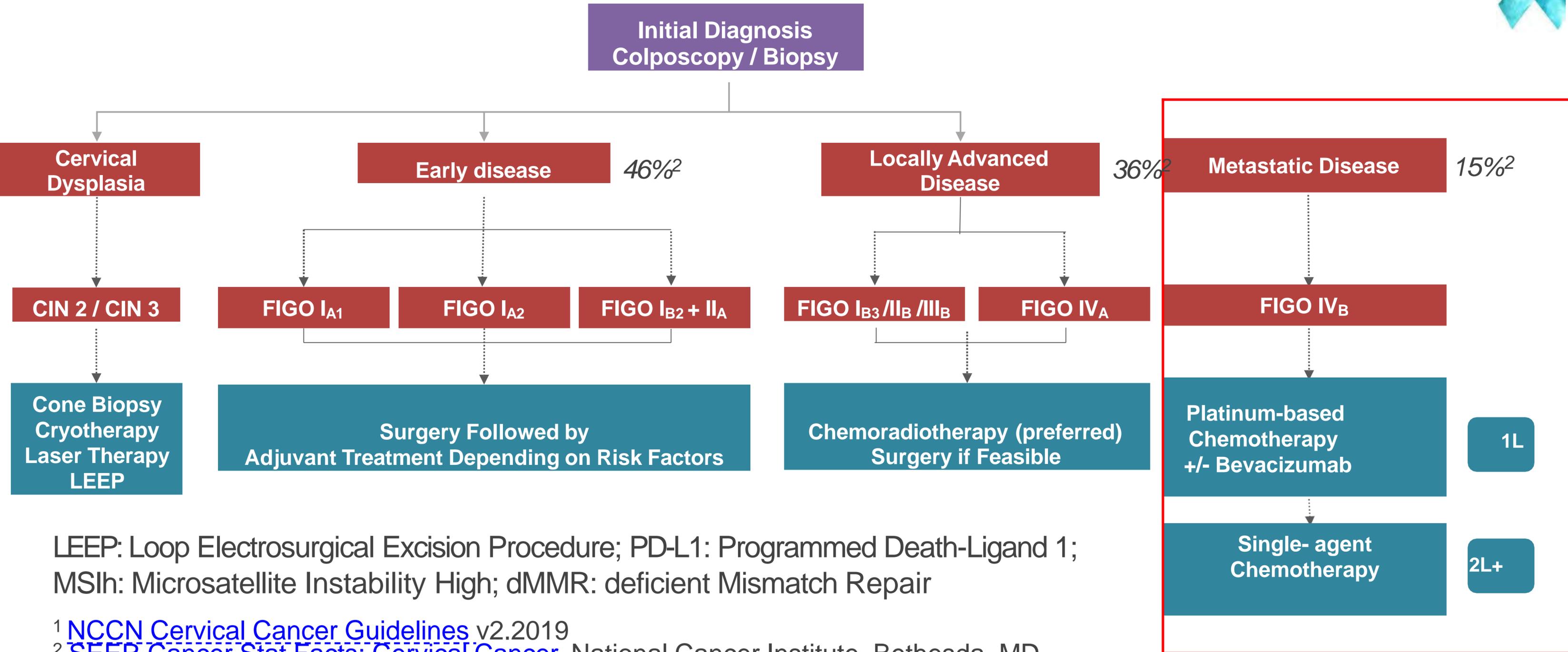
Regional: Regional lymph node metastasis (involvement of regional lymph nodes draining the primary organ, but no invasion of adjacent organs) or invasion of adjacent organs (direct invasion of adjacent organs, but no distant metastasis)

Distant metastasis: Metastasis or invasion to distant organs, distant lymph nodes, etc.

National Cancer Incidence Monitoring Summary: 2009-2011 Survival Rate Report (National Cancer Center Cancer Control Information Center, 2020)

National Cancer Center Research and Development Fund "Research on Improving the Accuracy and Utilization of Regional Cancer Registries" FY2010 Report

# Cervical Cancer: Summary of Treatment



LEEP: Loop Electrosurgical Excision Procedure; PD-L1: Programmed Death-Ligand 1; MSIh: Microsatellite Instability High; dMMR: deficient Mismatch Repair

<sup>1</sup> [NCCN Cervical Cancer Guidelines v2.2019](#)

<sup>2</sup> [SEER Cancer Stat Facts: Cervical Cancer](#). National Cancer Institute. Bethesda, MD



# **2/3 Line+ Recurrent**

# Regimen for 2L+ Metastatic Cervical Cancer

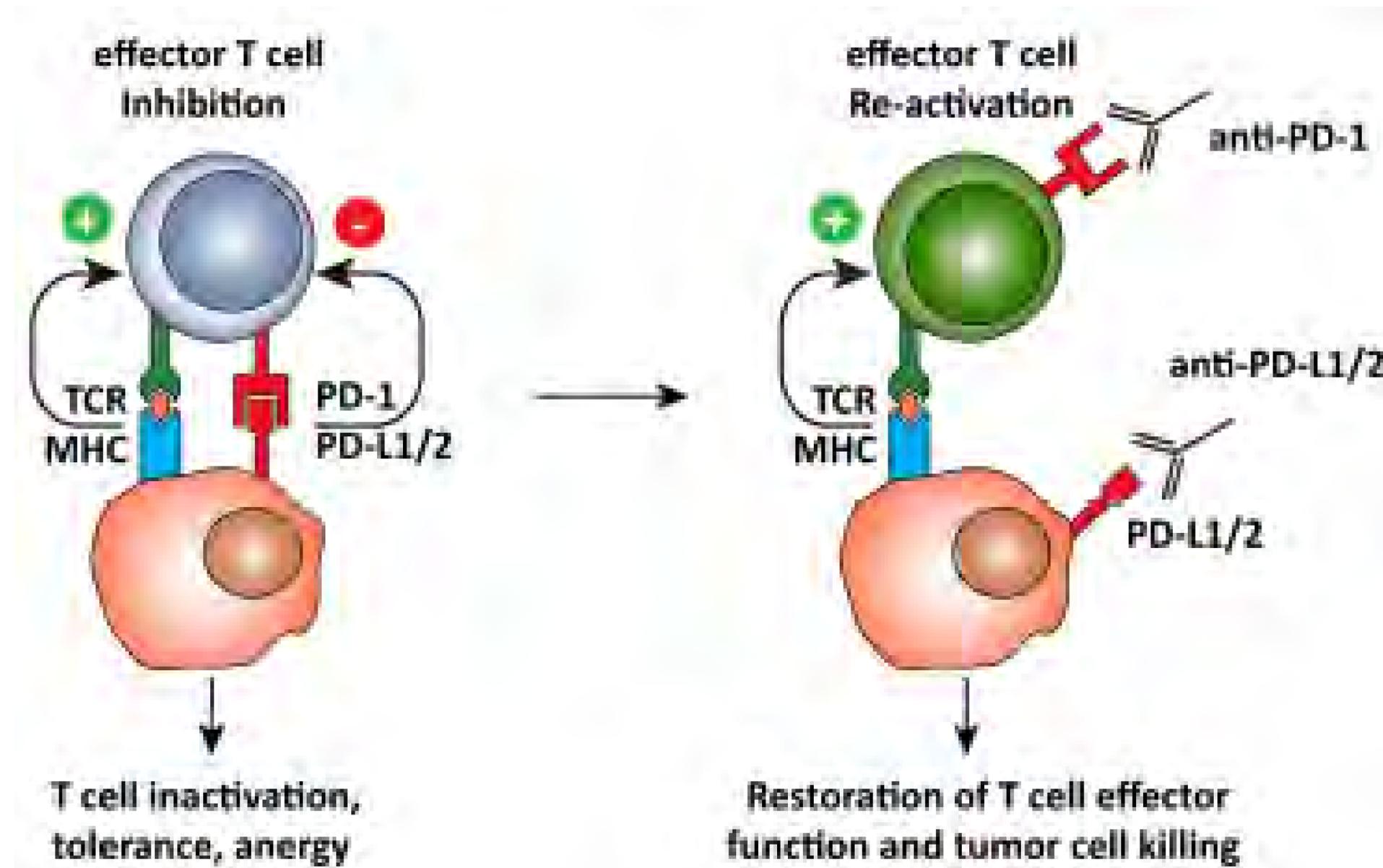


Design	N	ORR (%)	PFS (months)	OS (months)
Topotecan	45	12.5	2.1	6.6
Vinorelbine	44	13.7	NS	NS
<b>Pemetrexed</b>	<b>29</b>	<b>15</b>	<b>3.1</b>	<b>7.4</b>
<b>Pemetrexed</b>	<b>43</b>	<b>13.9</b>	<b>2.3</b>	<b>8.05</b>
<b>Docetaxel</b>	<b>27</b>	<b>8.7</b>	<b>3.8</b>	<b>7.0</b>
Gemcitabine	22	4.5	2.1	6.5
Bevacizumab	46	10.9	3.4	7.29

**Non-platinum PFS: 2-3 M OS: 6-7 M**

<sup>1</sup> [Yu et al., Am J Hematol Oncol 2015;11:27-31](#)

# Reactivation of anti-tumor immunity through PD-1/L1 pathway blockade



# PD-1/L1 pathway inhibition is effective against highly immunogenic tumors

## KEYNOTE-158 (NCT02628067) Cohort K – MSI-H Solid Tumors

**Key Eligibility Criteria**

- Any advanced solid tumor that is MSI-H<sup>a</sup>, excluding colorectal cancer
- Progression on or intolerance to ≥1 line of standard therapy for unresectable and/or metastatic disease
- Measurable disease per RECIST v1.1
- ECOG PS 0 or 1
- Provision of a tumor sample for biomarker assessments

N = 351<sup>b</sup>

**Pembrolizumab 200 mg IV Q3W  
Up to 35 cycles<sup>c</sup>**

**Primary Endpoint**

- ORR per RECIST v1.1 by BICR

**Secondary Endpoint**

- DOR and PFS per RECIST v1.1 by BICR
- OS
- Safety

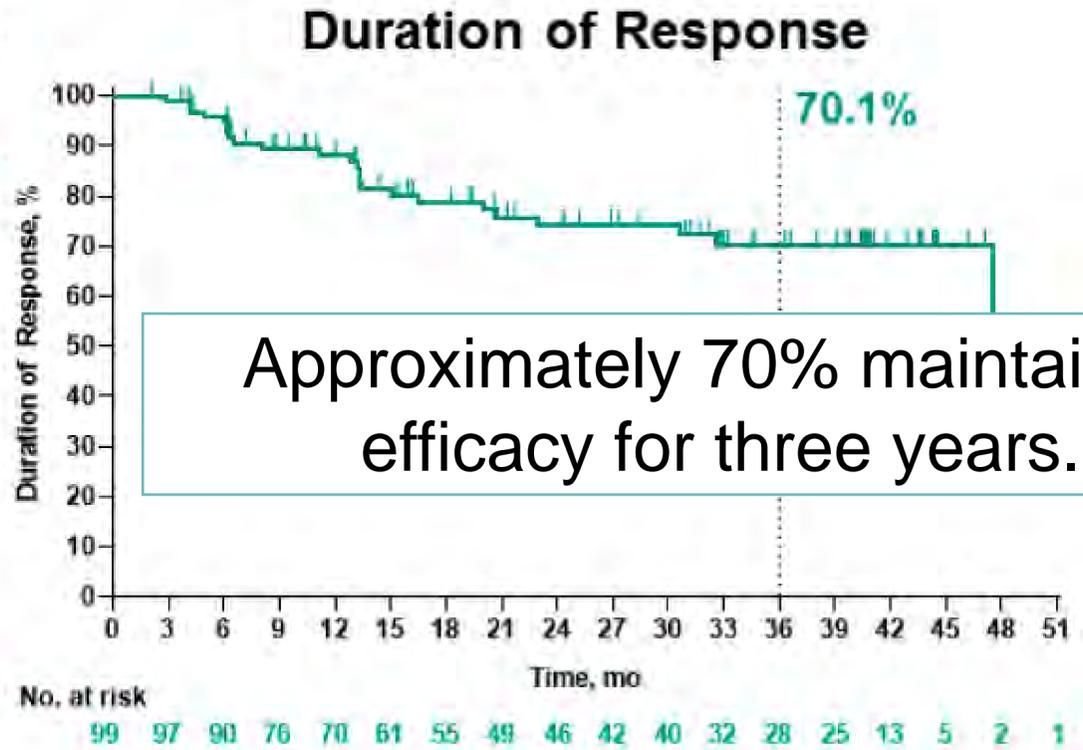
**Statistical Analysis Details**

- Efficacy assessed in all patients who received ≥1 dose of treatment with ≥6 mo follow-up
- Safety assessed in all treated patients

The effect lasts for a long time

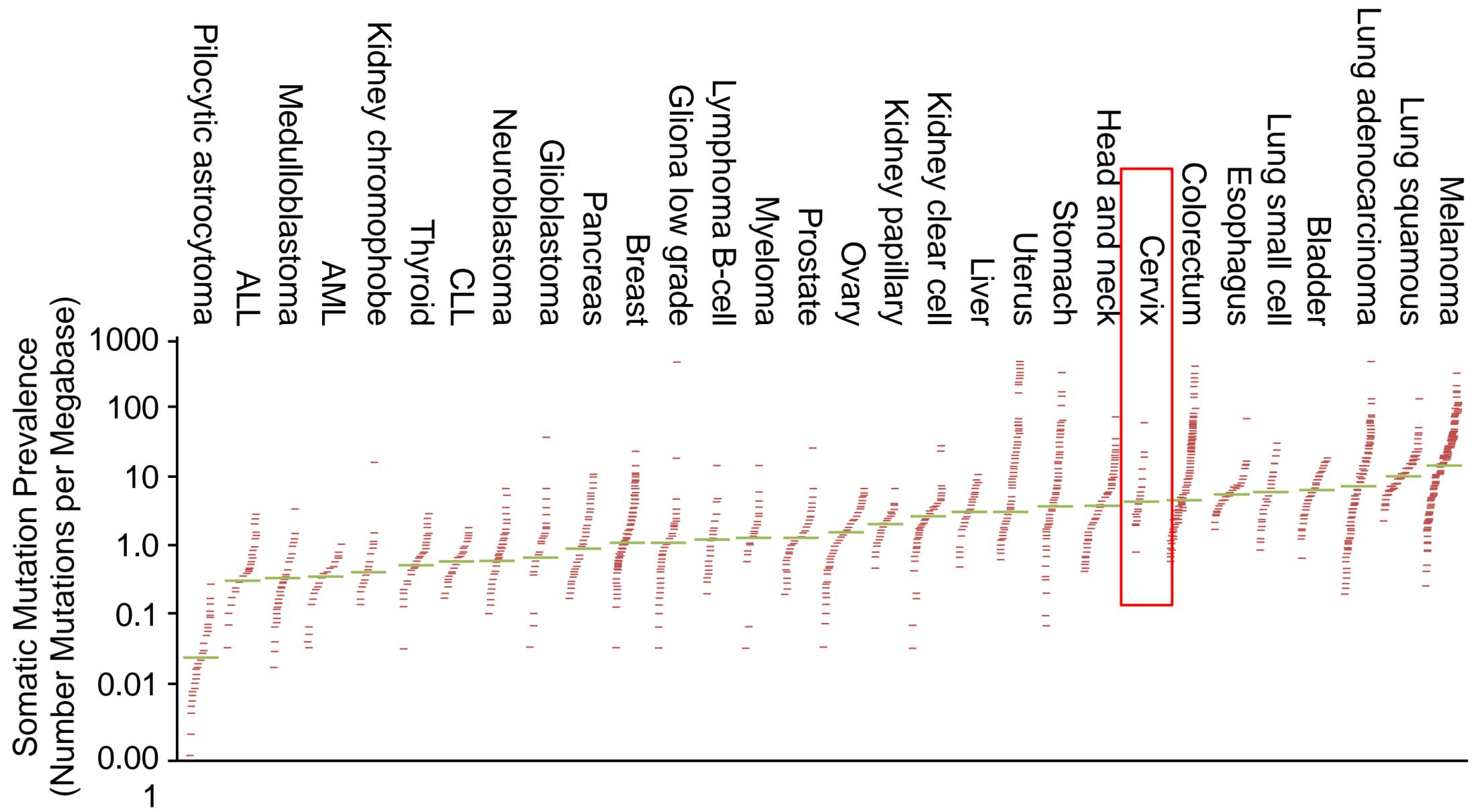
DOR is long

Efficacy Analysis Population	N = 321
<b>ORR, % (95% CI)</b>	<b>30.8 (25.8–36.2)</b>
CR	27 (8.4)
PR	72 (22.4)
SD	61 (19.0)
PD	131 (40.8)
Nonevaluable	3 (0.9)
No assessment <sup>a</sup>	27 (8.4)

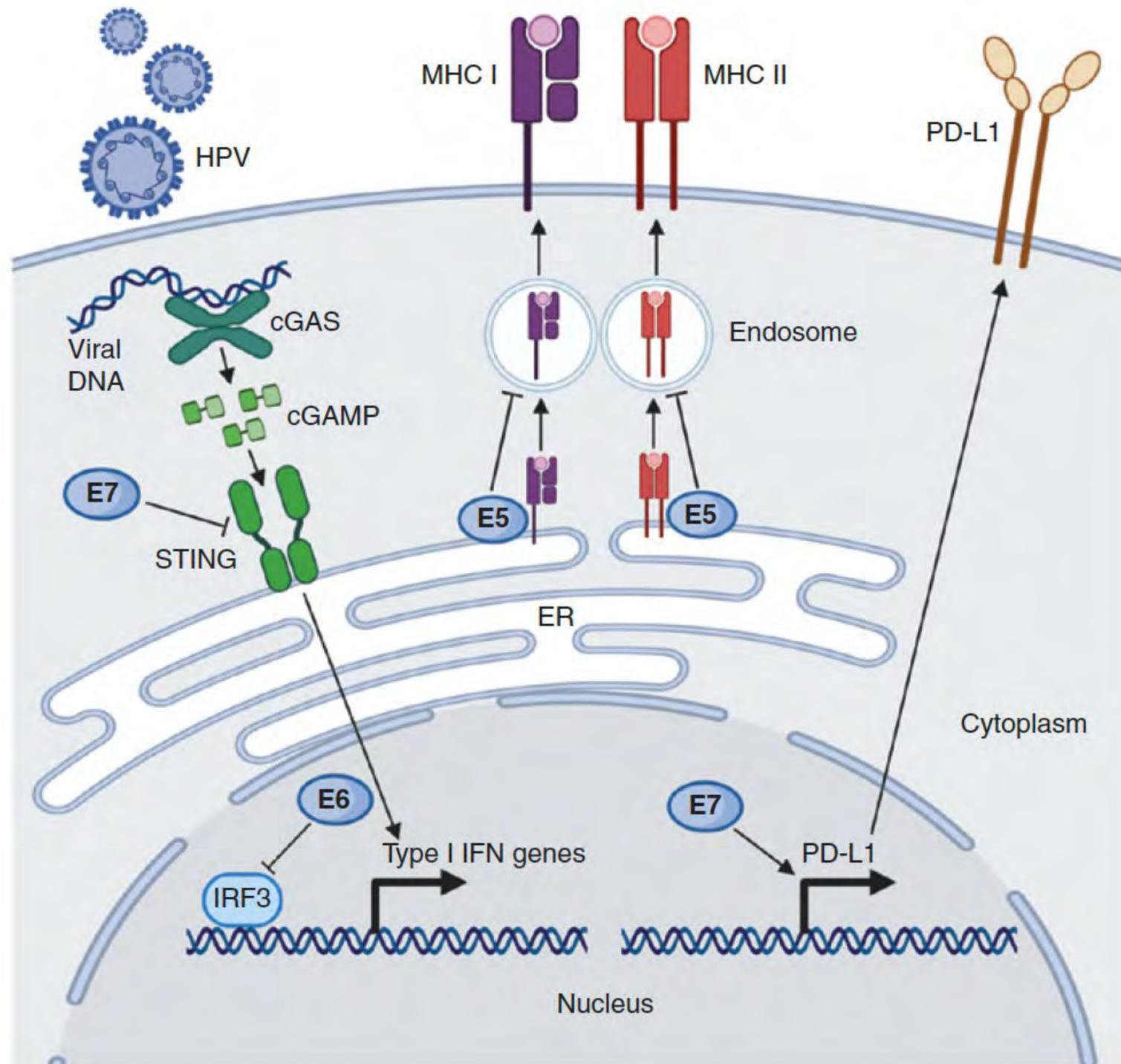




# Mutational Burden Compared With Other Tumors



# Cervical cancer is a target for immunotherapy



## Virus-related (HPV)

- Viral antigens also targeted
- PD-L1 Expression
- Activation of innate immune pathways

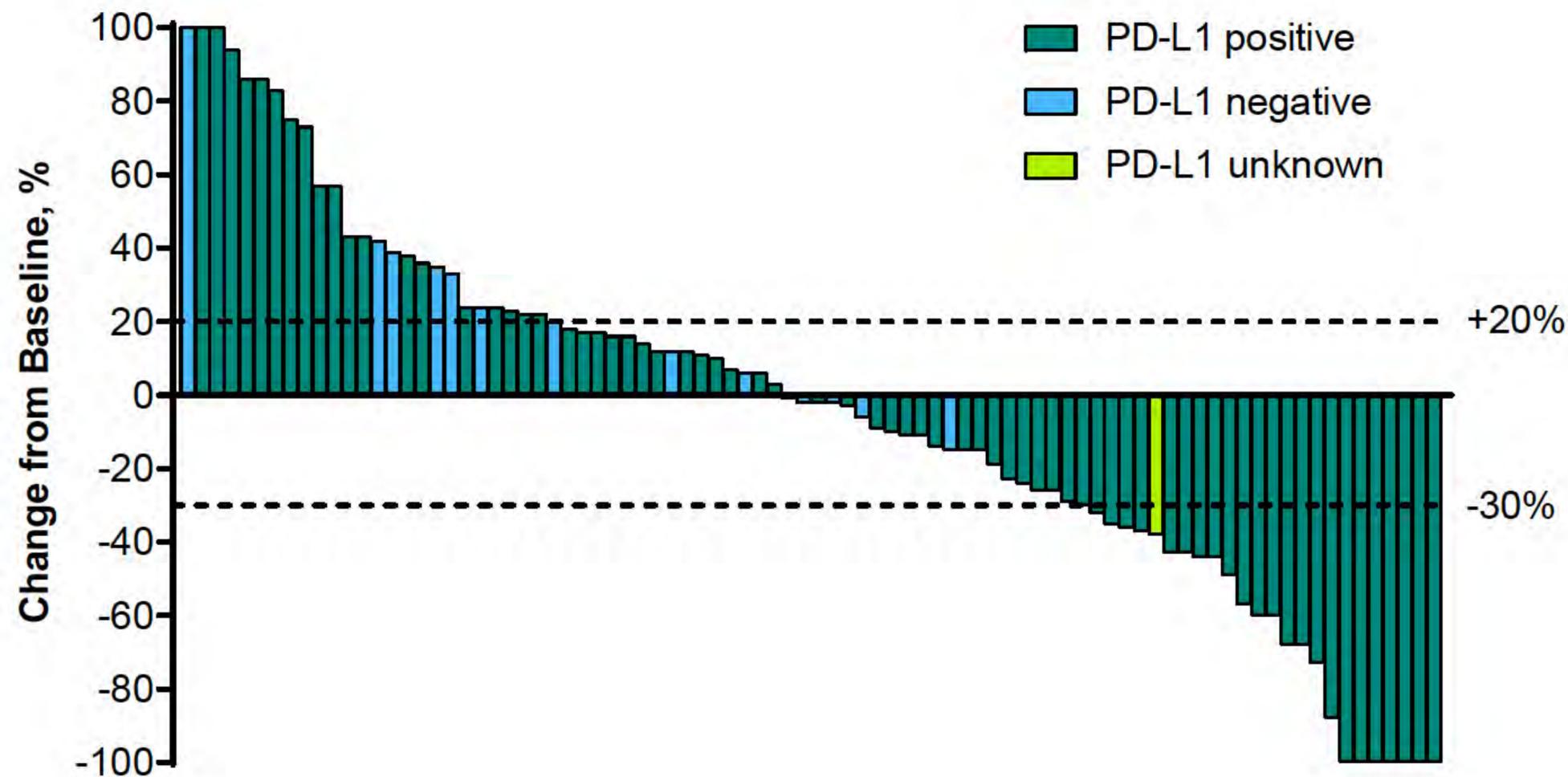
# Early Development of PD-1/L1 Pathway Inhibition for Cervical Cancer

Target	Drug (Study)	Ph	N. of pts	CR	PR	SD	ORR (%)	DCR (%)
PD-1	Pembrolizumab (KEYNOTE-028)	Ib	24	0	4	3	17	-
PD-1	Pembrolizumab (KEYNOTE-158)	II	98	3	9	18	12.2	-
PD-1	Nivolumab (CheckMate358)	II	19	1	4	8	26.3	68.4
PD-1	Nivolumab (ONO-4538-39)	II	20	0	5	10	25	75
PD-1	Nivolumab (NRG-GY002)	II	25	0	1	9	4	36
PD-1	Cemiplimab (R2810-1423)	Ib	10	0	1	3	10	40

- KEYNOTE-158 showed an ORR of 14.6% in PD-L1 positive cases, 0% in negative cases
- ONO-4538-39 showed an ORR of 33% in PD-L1 positive cases, 0% in negative cases

# Early Development of PD-1/L1 Pathway Inhibition for Cervical Cancer

## KEYNOTE-158

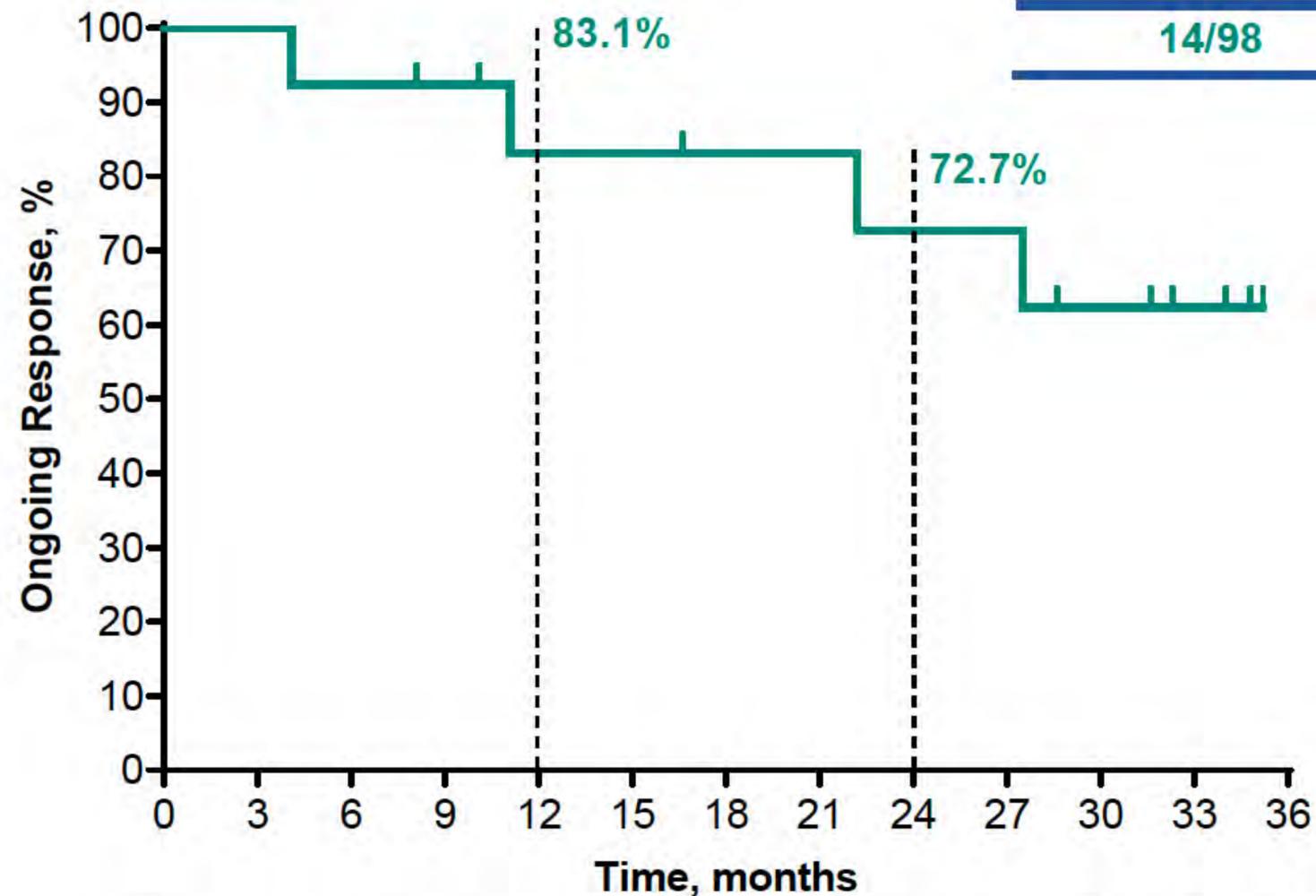


- ORR:14.3%
- PD-L1 +:17.1%
- PD-L1 -:0%
- TMB-h:31.1%
- non-TMB-h:11.9%

Includes patients with  $\geq 1$  evaluable post-baseline tumor assessment (n = 86). Data cutoff date: June 27, 2019.

# KEYNOTE-158

## DOR



Responders, n/N	Median, months (95% CI)
14/98	NR (3.7+ to 35.2+)

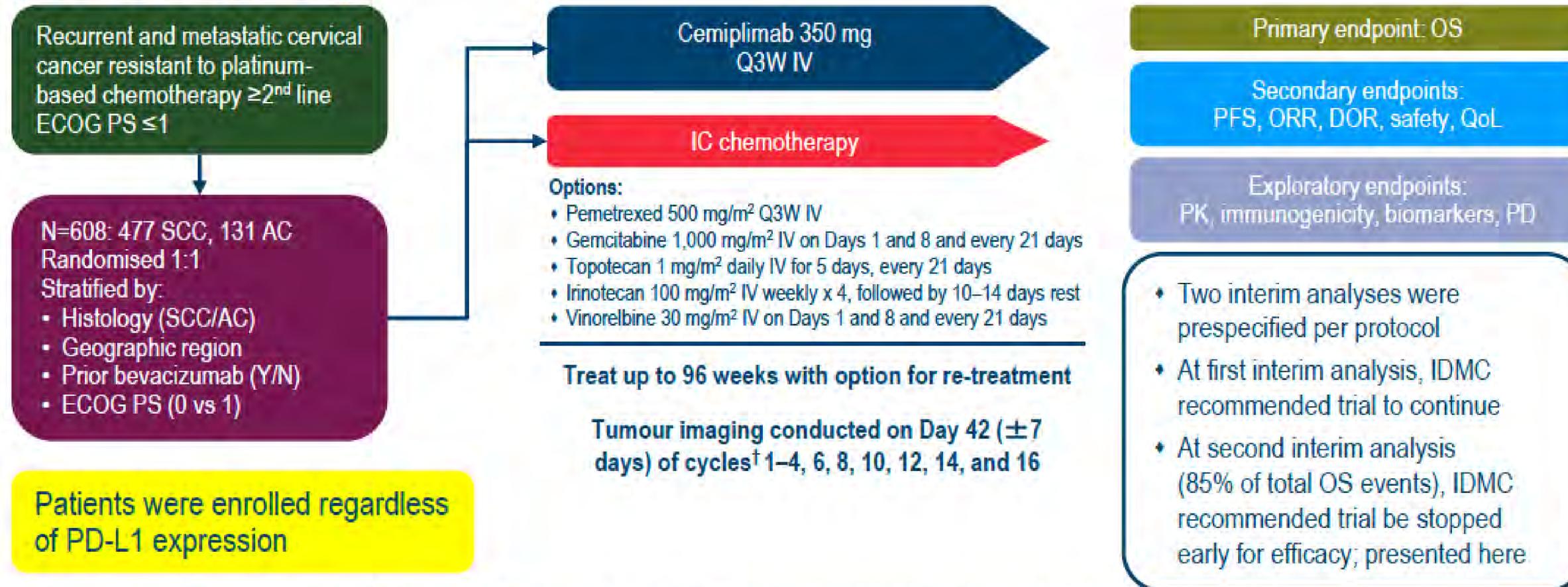
- Over 70% of cases showing efficacy maintained that response for more than two years.

No. at risk 14 14 12 11 9 9 8 8 7 7 5 3 0

Includes patients with best overall response of complete or partial response (n = 14). NR = Not Reached. "+" indicates there is no progressive disease by the time of last disease assessment. Data cutoff date: June 27, 2019.

# EMPOWER-CERVICAL 1

## EMPOWER-CERVICAL 1/GOG-3016/ENGOT-CX9 STUDY DESIGN\* (NCT03257267)



\*Performed according to ENGOT Model C.<sup>1†</sup>To account for differences in drug administration schedules, one cycle is defined as 6 weeks.

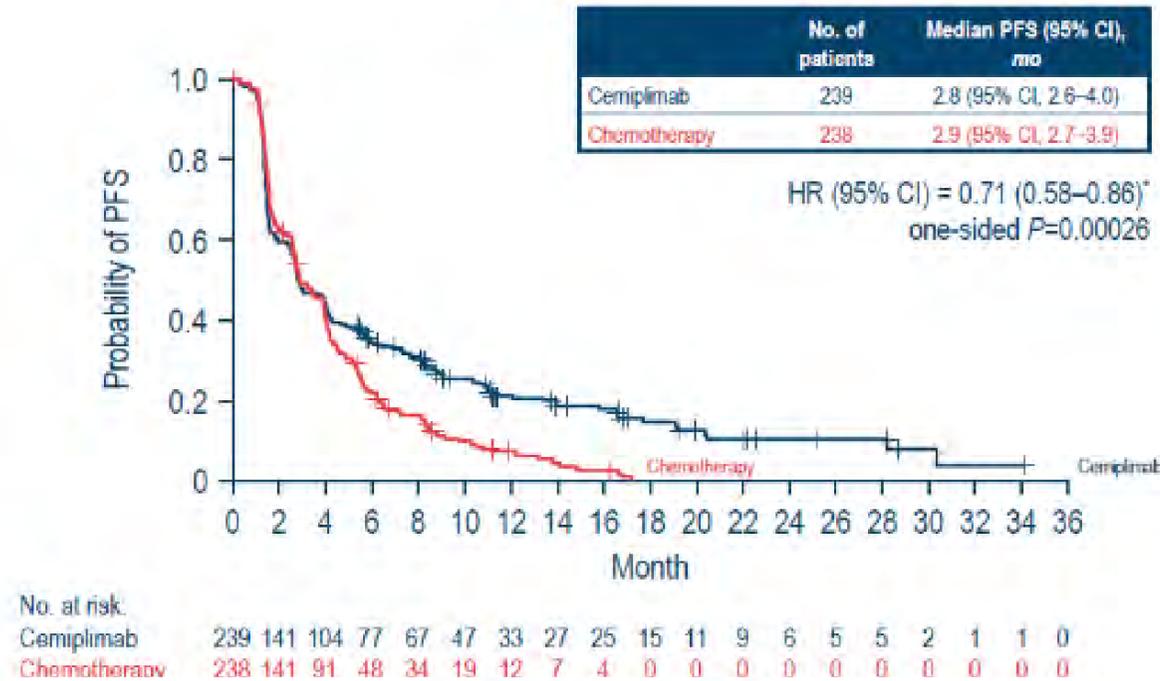
AC, adenocarcinoma or adenosquamous carcinoma; DOR, duration of response; ECOG PS, Eastern Cooperative Oncology Group performance status; IC, investigator's choice; IDMC, Independent Data Monitoring Committee; IV, intravenously; ORR, objective response rate; OS, overall survival; PD, pharmacodynamics; PD-L1, programmed cell death ligand 1; PFS, progression-free survival; PK, pharmacokinetics; Q3W, every 3 weeks; QoL, quality of life; SCC, squamous cell carcinoma.

1. Vergote I et al. *Int J Gynecol Cancer*. 2019;0:1–4.

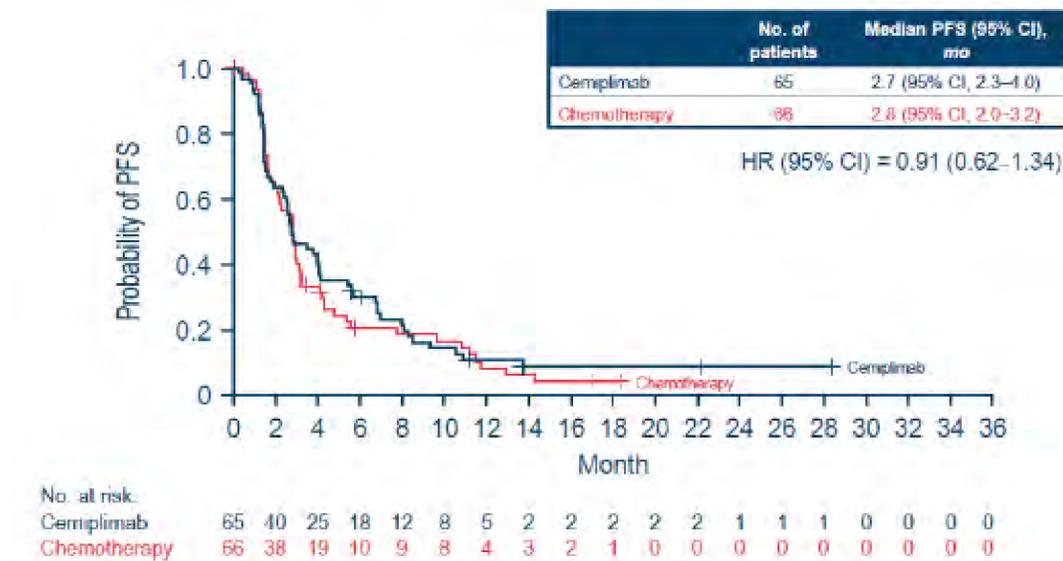
# EMPOWER-CERVICAL 1

## PROGRESSION-FREE SURVIVAL

### SCC Population

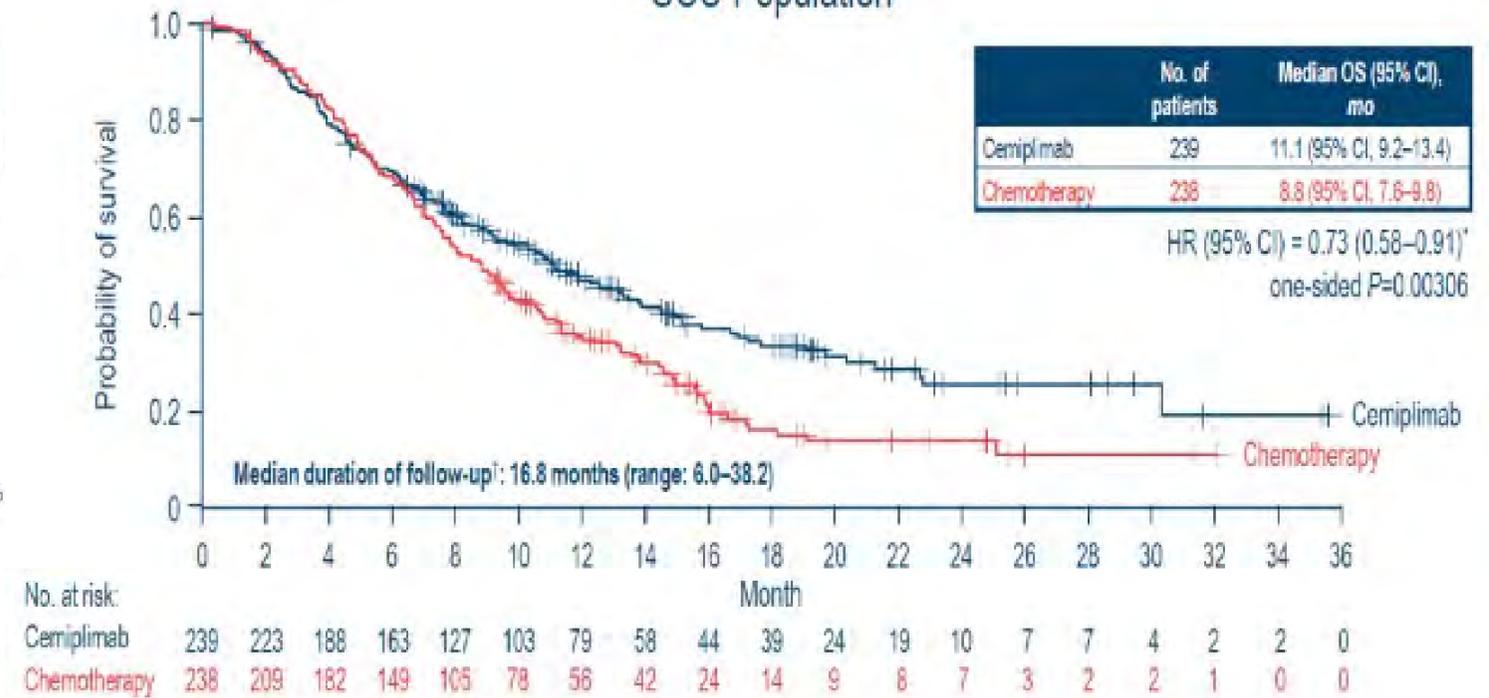


### AC Population



## OVERALL SURVIVAL

### SCC Population



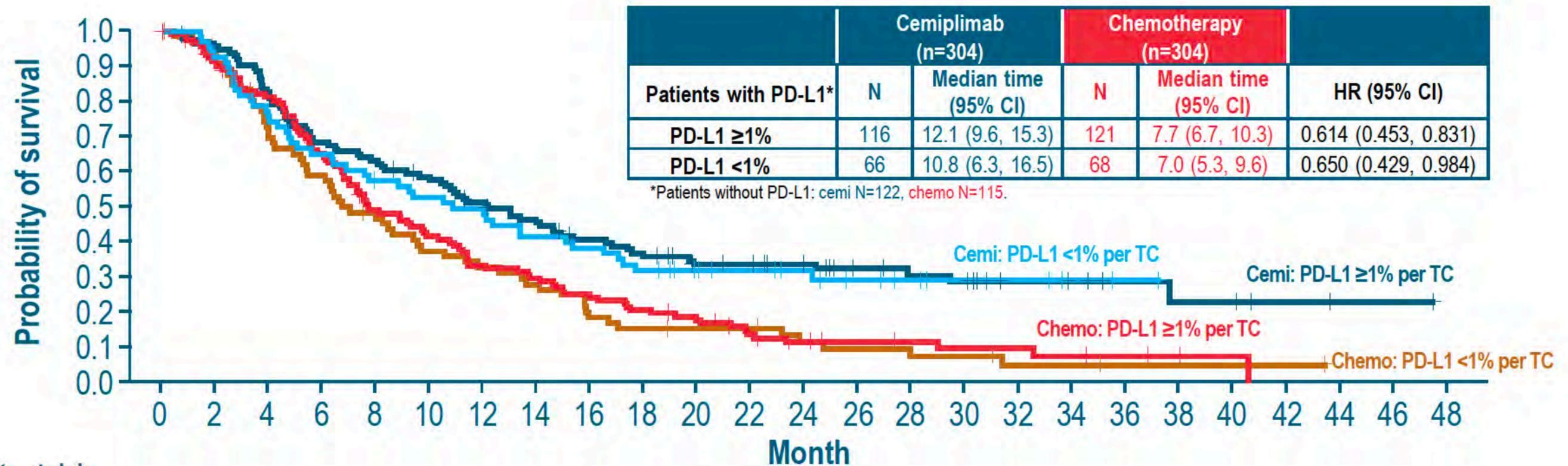
### AC Population





# EMPOWER-CERVICAL 1

**Cemiplimab monotherapy significantly improved OS vs chemotherapy regardless of PD-L1 status**

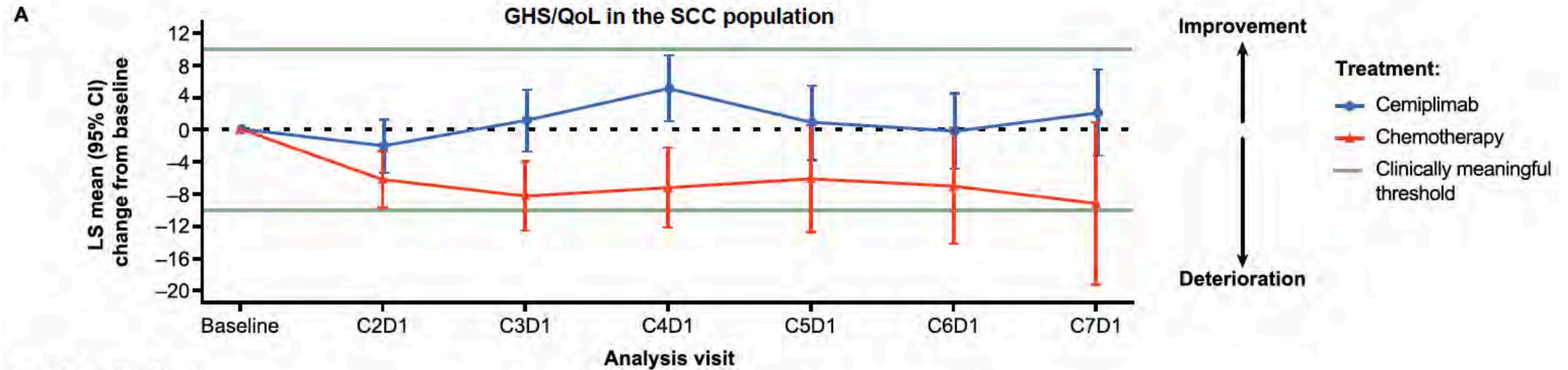


Patients at risk		0	2	4	6	8	10	12	14	16	18	20	22	24	26	28	30	32	34	36	38	40	42	44	46	48
Cemi: PD-L1 ≥1% per TC	116	110	93	77	71	63	55	48	41	36	30	29	25	20	17	16	10	9	5	4	4	2	1	1	0	
Cemi: PD-L1 <1% per TC	66	61	49	43	36	33	30	26	24	20	16	14	12	9	7	5	5	3	1	0	0	0	0	0	0	
Chemo: PD-L1 ≥1% per TC	121	107	92	73	54	46	37	33	27	23	19	13	9	7	6	5	5	4	3	2	1	0	0	0	0	
Chemo: PD-L1 <1% per TC	68	60	46	39	30	24	21	18	12	10	9	9	6	5	4	4	2	2	1	1	1	1	0	0	0	

Kaplan–Meier curves of overall survival in the full analysis set. **Data cutoff date: 4 Jan 2022**

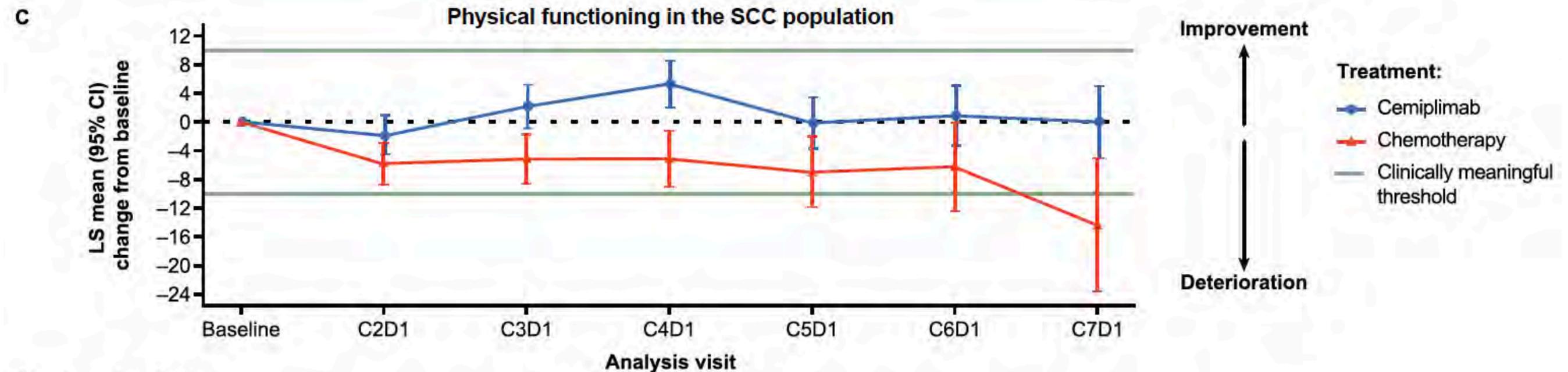
Cemi, cemiplimab; Chemo, chemotherapy; CI, confidence interval; HR, hazard ratio; OS, overall survival; PD-L1, programmed cell death-ligand 1; TC, tumour cell; PD-L1 expression was detected with the SP263 monoclonal antibody (Ventana; Tewari et al., NEJM, 2022)

# Patient-reported QOL



**Number of patients**

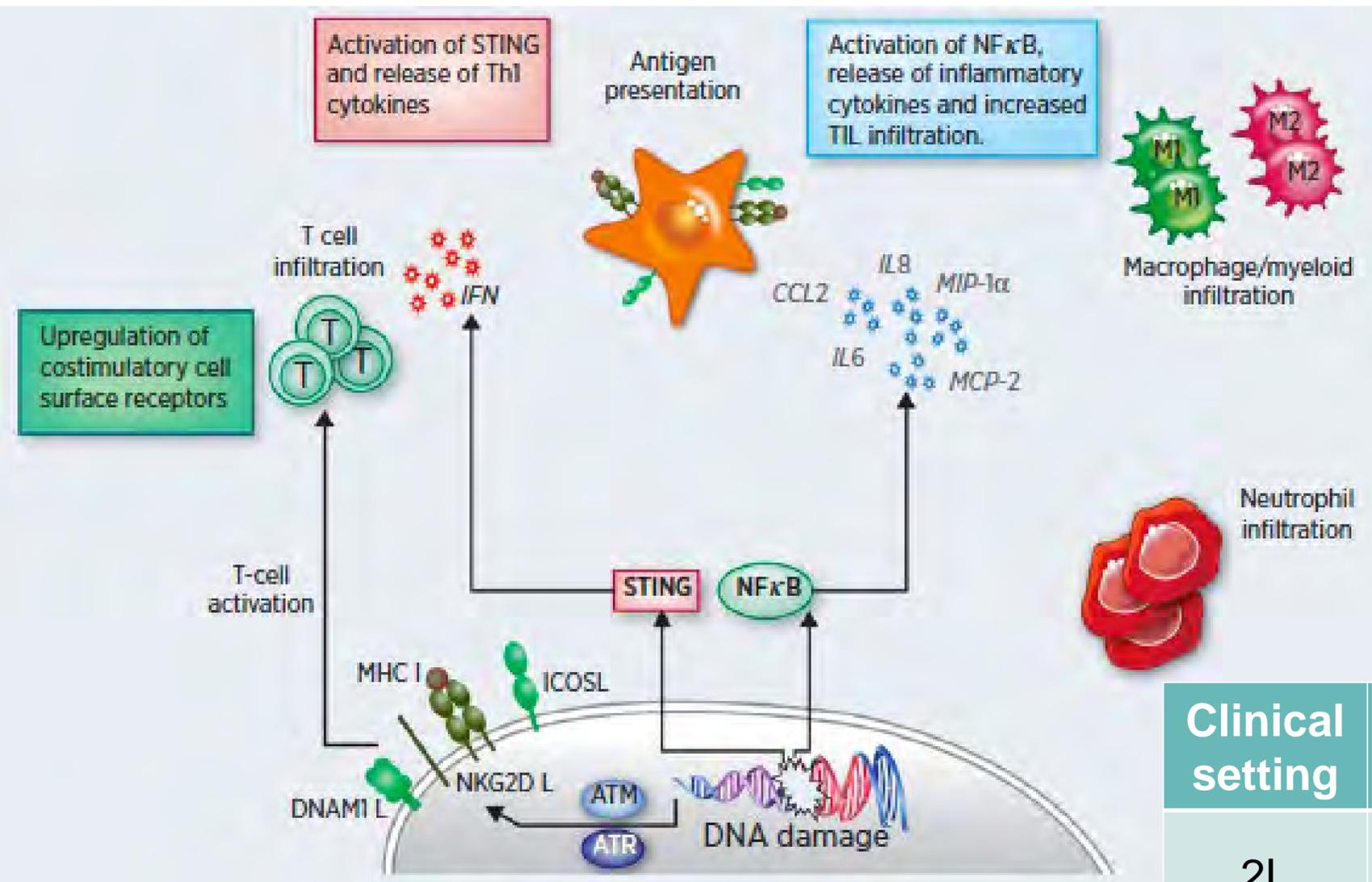
Cemiplimab	172	172	121	92	80	69	56
Chemotherapy	146	144	92	57	33	27	12



**Number of patients**

Cemiplimab	172	172	121	92	80	69	56
Chemotherapy	146	144	92	58	33	27	12

# PD-1/L1 pathway inhibitor + PARPi combination

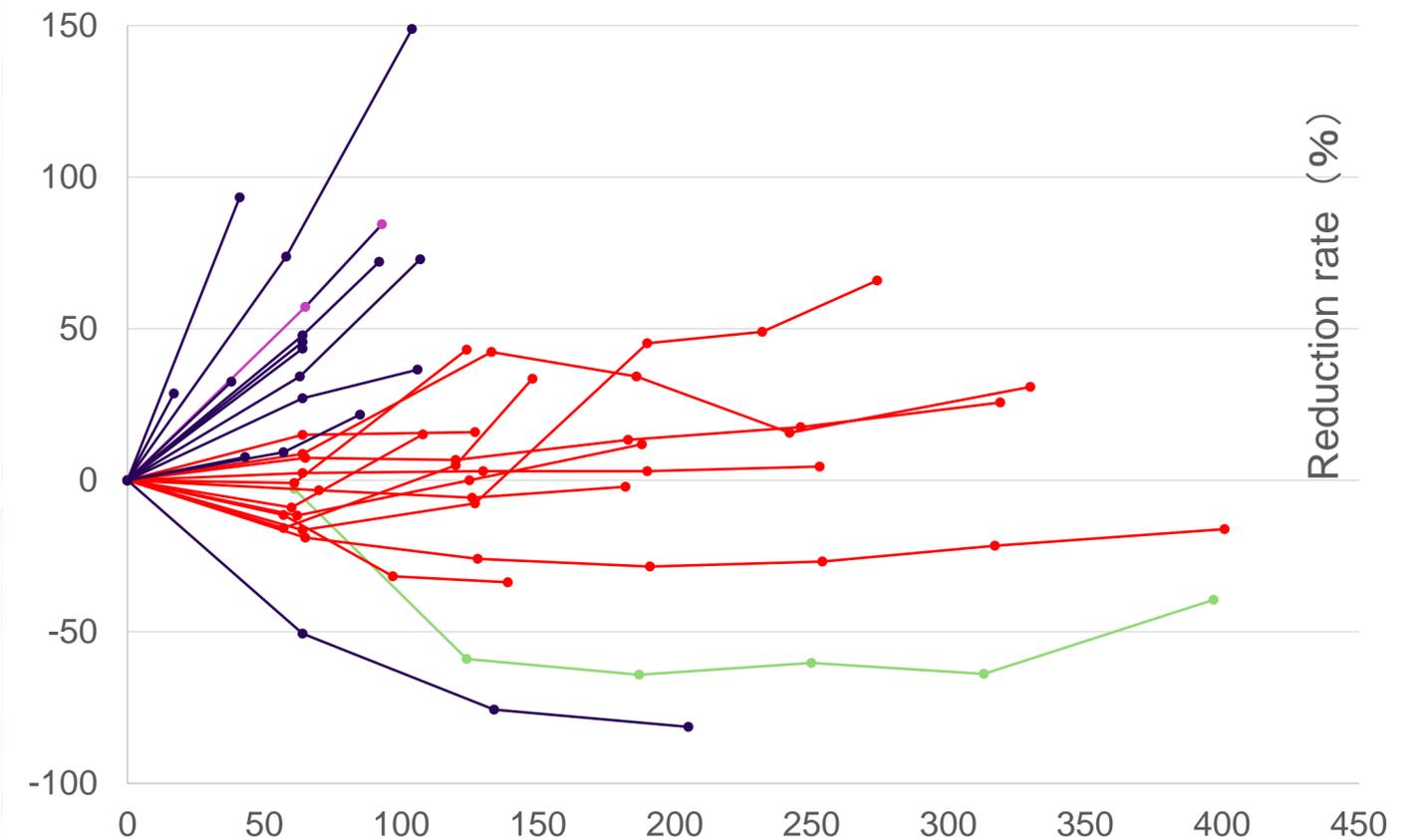
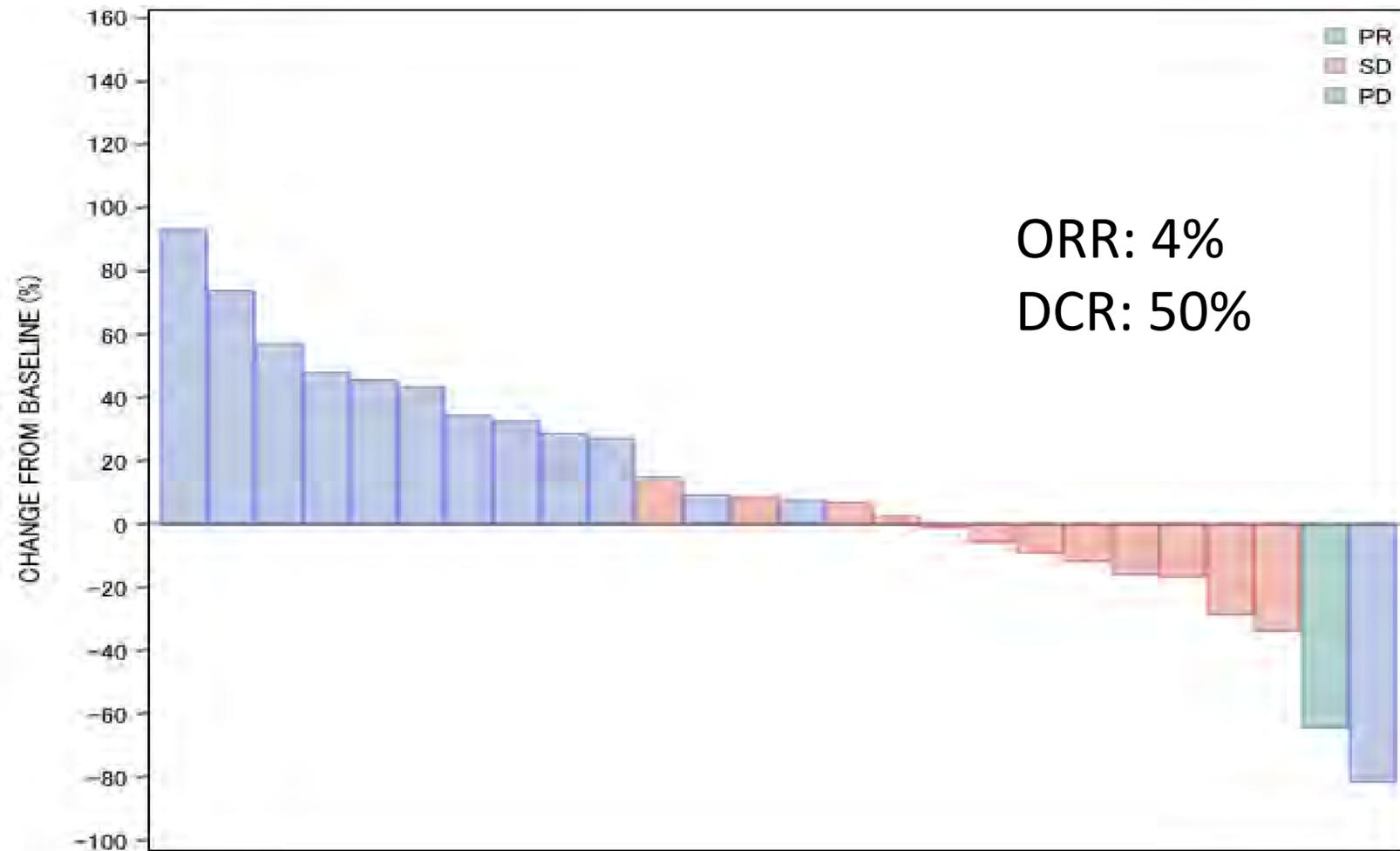


- PARPi induces immunomodulation
- cGAS/STING activation releases Th1 cytokines

Clinical setting	Study ID (Name)	Combination	N of pts	Endpoints
2L	NCT04068753	Dostal + Niraparib	66	•ORR •DOR
2L	NCT04483544	Pembro + Olaparib	48	•ORR •PFS
2L	NCT04641728 (GOTIC-025)	Pembro + Olaparib	28	•ORR •PFS

# Pembrolizumab+Olaparib for 2ndline CxCa

## GOTIC-025



Median PFS: 3.4M, median OS: 11.9M

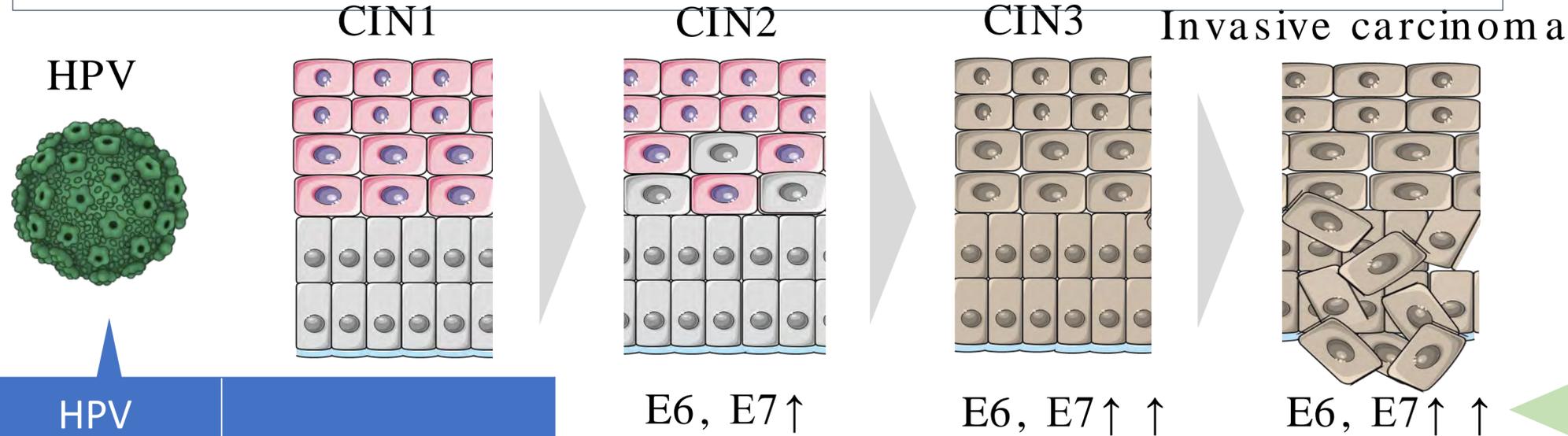


**Primary metastatic (4B),  
recurrent, persistent**

# Angiogenesis in Cervical Cancer

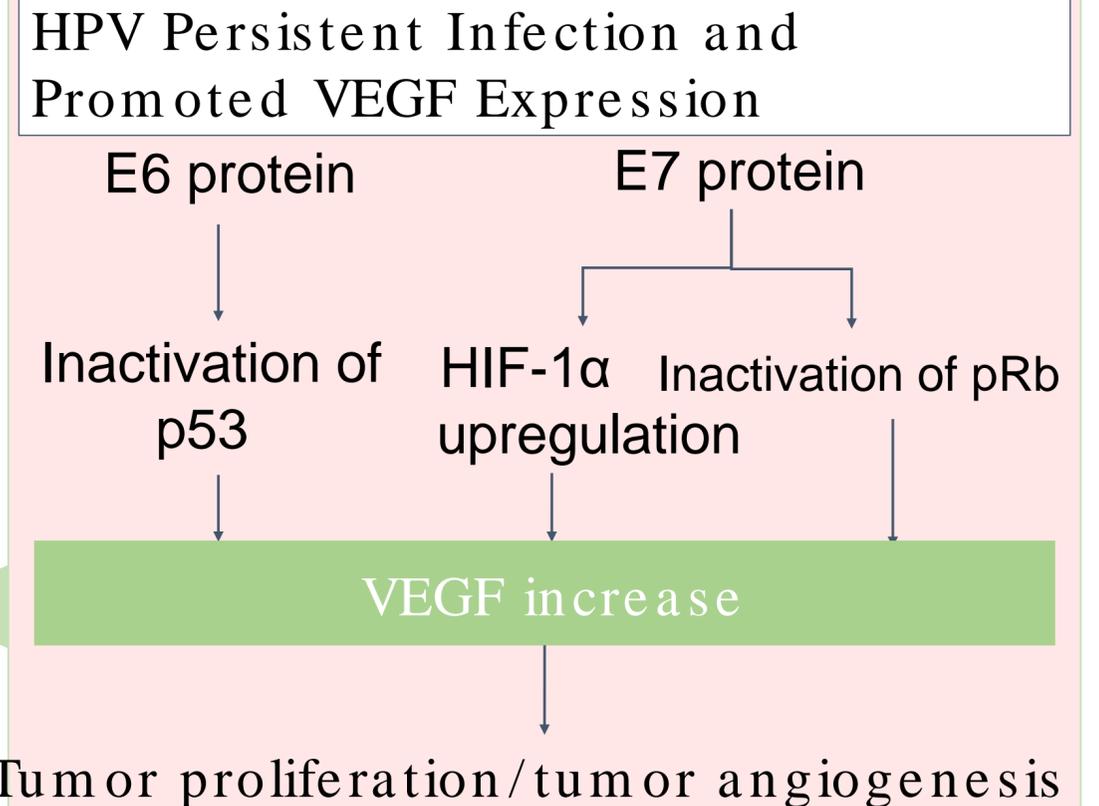
- In cervical cancer, the positivity rate for VEGF (Vascular Endothelial Growth Factor), which is involved in angiogenesis, is high at 72-94%. and it has been reported that high VEGF expression in resected cervical cancer tissue is a poor prognostic factor.

## Viral Genes and Molecular Biological Changes in the Carcinogenesis Mechanism of Cervical Cancer



HPV Gene Product	Function
E6, E7	Oncoprotein
E1, E2	Replication
L1, L2	Capsid synthesis

- Cell immortalization
- Loss of cell cycle control
- Promotion of epithelial-mesenchymal transition
- Apoptosis inhibition
- Immune response suppression
- Inhibition of cell differentiation
- Accumulation of genetic mutations

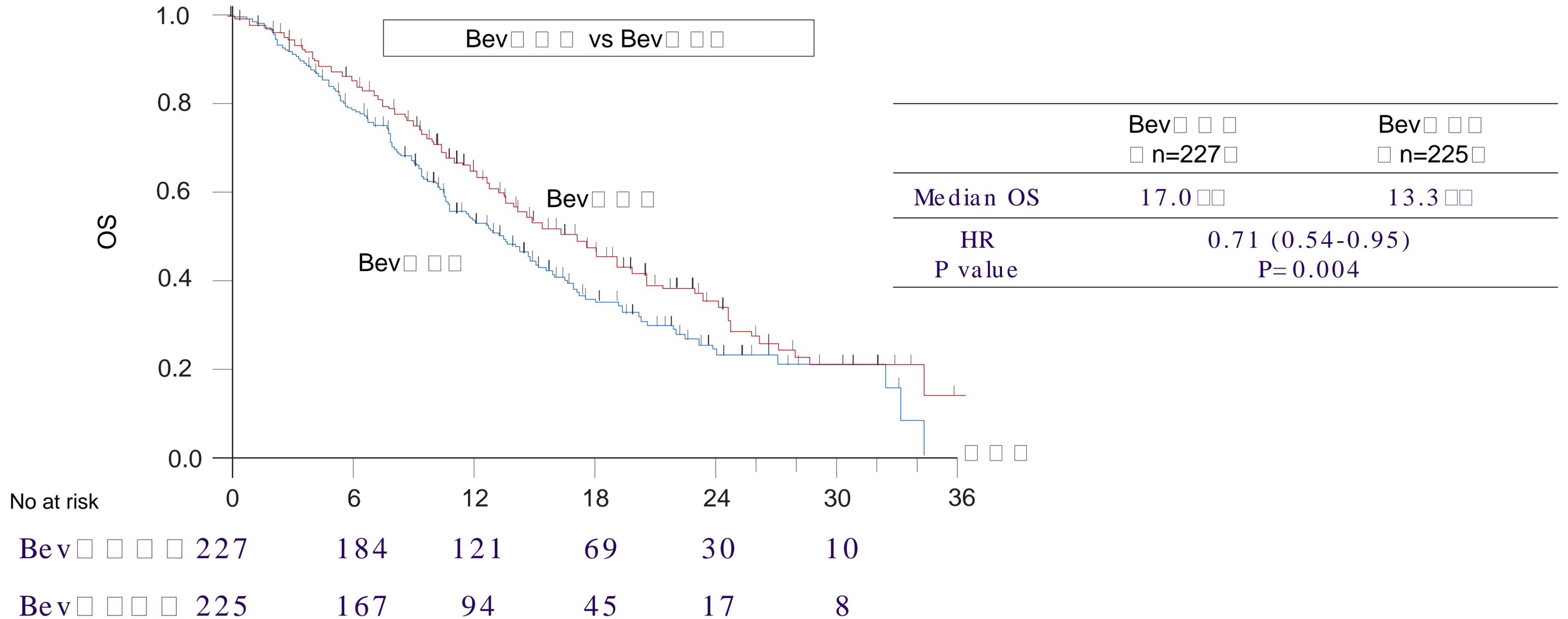


□ Bevacizumab □

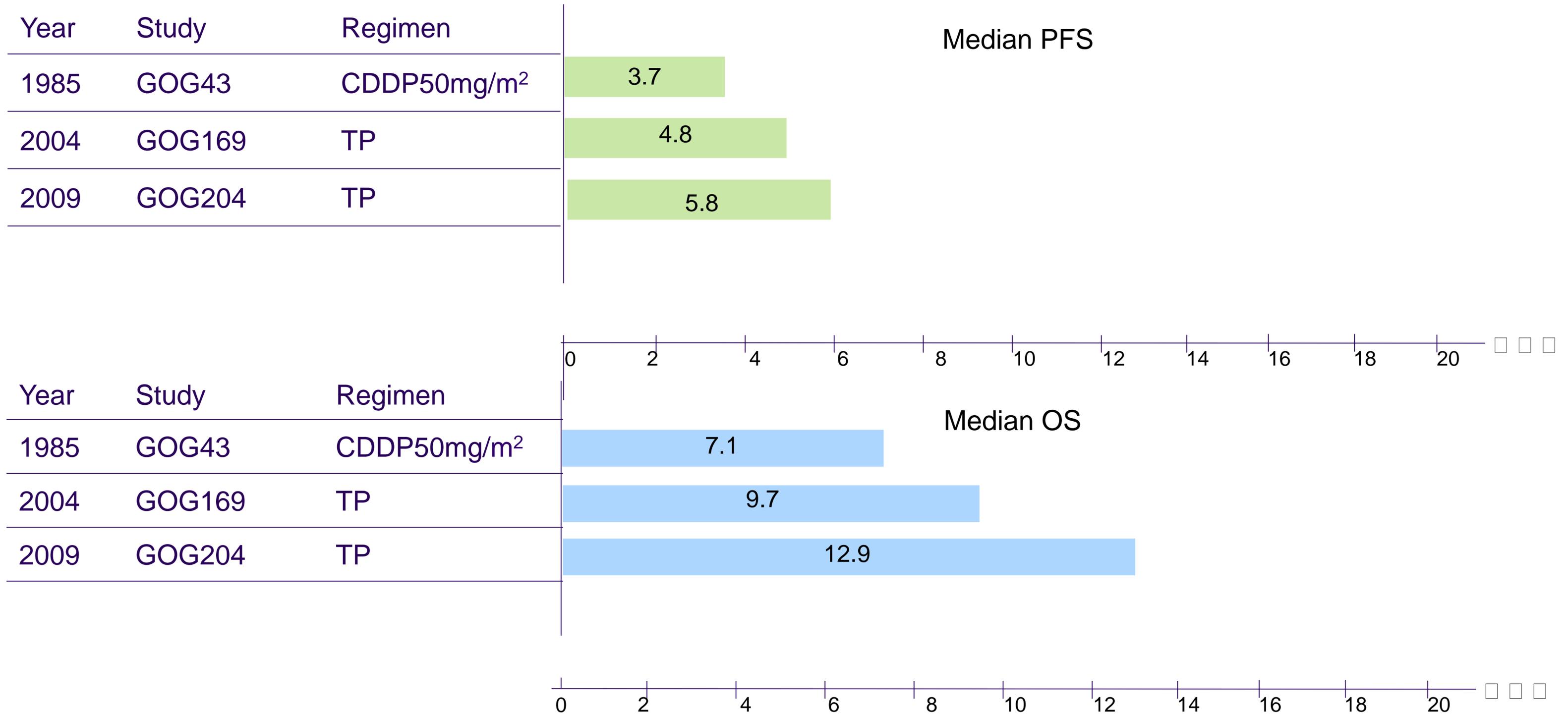
By specifically binding to VEGF, it inhibits VEGF and the binding of VEGF to its receptors on vascular endothelial cells

HPV: human papillomavirus; CIN: cervical intraepithelial neoplasia  
VEGF: vascular endothelial growth factor HIF-1α: hypoxia-inducible factor-1α

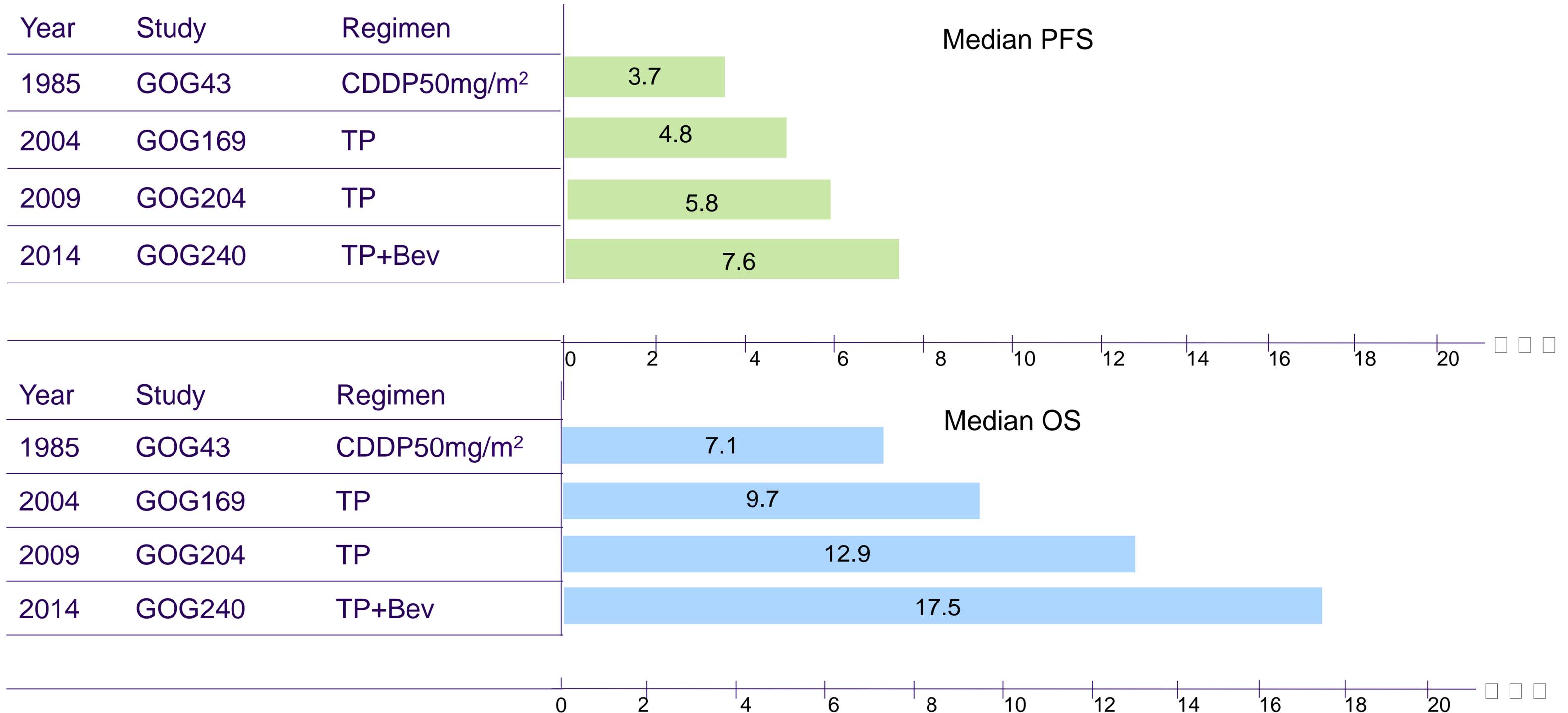
# GOG240: OS



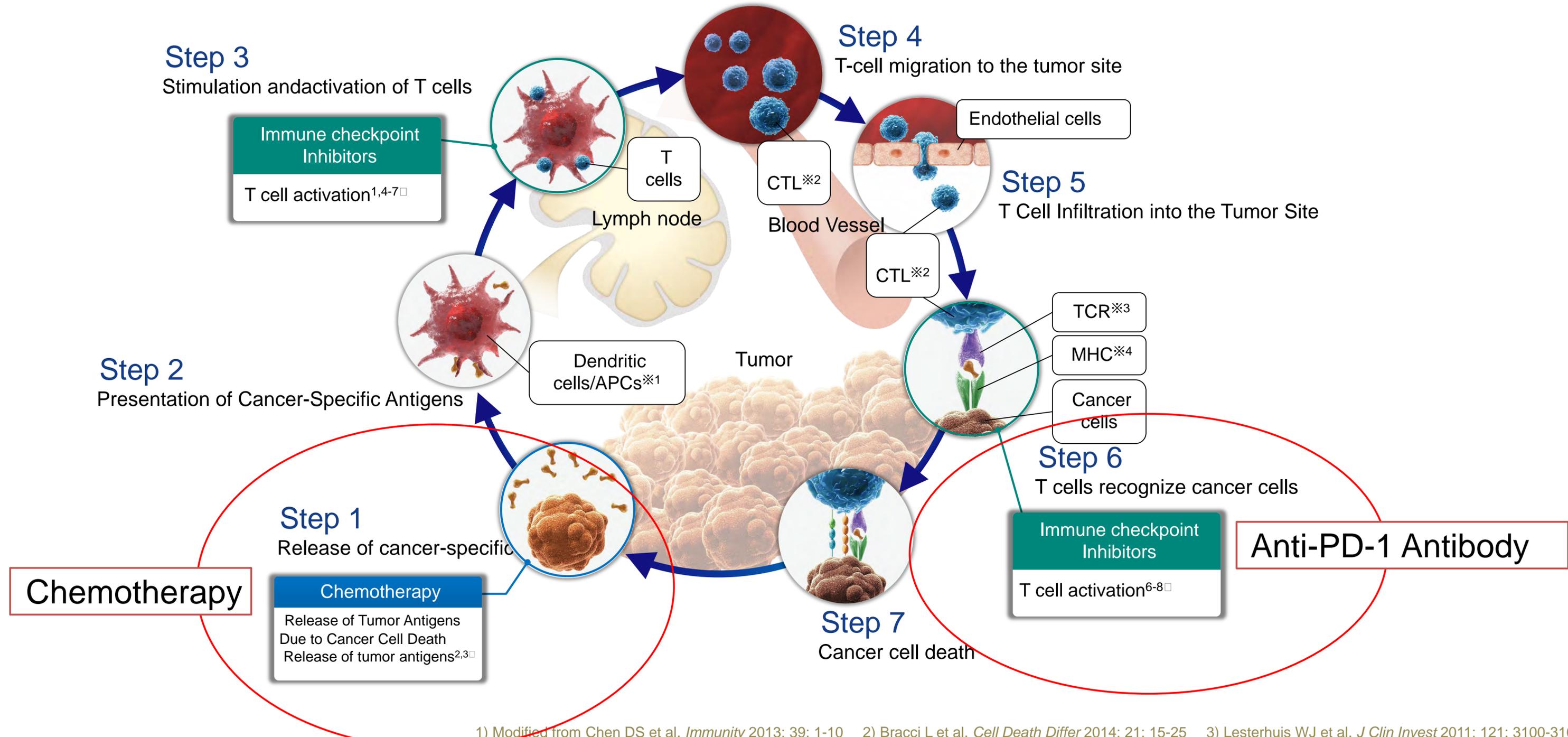
# Evolution of treatment for R/M cervical Cancer



# Evolution of treatment for R/M cervical Cancer



# Mechanisms of Action: Immune Checkpoint Inhibitors and Chemotherapy



※1 Antigen-presenting cells ※2 Cytotoxic T cells ※3 T cell receptor ※4 Major histocompatibility complex

1) Modified from Chen DS et al. *Immunity* 2013; 39: 1-10 2) Bracci L et al. *Cell Death Differ* 2014; 21: 15-25 3) Lesterhuis WJ et al. *J Clin Invest* 2011; 121: 3100-3108

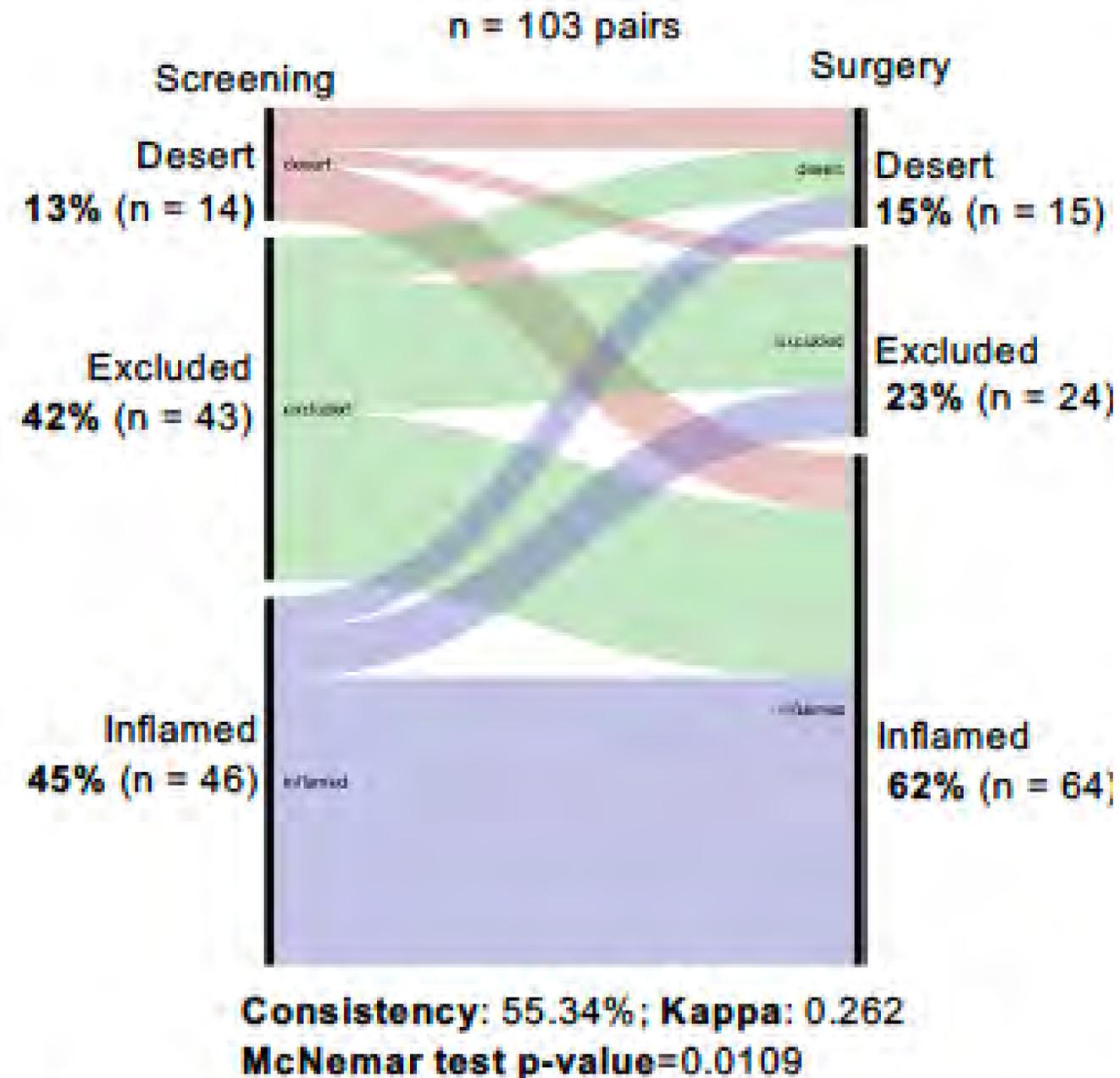
4) Clouthier DL et al. *Science* 2017; 355: 1373-1374 5) Mellman I et al. *Nature* 2011; 480: 480-489

6) Postow MA et al. *J Clin Oncol* 2015; 33: 1974-1982. Some authors received honoraria from MSD.

7) Pardoll DM. *Nat Rev Cancer* 2012; 12: 252-264 8) Topalian SL et al. *Curr Opin Immunol* 2012; 24: 207-212

# Chemotherapy may enhance local immunity

## Chemo+Bev



CD8/Pan-CK (IHC)

# KEYNOTE-826: Randomized, Double-Blind, Phase 3 Study

## Key Eligibility Criteria

- Persistent, recurrent, or metastatic cervical cancer not amenable to curative treatment
- No prior systemic chemotherapy (prior radiotherapy and chemoradiotherapy permitted)
- ECOG PS 0 or 1

## Stratification Factors

- Metastatic disease at diagnosis (yes vs no)
- PD-L1 CPS (<1 vs 1 to <10 vs ≥10)
- Planned bevacizumab use (yes vs no)

R  
1:1

**Pembrolizumab 200 mg IV Q3W**  
for up to 35 cycles

+

**Paclitaxel + Cisplatin or Carboplatin IV Q3W**  
for up to 6 cycles<sup>a</sup>

±

**Bevacizumab 15 mg/kg IV Q3W**

**Placebo IV Q3W**  
for up to 35 cycles

+

**Paclitaxel + Cisplatin or Carboplatin IV Q3W**  
for up to 6 cycles<sup>a</sup>

±

**Bevacizumab 15 mg/kg IV Q3W**

## End Points

- **Dual primary:** OS and PFS per RECIST v1.1 by investigator
- **Secondary:** ORR, DOR, 12-mo PFS, and safety
- **Exploratory:** PROs assessed per EuroQol EQ-5D-5L VAS

<sup>a</sup>Paclitaxel: 175 mg/m<sup>2</sup>. Cisplatin: cisplatin 50 mg/m<sup>2</sup>. Carboplatin: AUC 5 mg/mL/min. The 6-cycle limit was introduced with protocol amendment 2, although participants with ongoing clinical benefit who were tolerating chemotherapy could continue beyond 6 cycles after sponsor consultation.

CPS, combined positive score (number of PD-L1–staining cells [tumor cells, lymphocytes, macrophages] divided by the total number of viable tumor cells, multiplied by 100);

PROs, patient-reported outcomes; VAS, visual analog scale. KEYNOTE-826 ClinicalTrials.gov identifier, NCT03635567.

# Keynote 826

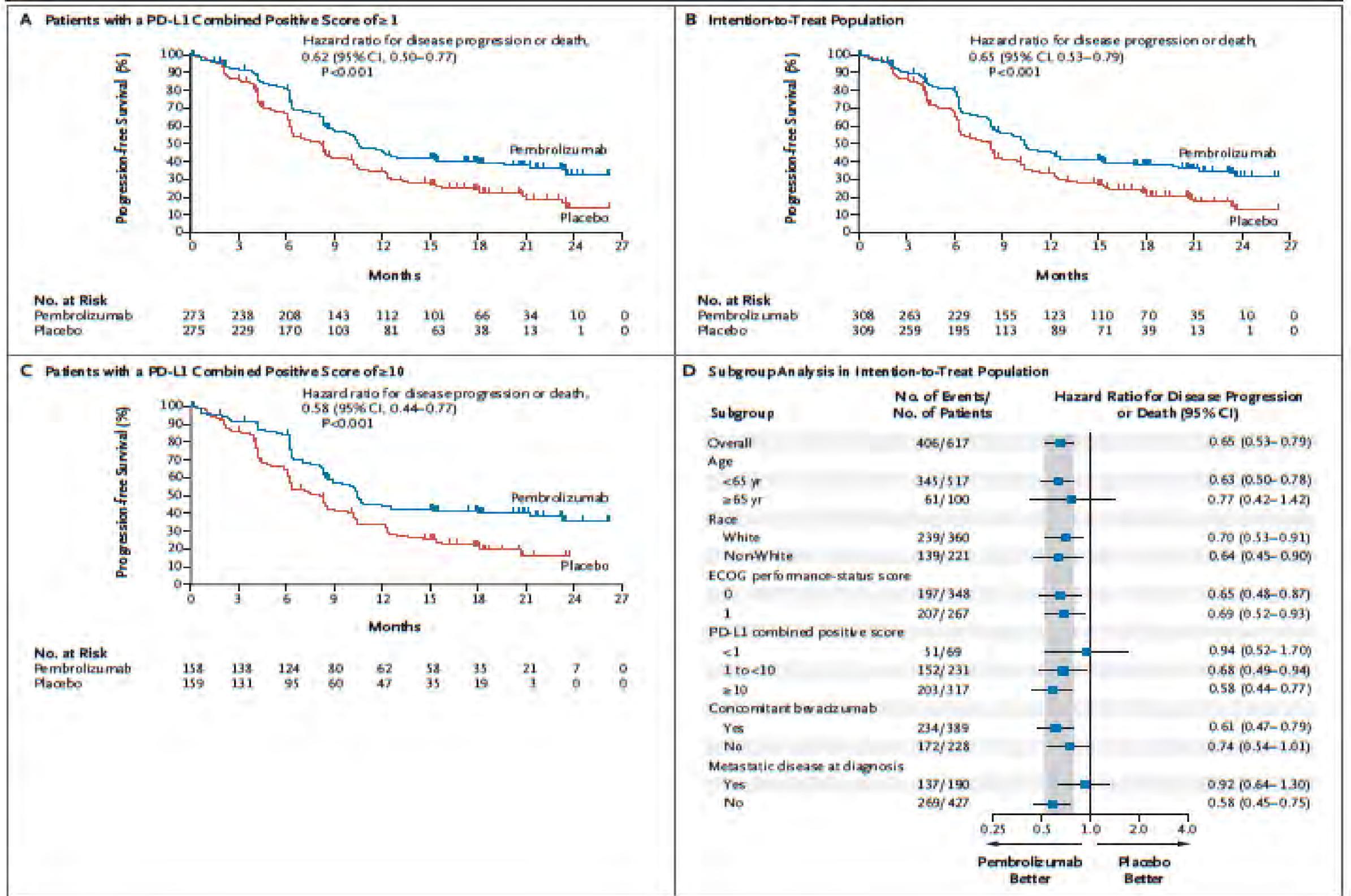
## PFS by PD-L1

### PFS HR

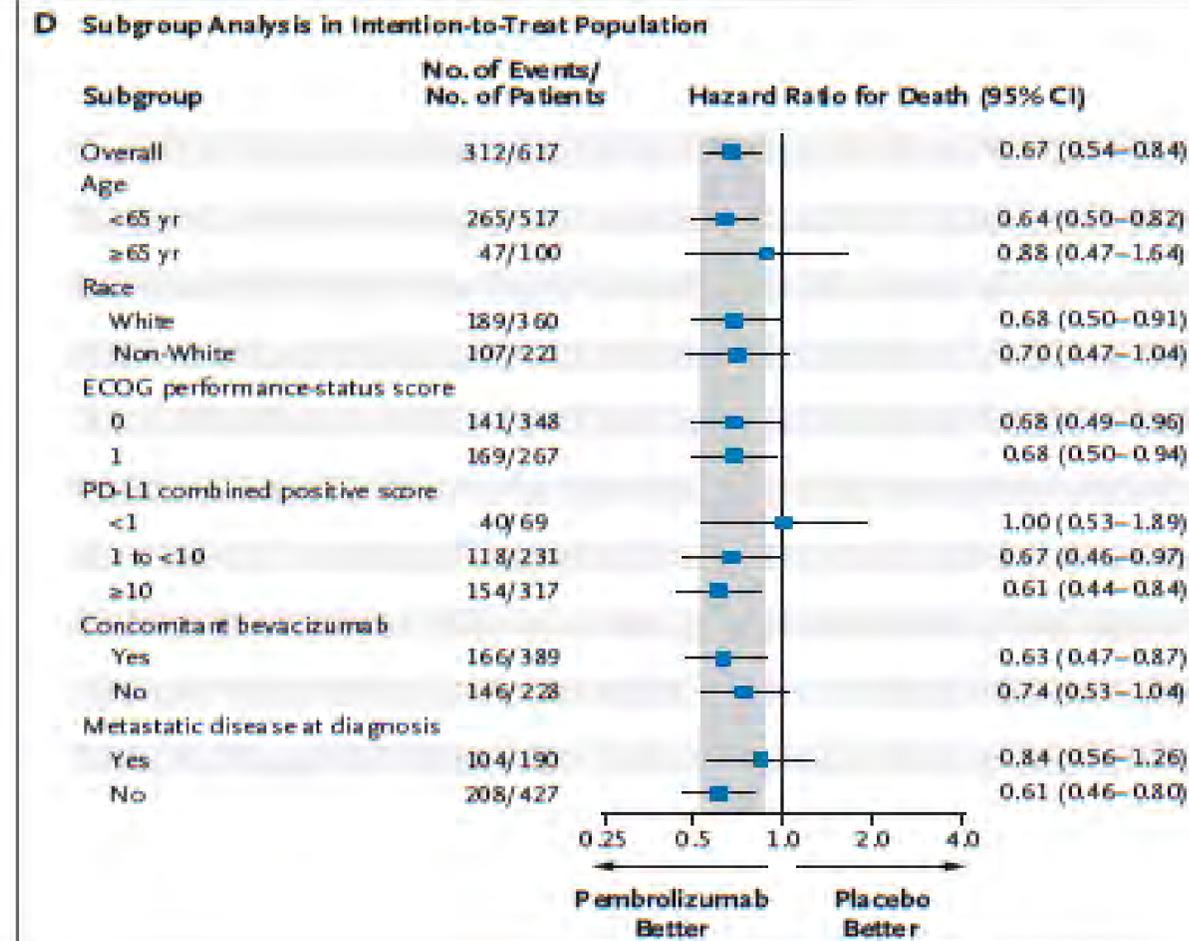
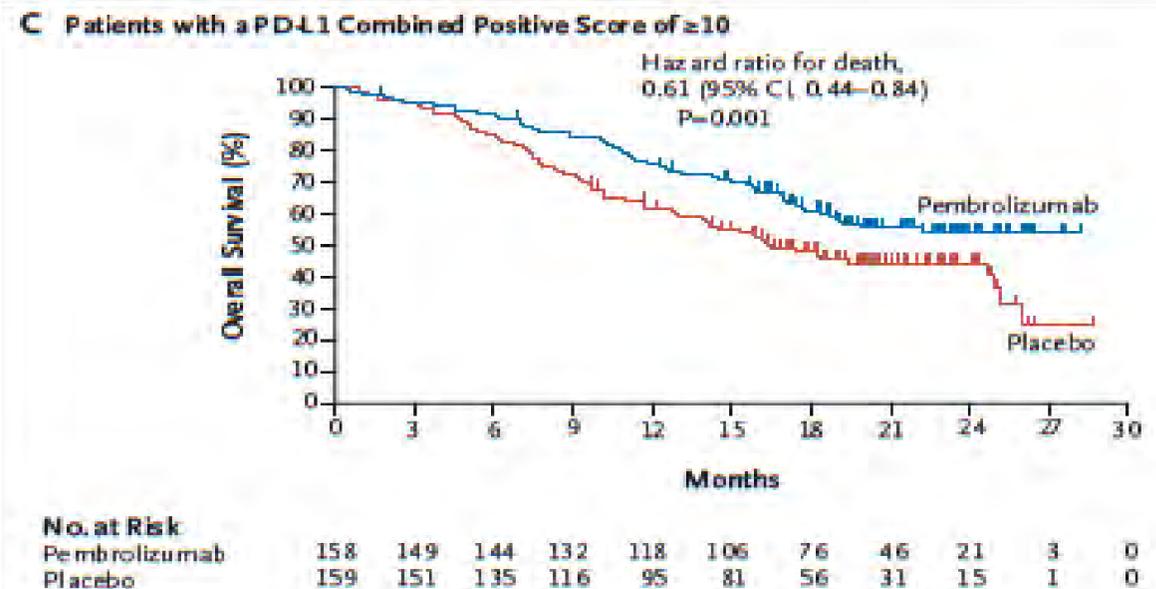
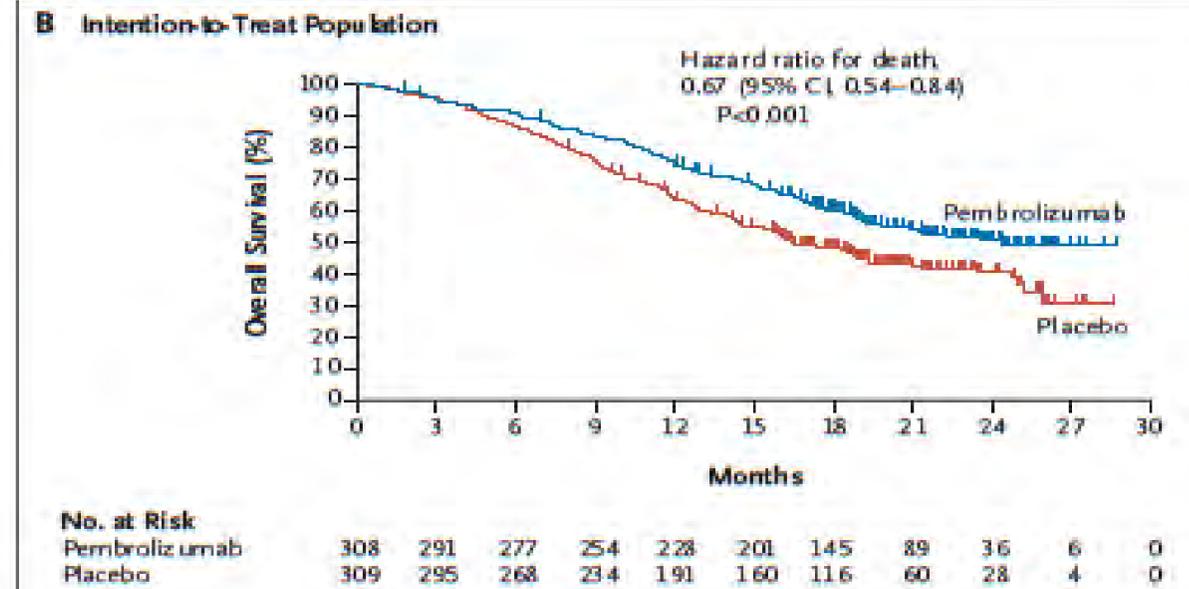
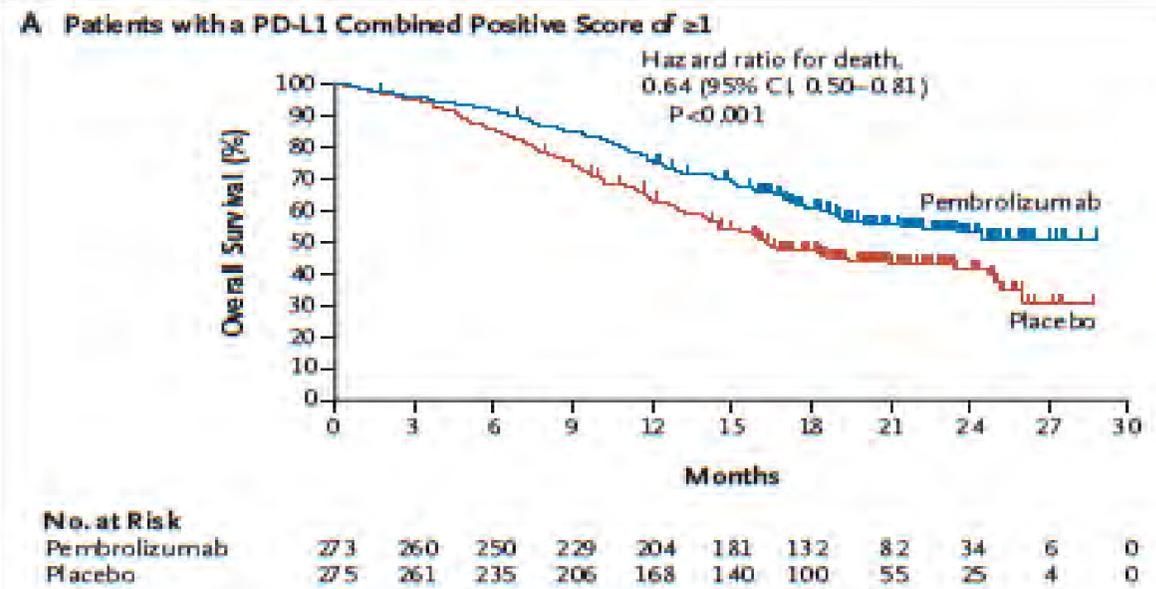
**CPS  $\geq 1$**       0.62

**ITT**      0.65

**CPS  $\geq 10$**       0.58



# Keynote 826 OS by PD-L1



**OS HR**

**CPS  $\geq 1$**       0.64

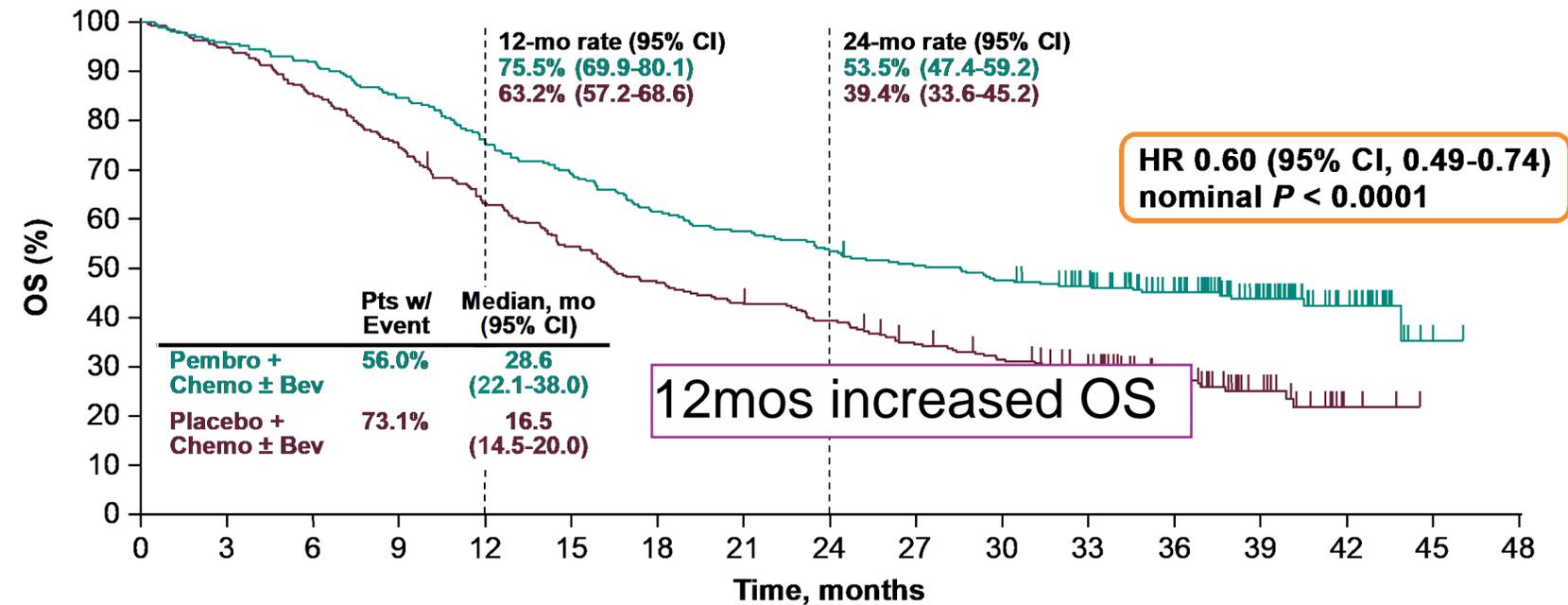
**ITT**              0.67

**CPS  $\geq 10$**       0.61

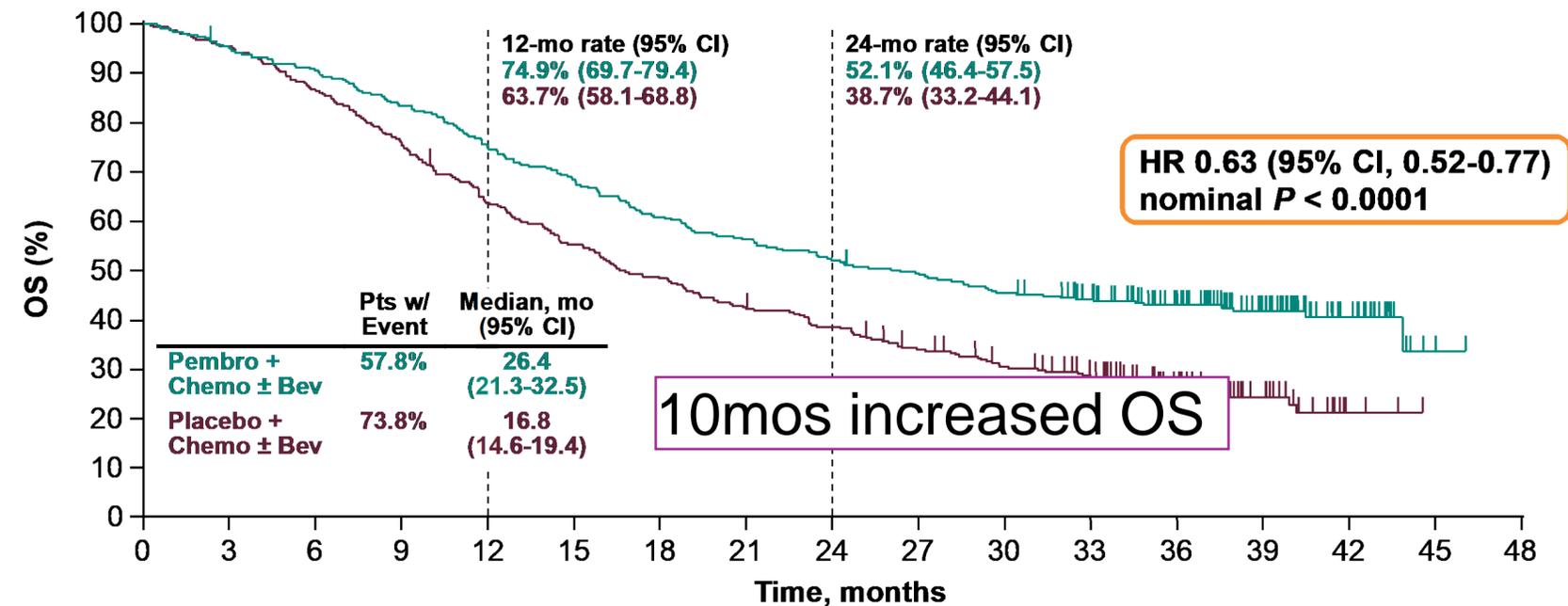
# KEYNOTE-826 Final OS

Median follow-up, 39.1 months

## Protocol-Specified Final OS: PD-L1 CPS $\geq 1$ Population



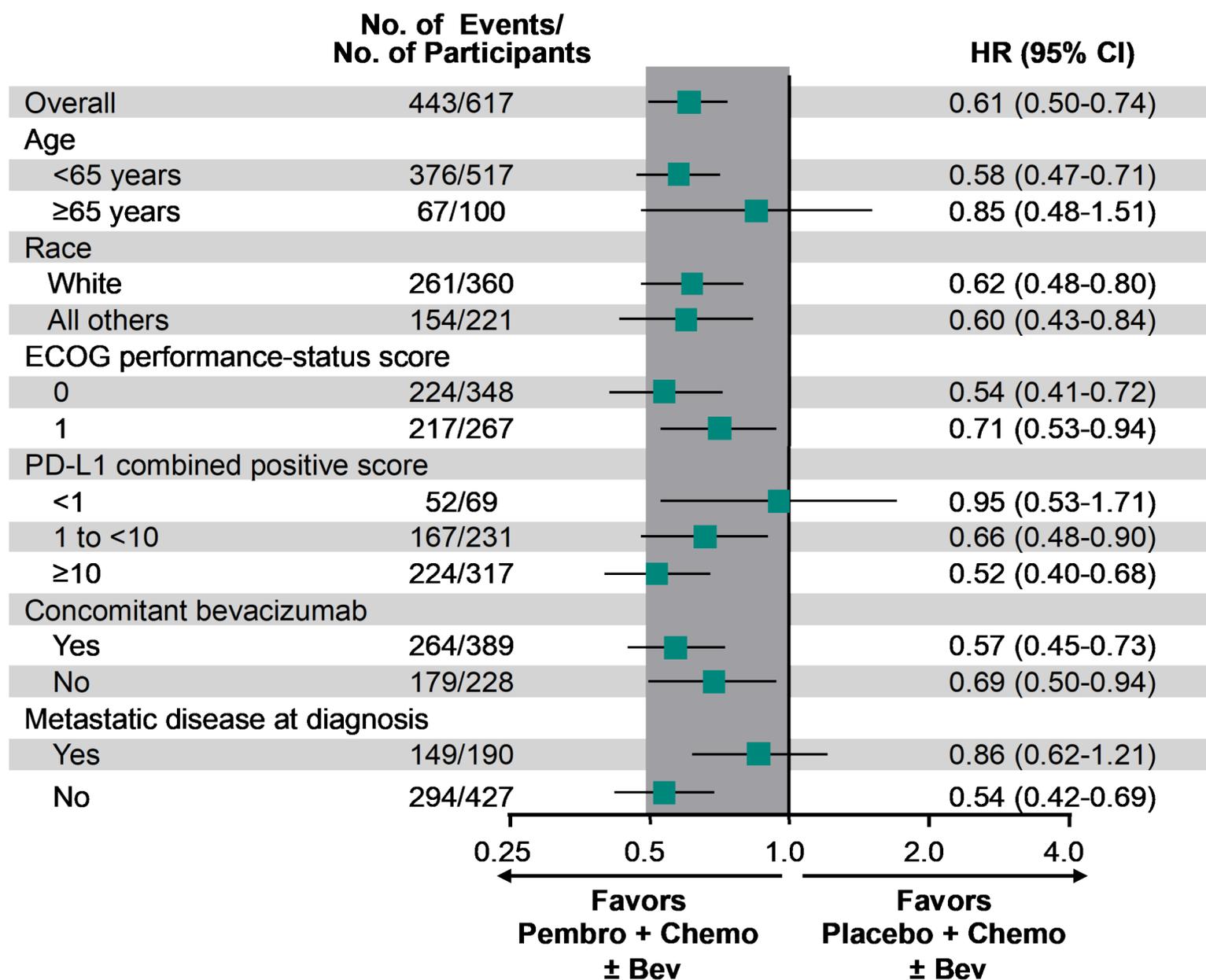
## Protocol-Specified Final OS: All-Comer Population



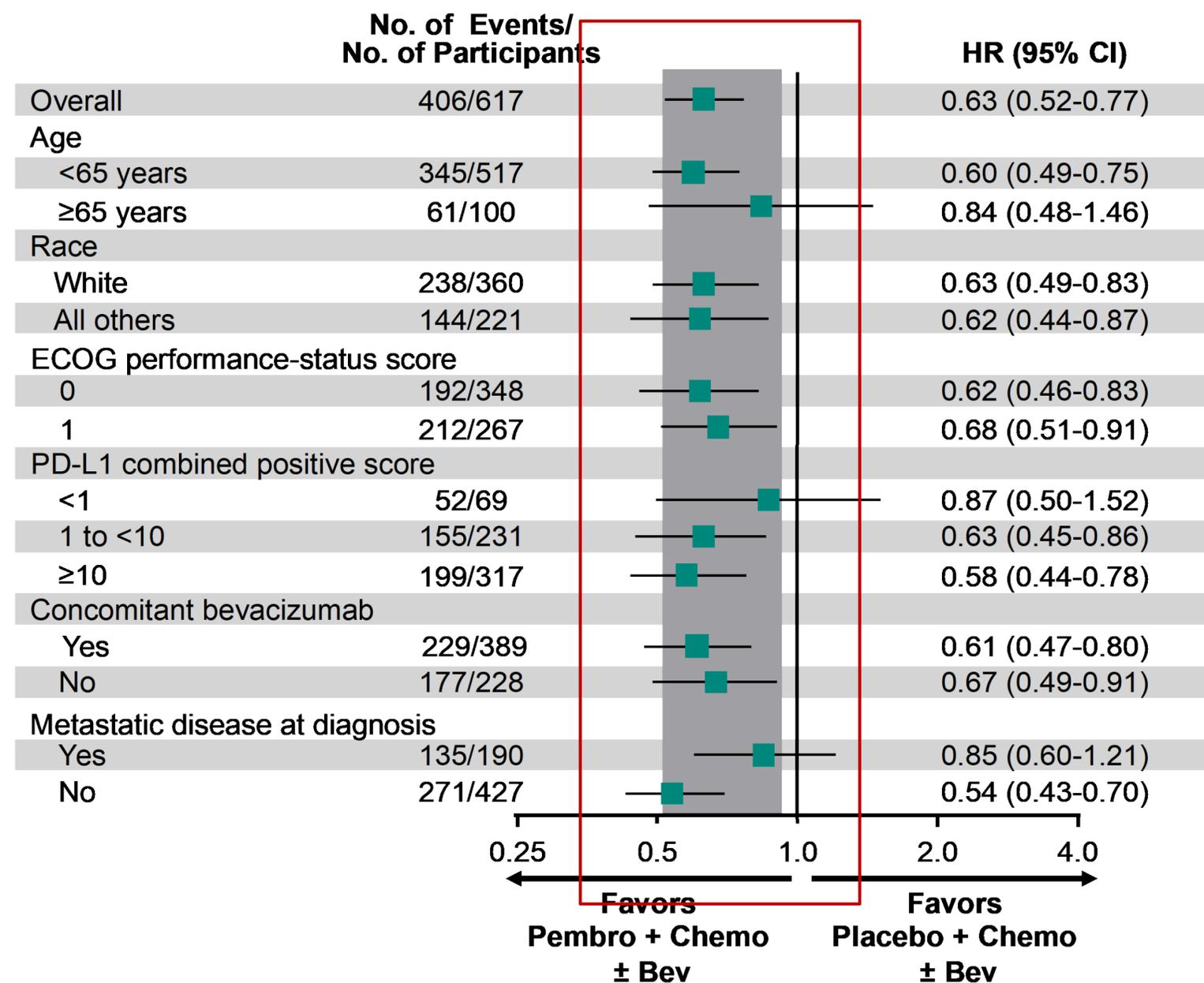
# KEYNOTE-826 Final analysis

Protocol-specified final analysis in subgroups

## PFS



## OS



# KEYNOTE-826 5-year follow-up

## Plenary: Cervical Cancer Oral Abstract Presentations

Session Type Plenary Session

Date Fri, 07.11.2025

Session Time 14:20 - 16:05

Room Hall A&B

Chair(s) [Hennie Botha \(South Africa\)](#), [Krishnansu S. S. Tewari \(United States of America\)](#)

Session Icon



**OP023 - PEMBROLIZUMAB PLUS CHEMOTHERAPY WITH OR WITHOUT BEVACIZUMAB IN PARTICIPANTS WITH PERSISTENT, RECURRENT, OR METASTATIC CERVICAL CANCER: 5-YEAR FOLLOW-UP RESULTS FROM KEYNOTE-826 (ID 426)**

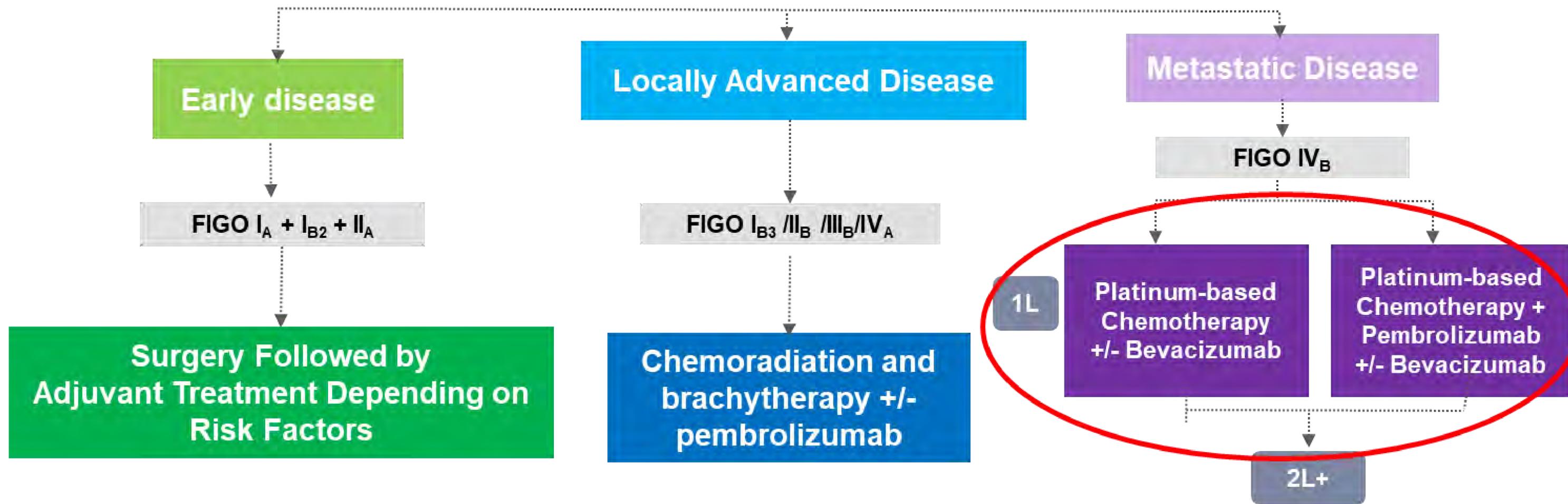
Presenter [Kosei Hasegawa \(Japan\)](#)

Lecture Time 15:19 - 15:27



# 1L Immunotherapy for CxCa

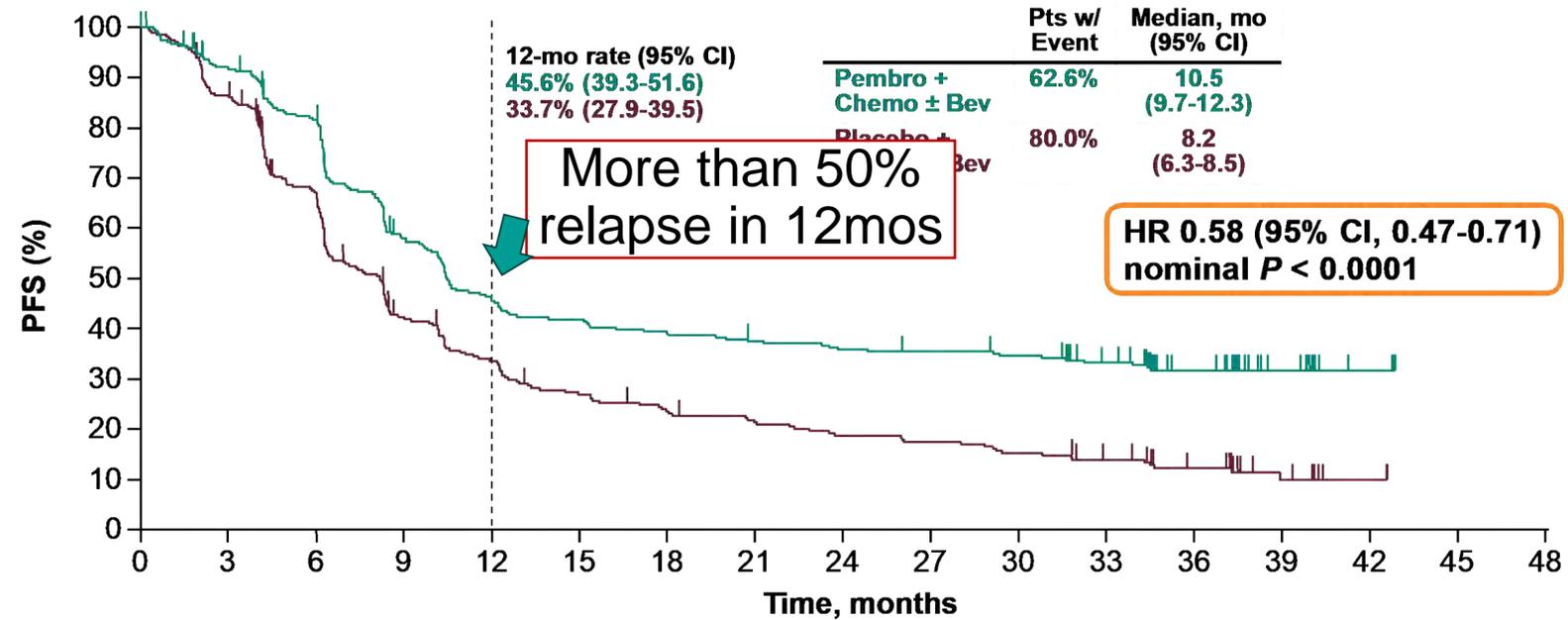
Trial (Immunotherapy)	N (Immuno/Control)	Primary Endpoint(s)	PFS (months, HR)	OS (months, HR)	Key Factors/Population
<b>KEYNOTE-826 (Pembrolizumab)</b>	308/309	PFS, OS (dual primary)	Median PFS: 10.4 vs 8.2; HR 0.65	Median OS: 24.4 vs 16.5; HR 0.67	Chemo ± bevacizumab; PD-L1+ and all-comers
<b>BEATcc (Atezolizumab)</b>	205/205	OS (primary), PFS	Median PFS: 13.0 vs 10.4; HR 0.73	Median OS: 32.1 vs 22.8; HR 0.68	Chemo + bevacizumab; all-comers
<b>COMPASSION-16 (Cadonilimab)</b>	222/223	PFS, OS (dual primary)	Median PFS: 10.7 vs 8.5; HR 0.67	Median OS: 23.9 vs 17.9; HR 0.67	Chemo ± bevacizumab; PD-L1-agnostic, China



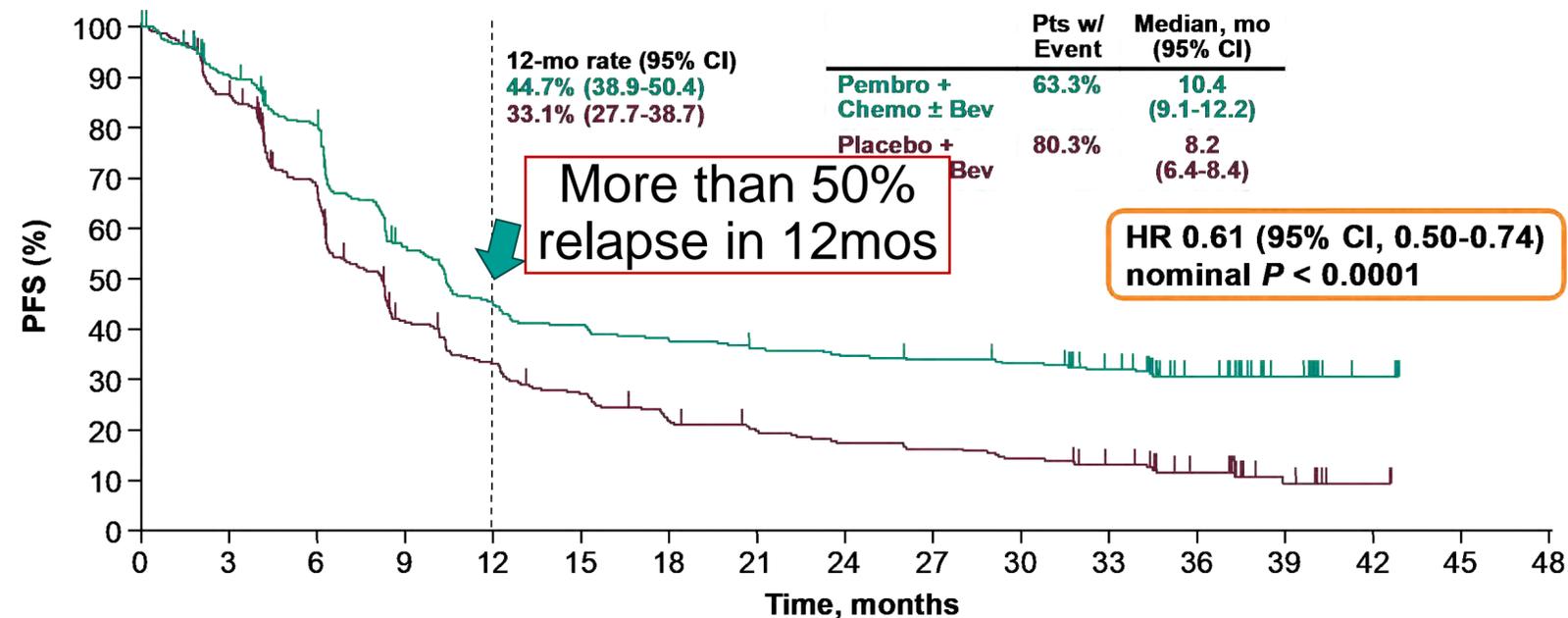
# KEYNOTE-826 Final PFS

Median follow-up, 39.1 months

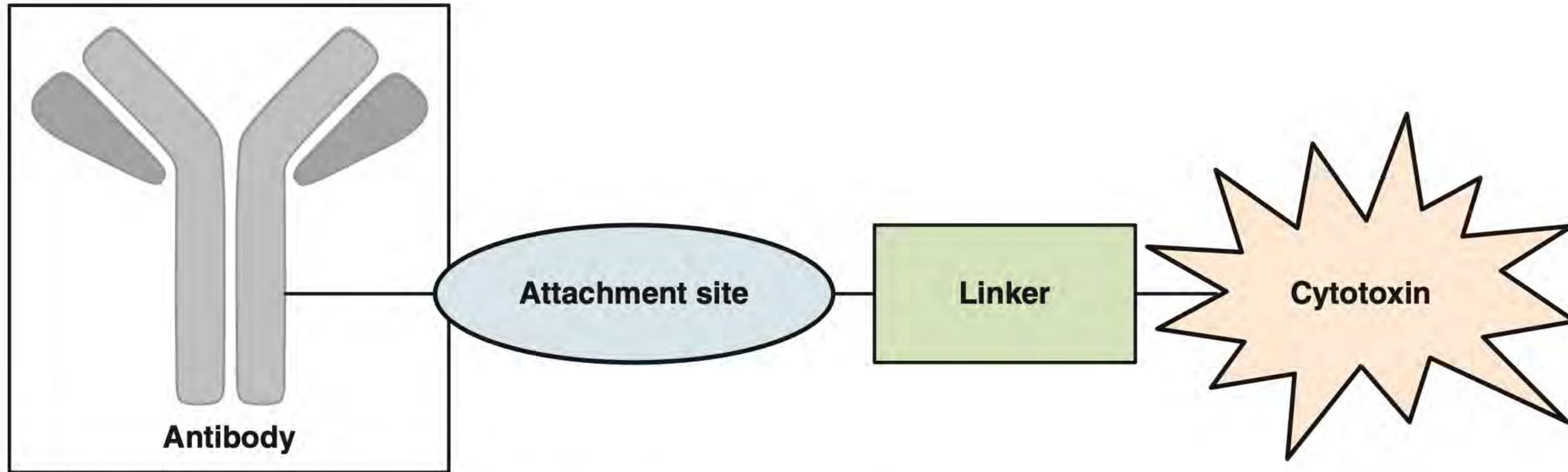
## Protocol-Specified Final PFS: PD-L1 CPS $\geq 1$ Population



## Protocol-Specified Final PFS: All-Comer Population



# Antibody Drug Conjugate □ ADC □



- Targets a well-characterized antigen with high tumor expression and limited normal tissue expression
- Maintains binding, stability, internalization, PK, etc. when conjugated to a cytotoxin
- Minimal nonspecific binding

- Typically through nonspecific modification of cysteine or lysine residues on the antibody
- Mixture of conjugates with variable drug:antibody ratios
- Site-selective conjugation technologies can produce more homogeneous ADCs

- Cleavable or noncleavable
- Stable in circulation
- Selective intracellular release of drug (e.g. via enzymatic cleavage or antibody degradation)

- Highly potent
- Non-immunogenic
- Amenable to modifications for linker attachment
- Defined mechanism of action



# The Role of Antibody Drug Conjugates (ADCs) in Cervical Cancer

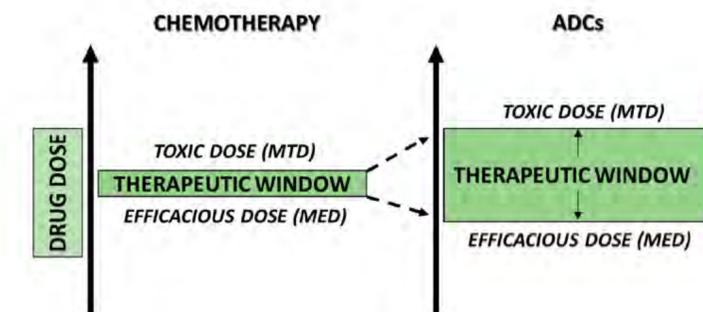
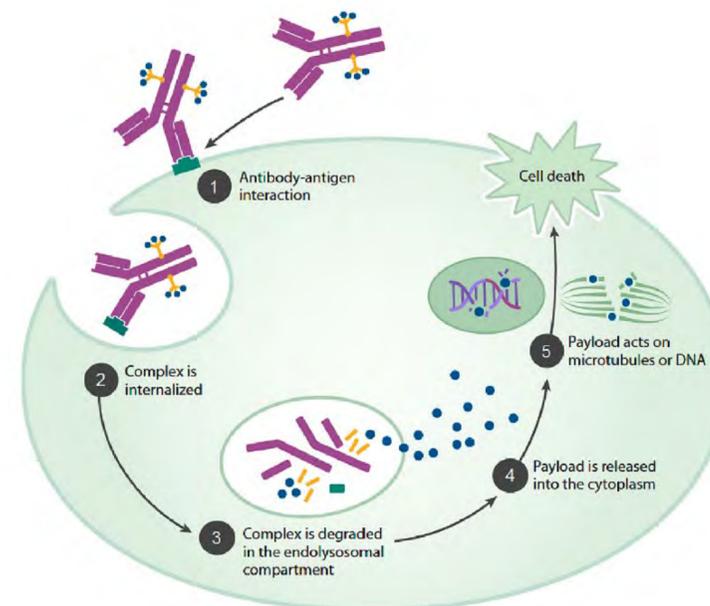
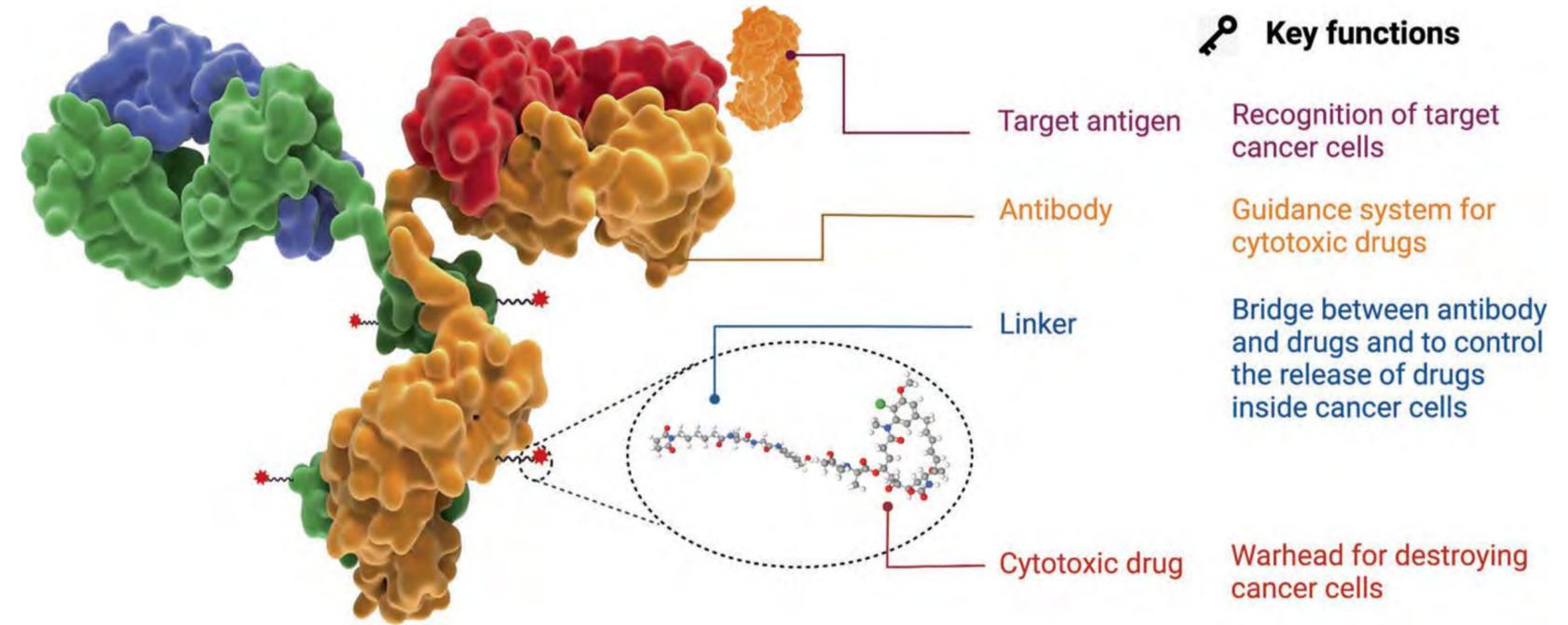
**Ritu Salani, MD, MBA**

UCLA Health

Los Angeles, California, USA



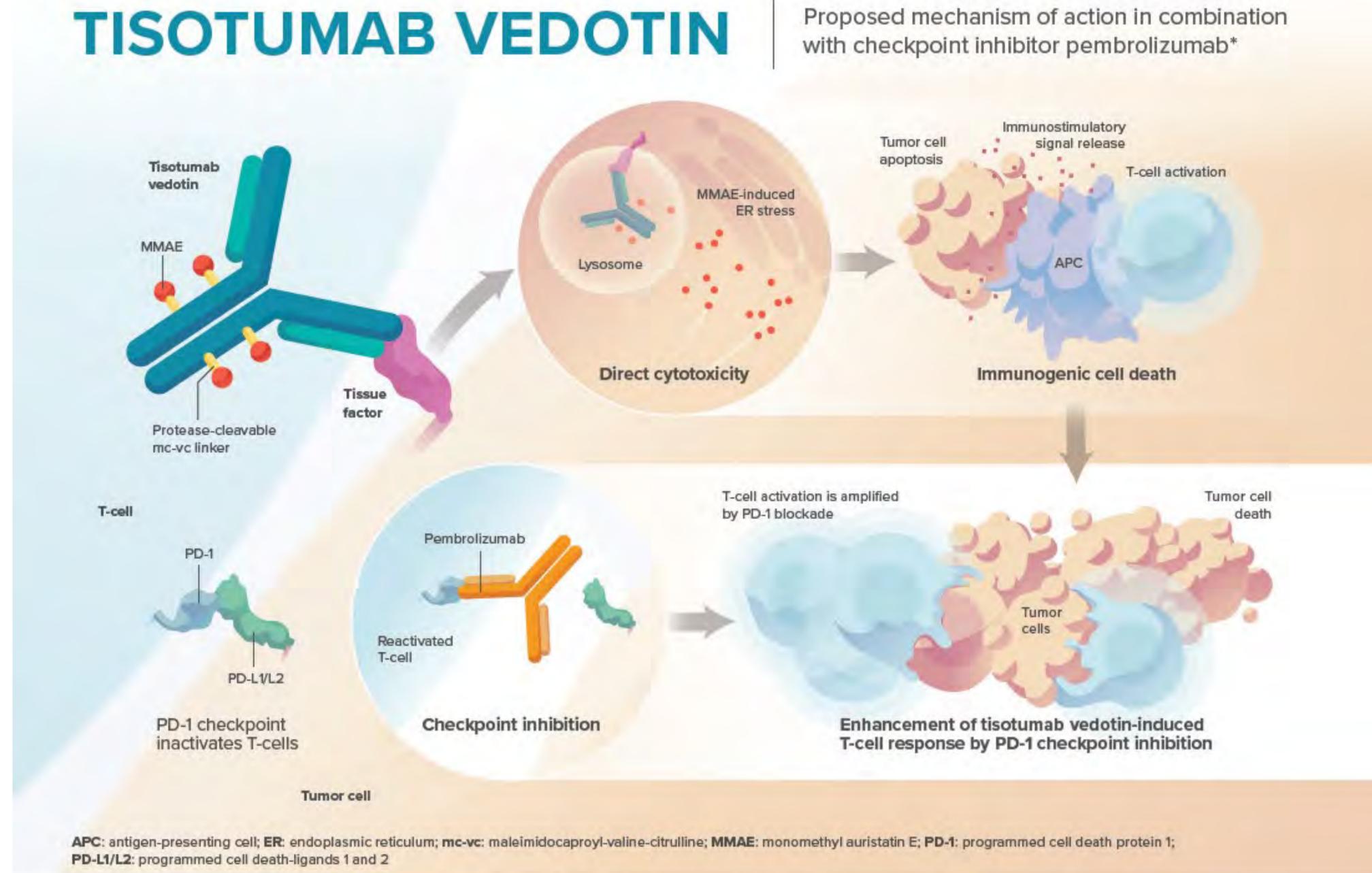
# Antibody Drug Conjugate



# Tisotumab Vedotin (TV)

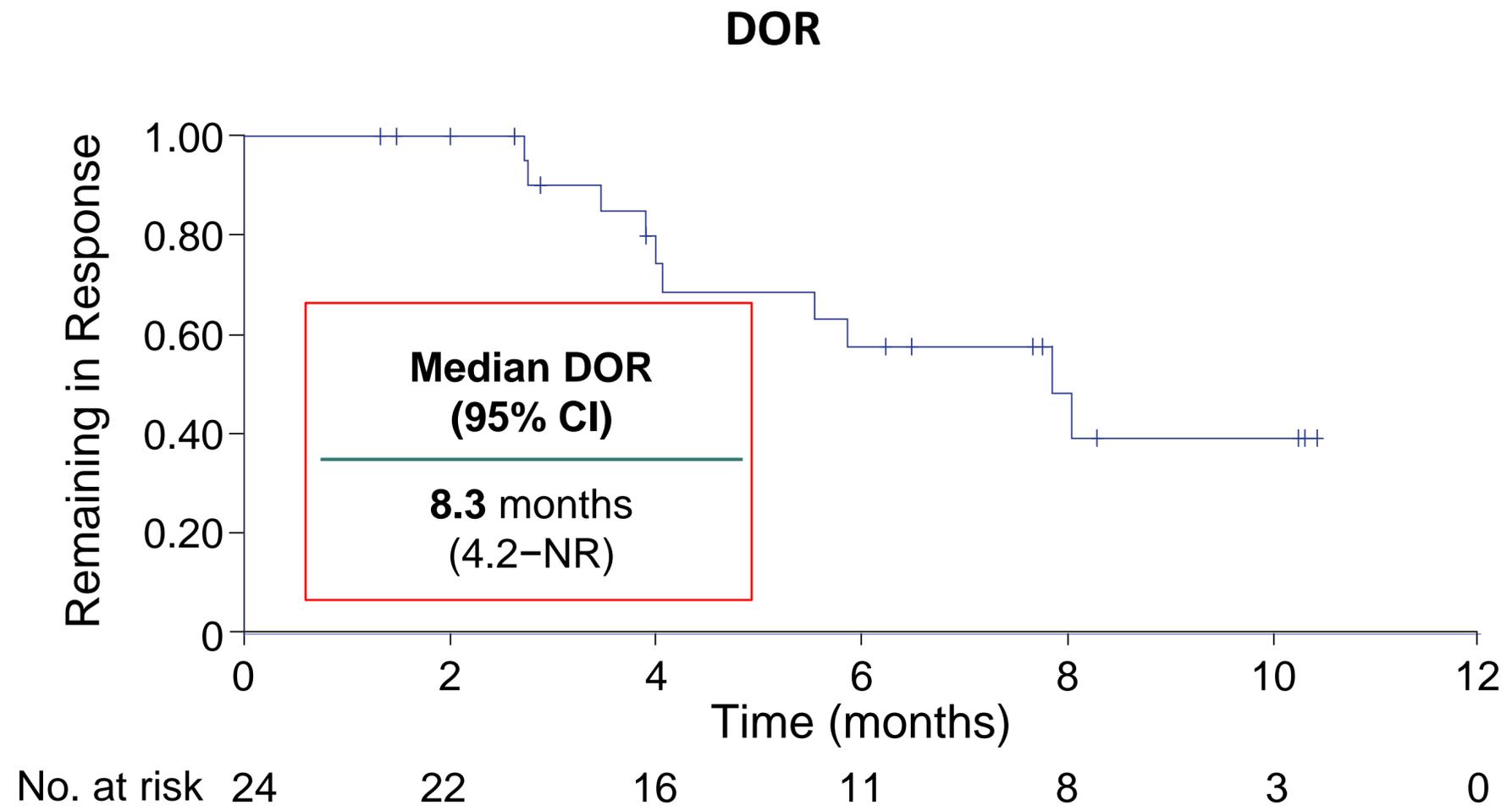
## Tissue factor (TF) ADC

- Microtubule-disrupting agent  
**MMAE**
- Highly expressed in cervix cancer
- Involved in progression and metastases



# Phase 2: innovaTV 204

	<b>N = 101</b>
<b>Confirmed ORR (95% CI),<sup>a</sup> %</b>	<b>24 (15.9–33.3)</b>
CR, n (%)	7 (7)
PR, n (%)	17 (17)
SD, n (%)	49 (49)
PD, n (%)	24 (24)
Not evaluable, n (%)	4 (4)



**Clinically meaningful and durable responses were observed**  
**Accelerated approval**

# InnovaTV 301/ENGOT-cx12/GOG-3057

## Key eligibility criteria

- Recurrent or metastatic cervical cancer
- Disease progression on or after chemotherapy doublet ± Bev and an anti-PD-(L)1 agent, if eligible and available
- ≤2 prior therapies for recurrent/metastatic disease
- ECOG PS 0-1

**N=502**

**R  
1:1**

**Primary endpoint: OS**

**Key secondary endpoints: PFS, ORR, safety**

**Stratification:**

- ECOG PS
- Prior Bev
- Prior anti-PD(L)1 therapy
- Region

**Tisotumab vedotin  
(n=253)  
2.0 mg/kg IV q3w**

**Chemotherapy  
(n=249)  
Topotecan, vinorelbine,  
gemcitabine, irinotecan, or  
pemetrexed  
(investigator's choice)**

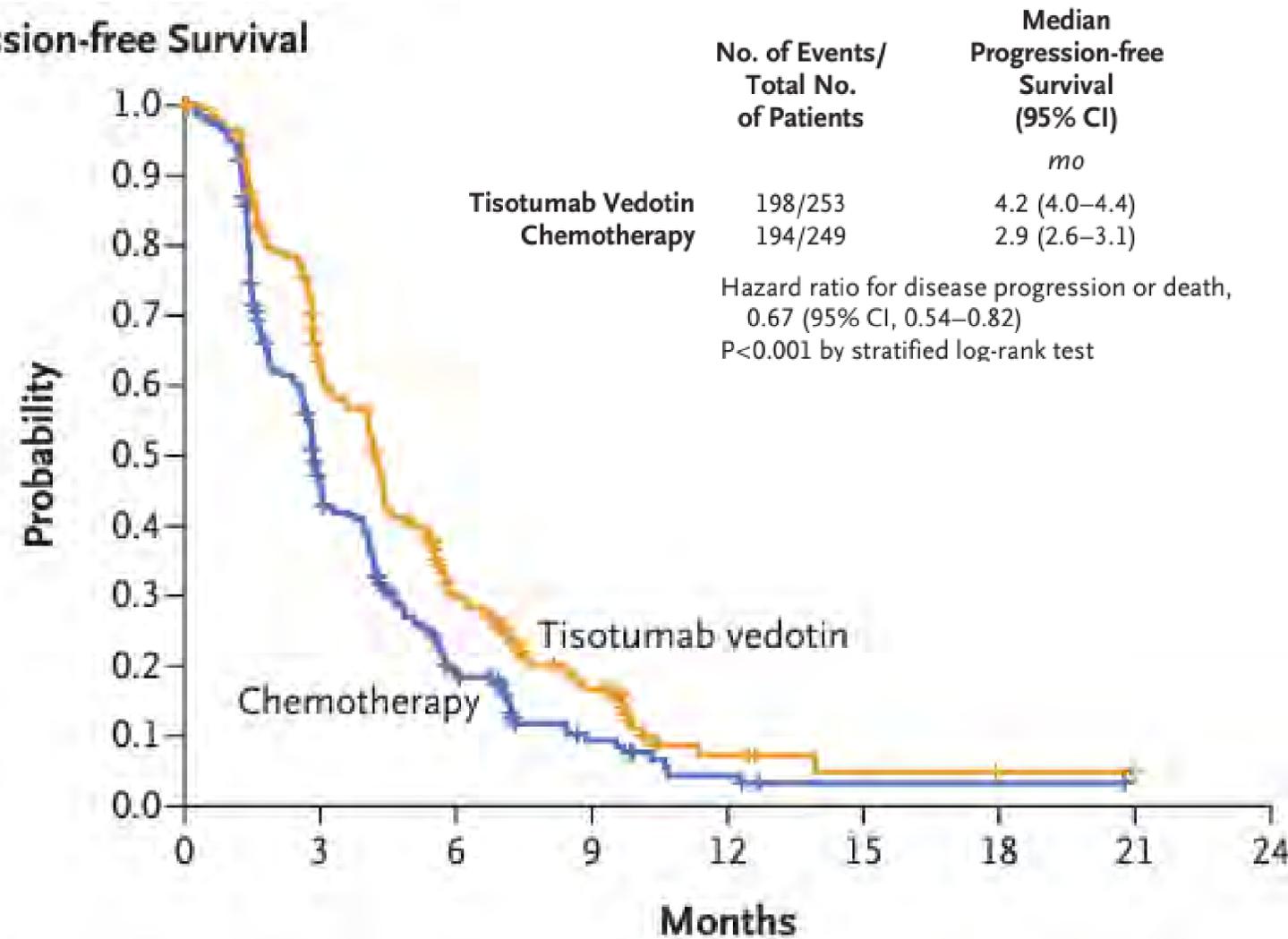
	<b>TV (N=253)</b>	<b>Chemo (N=249)</b>
<b>Age, years, median (range)</b>	51 (26-80)	50 (27-78)
<b>Baseline ECOG 0</b>	137 (54.2)	136 (54.6)
<b>Histology, n (%)</b>		
Squamous cell carcinoma	160 (63.2)	157 (63.1)
Adenocarcinoma	85 (33.6)	75 (30.1)
Adenosquamous	8 (3.2)	17 (6.8)
<b>Prior regimens, n (%)</b>		
1	159 (62.8)	149 (59.8)
2	93 (36.8)	100 (40.2)
<b>Prior bevacizumab, n (%)</b>	164 (64.8)	157 (63.1)
<b>Prior anti-PD-(L)1 therapy, n (%)</b>	71 (28.1)	67 (26.9)
<b>Prior radiation therapy for cervical cancer, n (%)</b>	205 (81.0)	203 (81.5)

# InnovaTV 301: Survival Outcomes



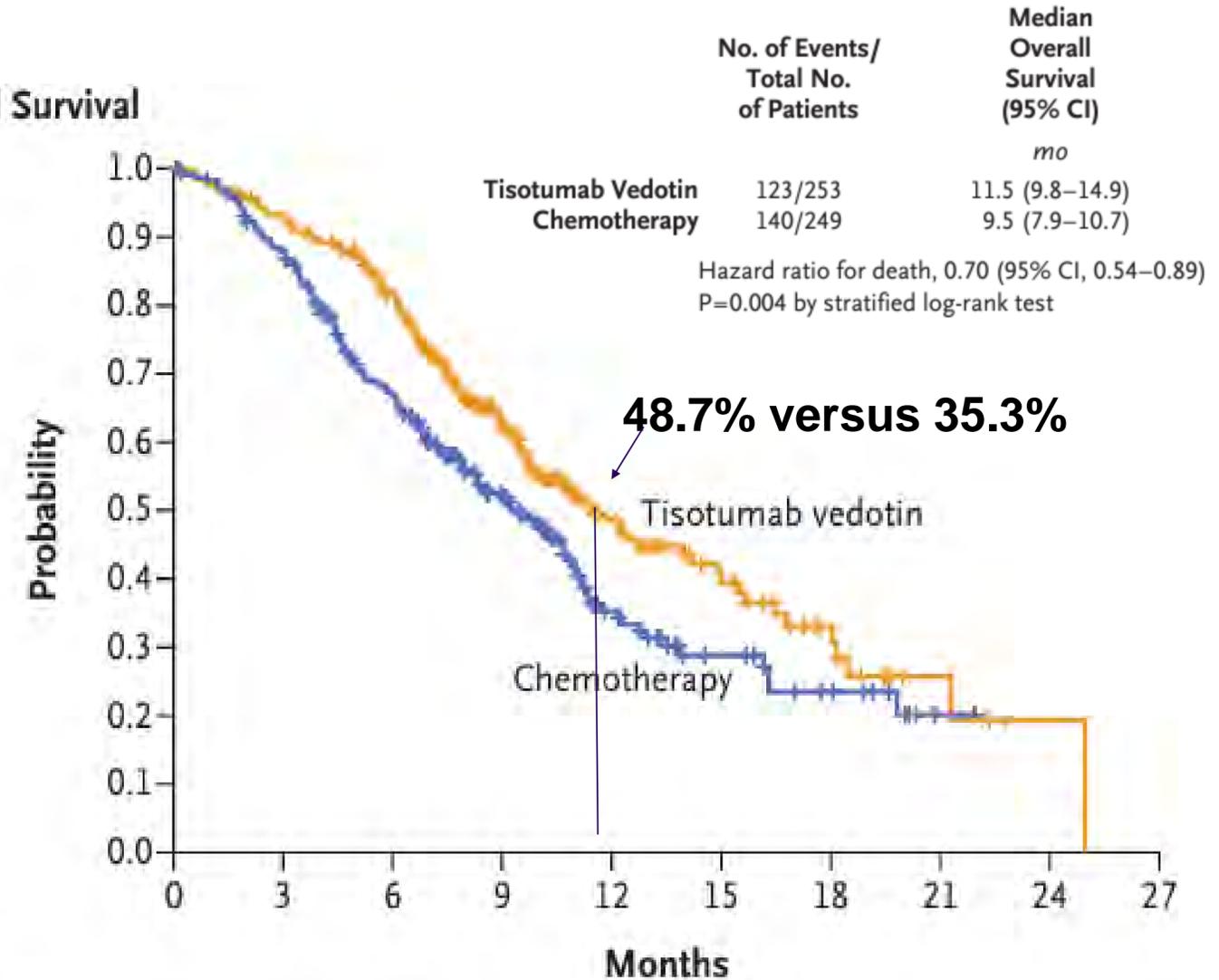
Tisotumab Vedotin as Second- or Third-Line Therap...  
www.nejm.org

**Progression-free Survival**



No. at Risk		0	3	6	9	12	15	18	21	24
Tisotumab vedotin	253	148	62	25	5	2	1	0	0	0
Chemotherapy	249	96	34	11	4	1	1	0	0	0

**Overall Survival**

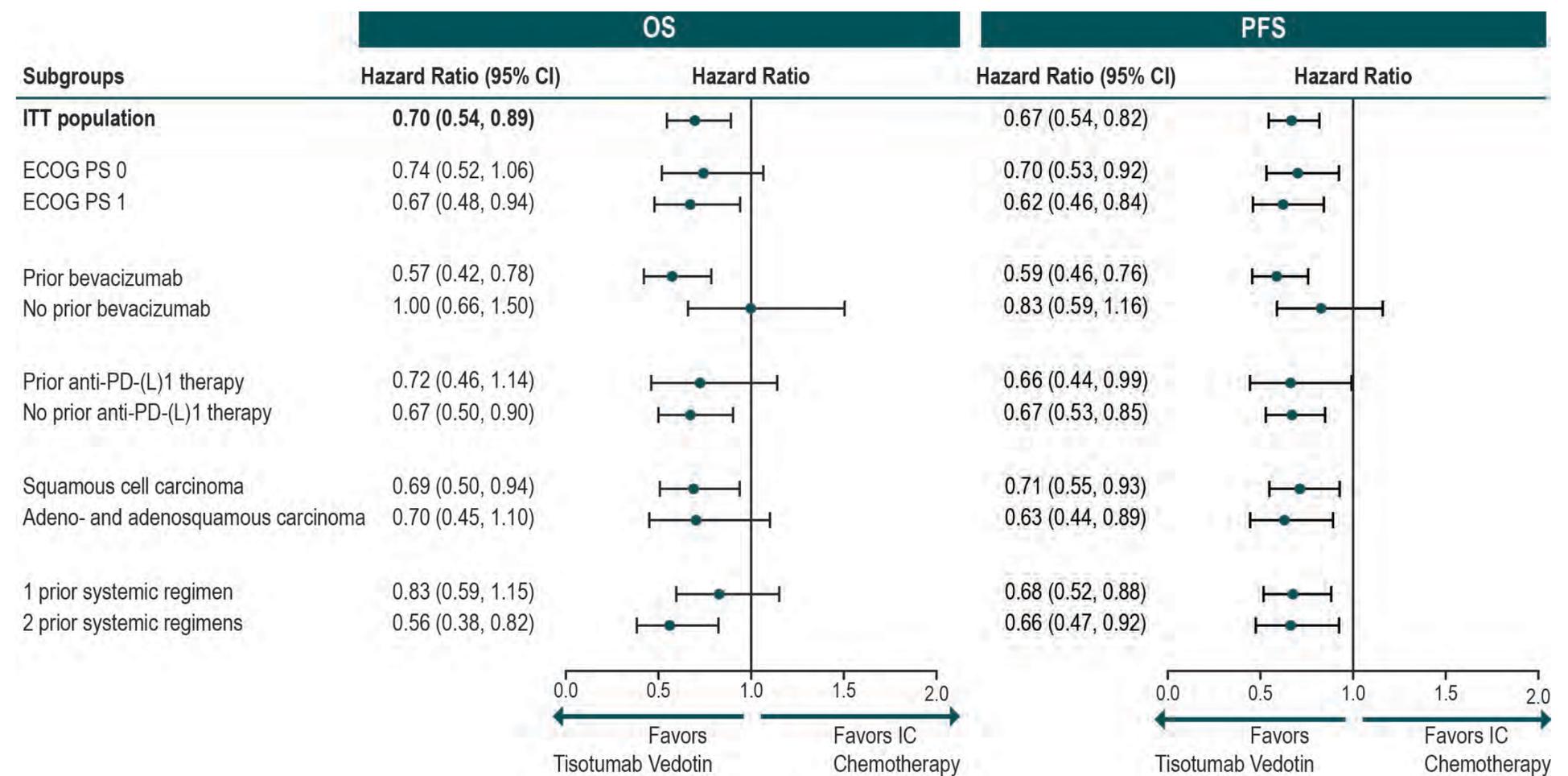


No. at Risk		0	3	6	9	12	15	18	21	24	27
Tisotumab vedotin	253	234	191	109	52	29	14	4	1	0	0
Chemotherapy	249	212	150	87	37	19	11	1	0	0	0

# InnovaTV 301: Outcomes

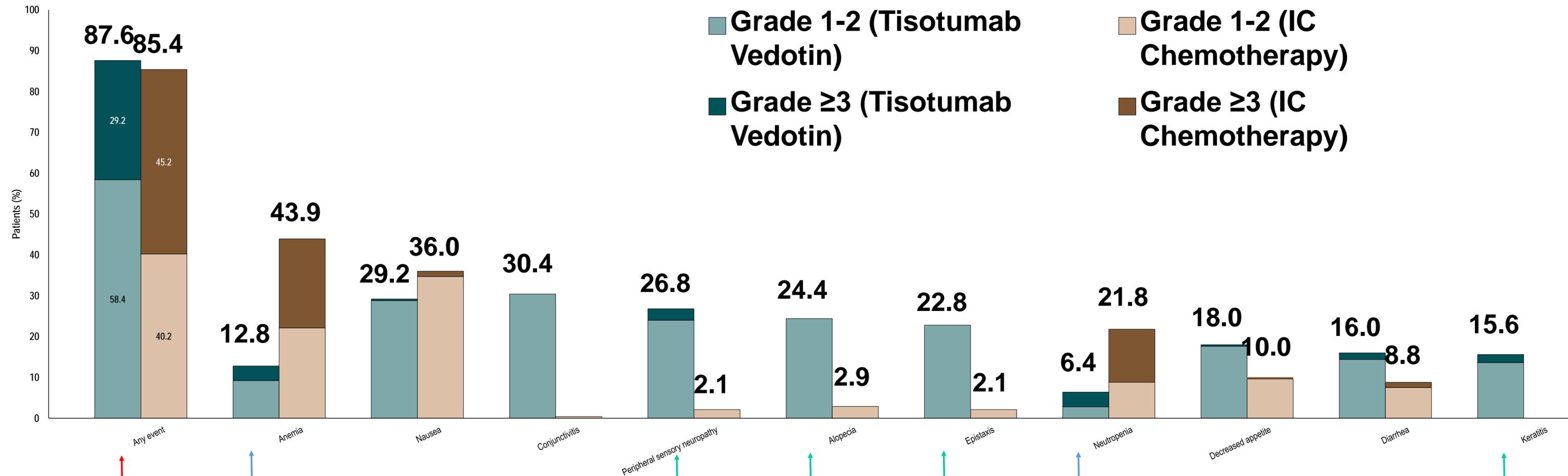
## Subgroup Analysis

	TV (N=253)	Chemo (N=249)
<b>ORR, % (95% CI)</b>	<b>17.8 (13.3-23.1)</b>	<b>5.2 (2.8-8.8)</b>
Odds ratio (95% CI) P value	4.0 (2.1-7.6) p<0.0001	
<b>Best Overall Response, n (%)</b>		
CR	6 (2.4)	0
PR	39 (15.4)	13 (5.2)
SD	147 (58.1)	132 (53.0)
PD	46 (18.2)	74 (29.7)
Not evaluable/Not available	15 (5.9)	30 (12.0)
<b>DCR, % (95% CI)</b>	<b>75.9 (70.1-81.0)</b>	<b>58.2 (51.8-64.4)</b>
Median DOR (95% CI)	5.3 (4.2-8.3)	5.7 (2.8-NR)



OS and PFS benefit was generally consistent across key subgroups

# Most Common Treatment-Related Adverse Events<sup>a</sup>



- Grade 5 TRAEs occurred in 2 (0.8%) and 1 (0.4%) patients in the tisotumab vedotin and IC chemotherapy arms, respectively<sup>b</sup>
- Median relative dose intensity was 96.1% and 90.0% in the tisotumab vedotin and IC chemotherapy arms, respectively

<sup>a</sup>TRAEs listed are those occurring in ≥15% of patients on either arm; <sup>b</sup>Grade 5 TRAEs included acute kidney injury (n=1) and Stevens-Johnson syndrome (n=1) in the tisotumab vedotin arm and pancytopenia (n=1) in the IC chemotherapy arm.

# Ocular Toxicity

- Causes changes in corneal epithelium and conjunctiva
- May result
  - Vision changes/loss
  - Keratitis
  - Conjunctivitis
  - Corneal ulceration



- Conduct ophthalmic examination at baseline, prior to each dose, and as clinically indicated
- Eye care plan including
  - Vasoconstrictor drops
  - Steroid eye drops
  - Lubricating drops
  - Ice packs
- Withhold until improvement and resume, reduce the dose, or permanently discontinue, based on severity

# GOG-3116/C5721005

PI: Scott Jordan, MD

Single-arm, prospective, low-interventional study of tisetumab vedotin in adult participants in the US with r/mCC who have received prior systemic therapy for recurrent or metastatic disease

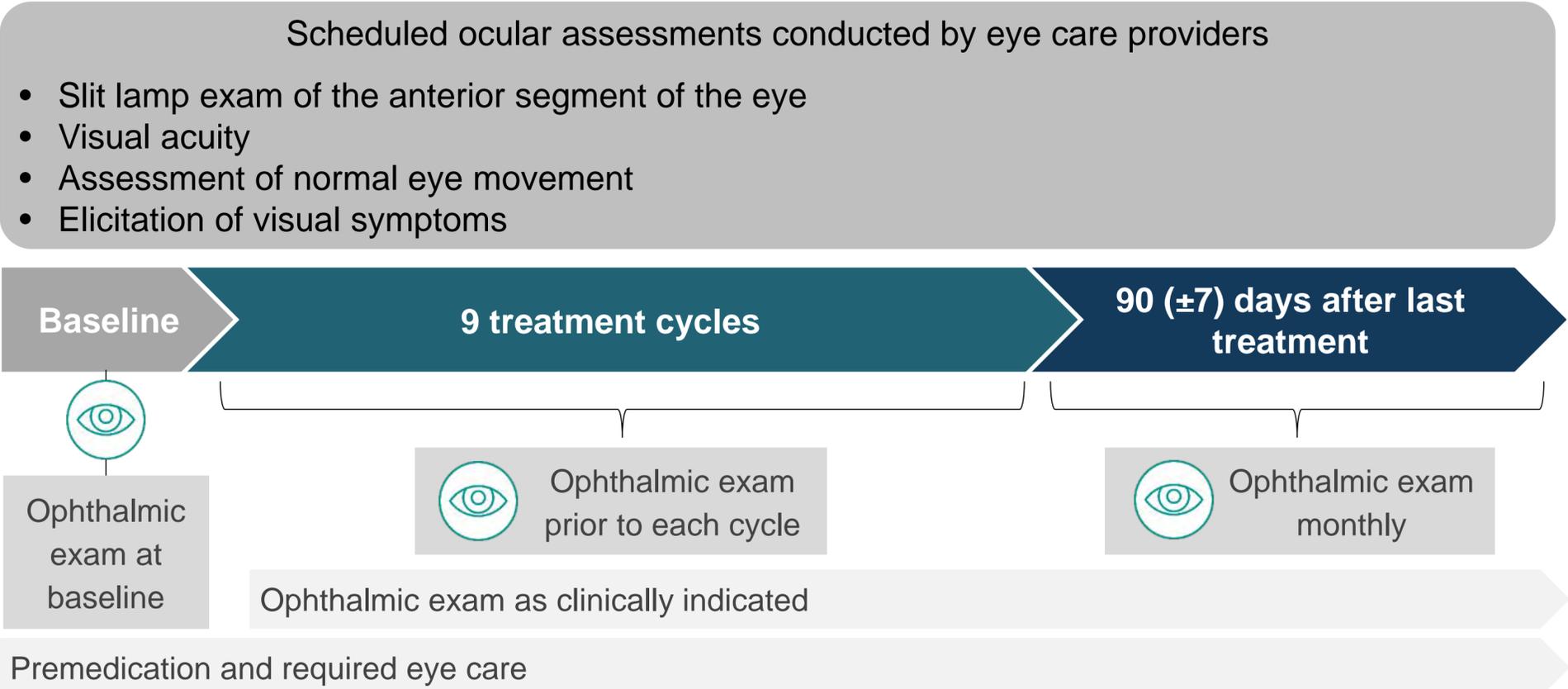
## Patient Population

r/m CC with disease progression on or after chemotherapy, following prior systemic therapy  
No active ocular disease at baseline  
No prior TV treatment

**Tisetumab vedotin**  
2.0 mg/kg IV on Day 1 of a 21-day cycle (Q3W)

**N=100**

## Intervention



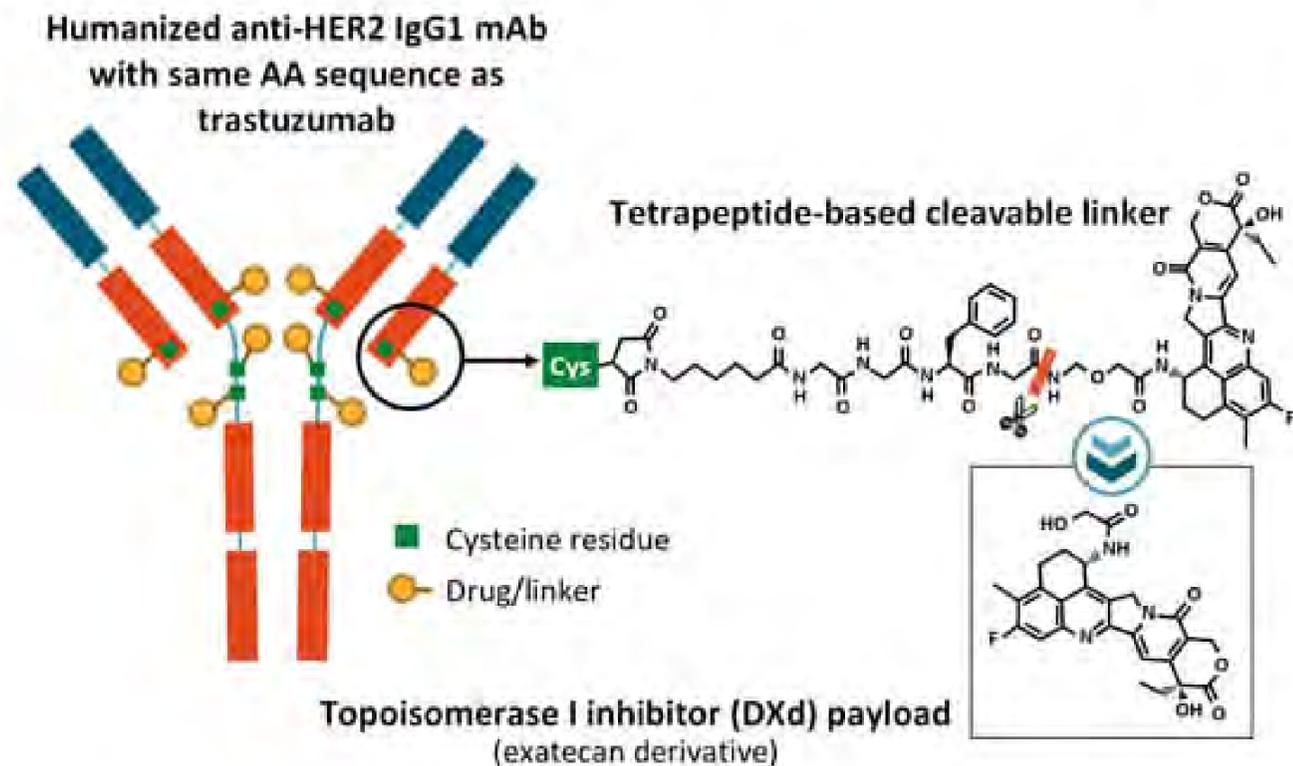
## Endpoints

- Primary endpoint**
- Type, incidence and severity of ocular AEs
- Secondary endpoints**
- Time to onset, time to resolution, and outcome of ocular AEs
  - Incidence of serious AEs
  - AEs leading to dose modifications including treatment discontinuation

**Objective:** to further characterize the incidence and severity of tisetumab vedotin-related ocular events with prospectively pre-specified, scheduled ocular assessments in patients receiving tisetumab vedotin for r/mCC

# Emerging ADC Options

## HER2-Targeted ADC: Trastuzumab Deruxtecan



- High drug:antibody ratio: ~8
- Stable linker-payload
- Tumor-selectable cleavable linker
- High potency, membrane-permeable payload with short systemic half-life
- Bystander killing effect

### HER2 IHC 3+ and 2+ prevalence

Site	Icon	IHC 3+ Prevalence	IHC 2+ Prevalence
Endometrial		IHC 3+ 6–17% <sup>5,8</sup>	IHC 2+ 13–39% <sup>5,8</sup>
Cervical		IHC 3+ 4–11% <sup>1,9</sup>	IHC 2+ 18% <sup>9</sup>
Ovarian		IHC 3+ 2–5% <sup>1,10</sup>	IHC 2+ 8–18% <sup>10,11</sup>

# DESTINY-PanTumor02: Study Design

A phase 2, multicenter, open-label study to evaluate the efficacy and safety of trastuzumab deruxtecan for the treatment of selected HER2 expressing tumors<sup>1,2</sup>

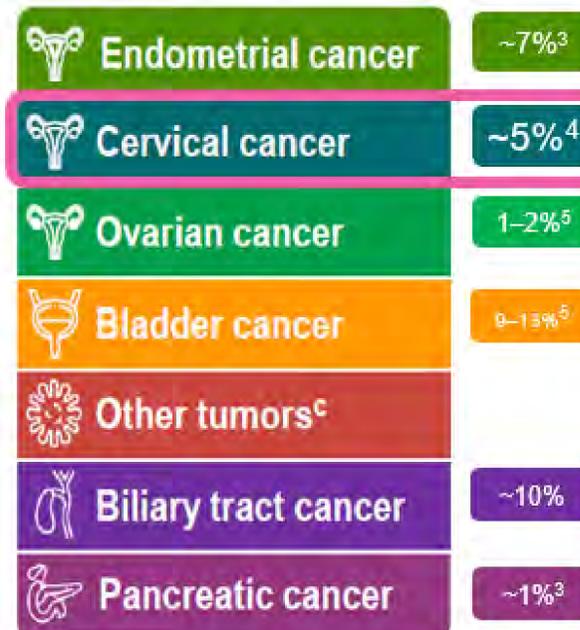
## Key Eligibility Criteria

- Advanced solid tumors not eligible for curative therapy
- 2L+ patient population
- HER2 expression (IHC 3+ or 2+)
- Local test or central test by HercapTest if local test not feasible (ASCO/CAP gastric cancer scoring)
- Prior HER-targeting therapy allowed
- ECOG/WHO PS ≤1

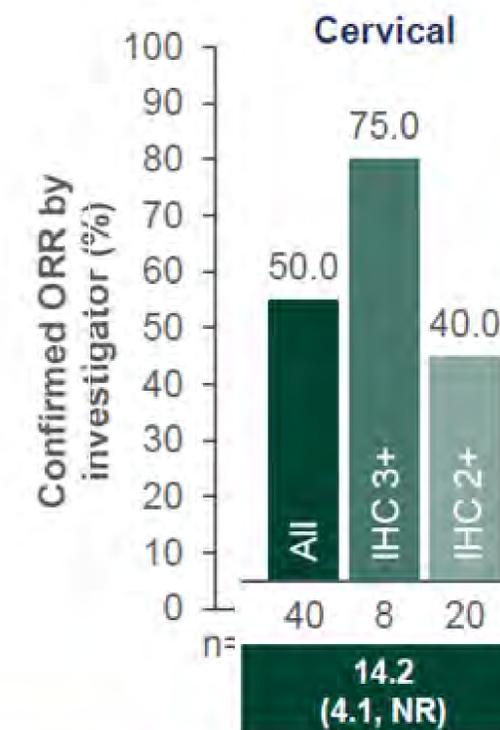
T-DXd  
5.4 mg/kg Q3W

40 per cohort<sup>b</sup>

Frequency of HER2 mutations across tumor types



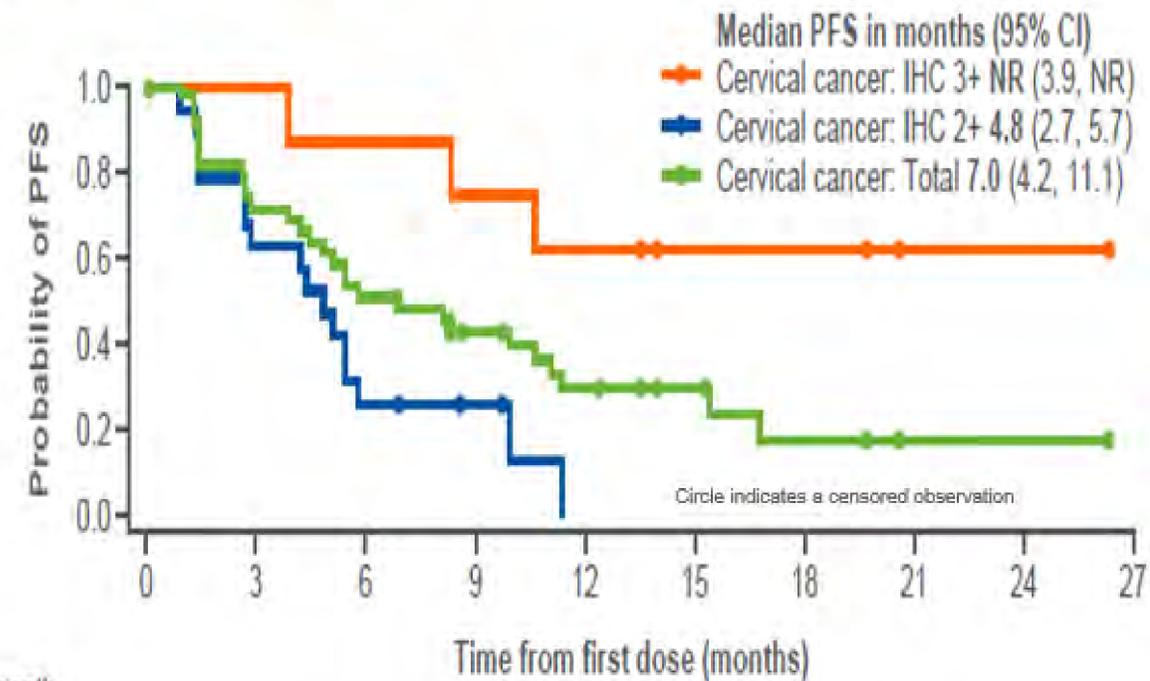
## ORR by HER2 status in Cervical Cancer



All patients were HER2-positive per local determination

# DESTINY-PanTumor02:

## PFS by HER2 status in Cervical Cancer



Number at risk, month	0	3	6	9	12	15	18	21	24	27
Cervical cancer: IHC 3+	8	8	7	6	5	3	3	1	1	0
Cervical cancer: IHC 2+	20	12	5	3	0					
Cervical cancer: Total	40	28	20	14	9	6	3	1	1	0

TEAEs	All patients (N=267); n (%)
Any drug-related TEAEs	226 (84.6)
Drug-related TEAEs Grade ≥3	109 (40.8)
Serious drug-related TEAEs	36 (13.5)
Drug-related TEAEs associated with dose discontinuations	23 (8.6)
Drug-related TEAEs associated with dose interruptions	54 (20.2)
Drug-related TEAEs associated with dose reductions	54 (20.2)
Drug-related TEAEs associated with deaths	4 (1.5) <sup>a</sup>

Most common TEAEs	Any Grade	Grade ≥3
Nausea	55.1	3.7
Fatigue	40.1	7.1
Neutropenia	32.6	19.1
Anemia	27.7	10.9
Diarrhea	25.8	3.7
Vomiting	24.7	1.5
Decreased appetite	17.6	1.5
Thrombocytopenia	17.2	5.6
Alopecia	16.9	
Increased transaminases	10.1	0.4
Leukopenia	10.1	2.6

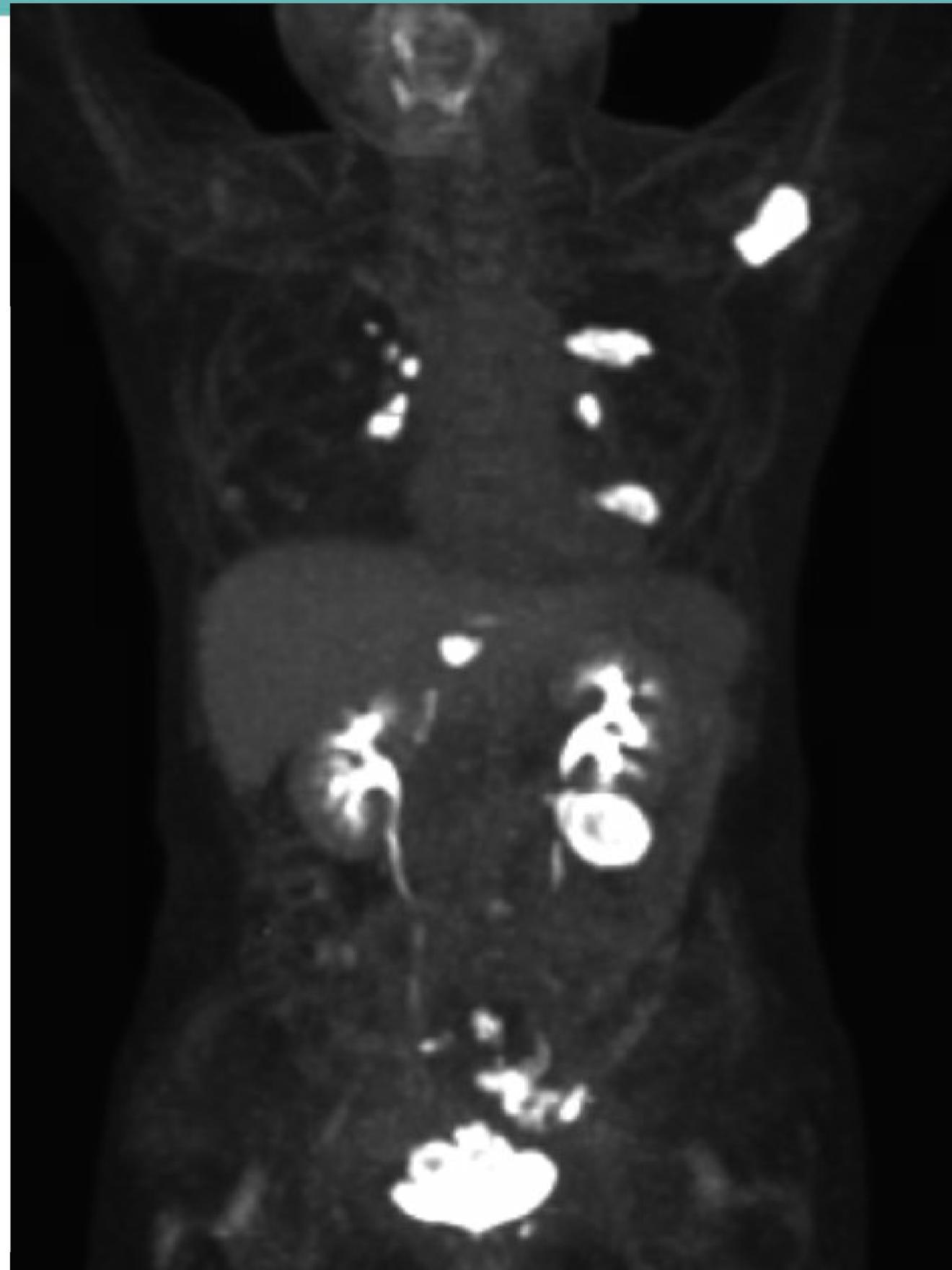
ILD/pneumonitis adjudicated as T-DXd related, n (%)	Grade 1	Grade 2	Grade 3	Grade 4	Grade 5	Any grade
All patients (N=267)	7 (2.6)	17 (6.4)	1 (0.4)	0	3 (1.1)	28 (10.5)

**FDA accelerated approval: patients with unresectable or metastatic HER2+ (IHC 3+) solid tumors who have received prior systemic treatment and have no satisfactory alternative treatment options**

# Clinical Case

## Clinical History

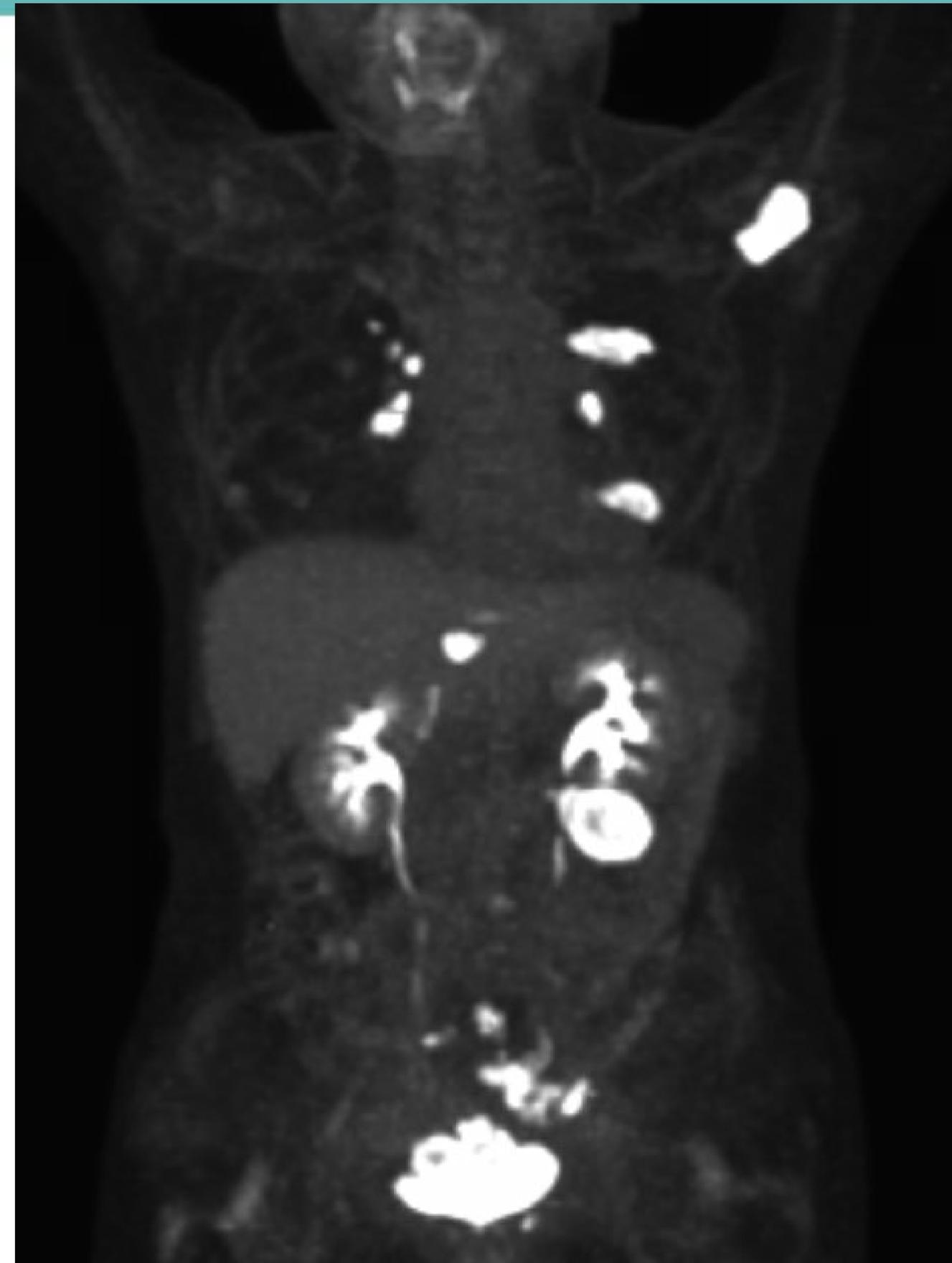
- 36 year from China who presented with advanced cervical cancer (non-vaccinated)
- **Symptom:** pelvic pain /vaginal bleeding
- **Pelvic MRI with Contrast:** 3.1 x 1.5 cm mass with ill-defined border to left bladder wall/enlarged lymph nodes adjacent to bilateral iliac vessels, largest one about 1.8 x 1.3 cm in size.
- **Cystoscopy** with space-occupying lesion in bladder
- **Biopsy:** Squamous cell carcinoma, poorly differentiated, non-keratinizing type, involving colonic submucosa. **PD-L1 positive** , **Her 2 neu negative (0)**
- **PET Scan:** FDG avid cancer involving lower segment of uterus and bladder, multiple lymph node metastases, abdominal, and pelvic metastases



# Clinical Case

## Treated Locally

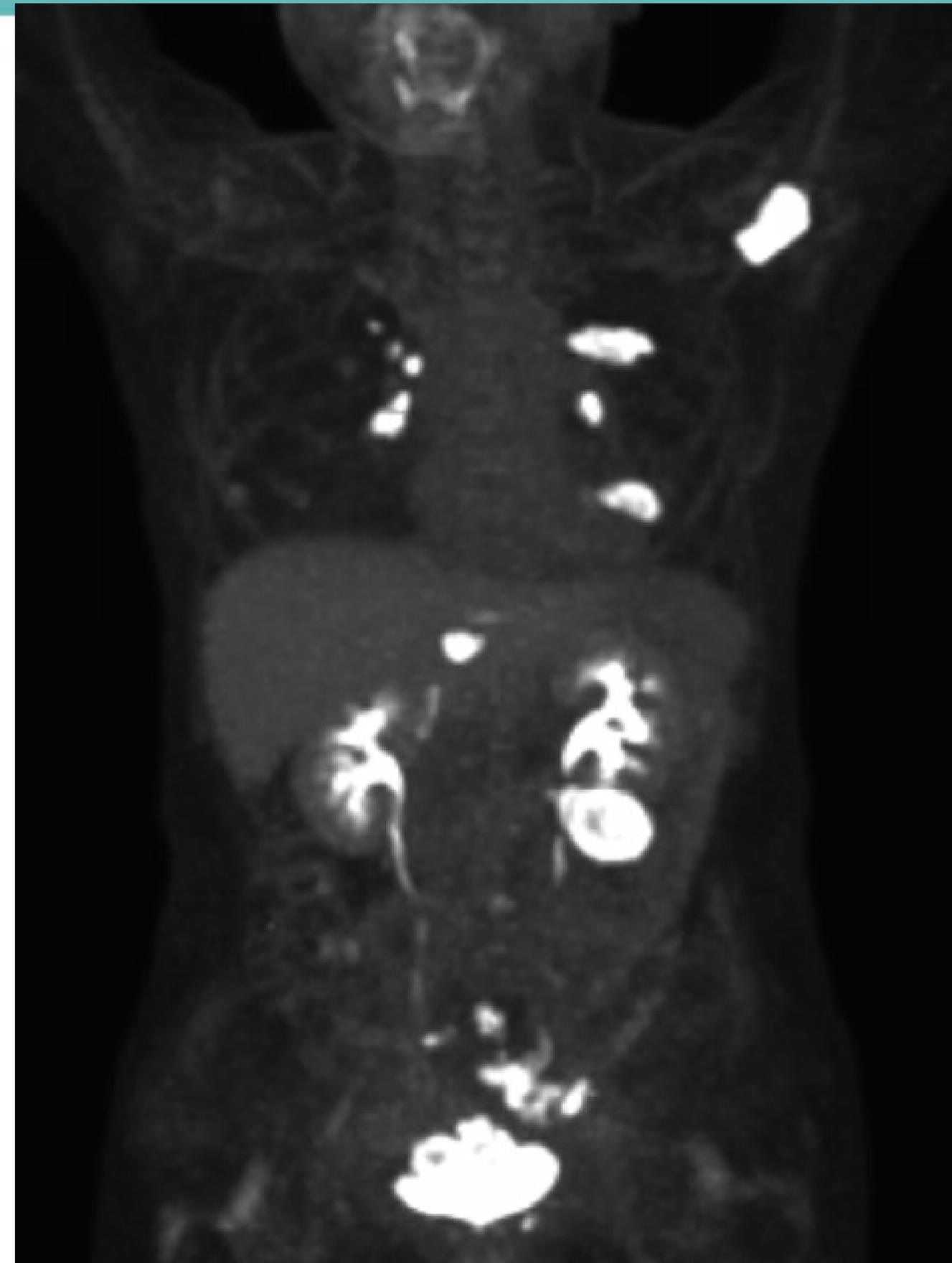
- **Carboplatin and paclitaxel**
- **Radiation Therapy** to the cervical lesion and pelvic lymph nodes
- **Post Therapy Imaging:** Cervical mass involving left bladder wall and lymph nodes significantly regressed compared with previous scan. Multiple heterogeneous masses in peritoneum of left upper abdomen.
- Treatment: Paclitaxel + Carboplatin + Pembrolizumab x 6 cycles
  - After completion of therapy, PET CT done
  - Multifocal areas of FDG avid metastatic disease involving bilateral hilar and right internal mammary lymph nodes, multifocal soft tissue implants in the abdomen and pelvis, and multiple osseous metastases.



# Clinical Case

**Q: Outside of clinical trials, which treatment would you consider next for this patient?**

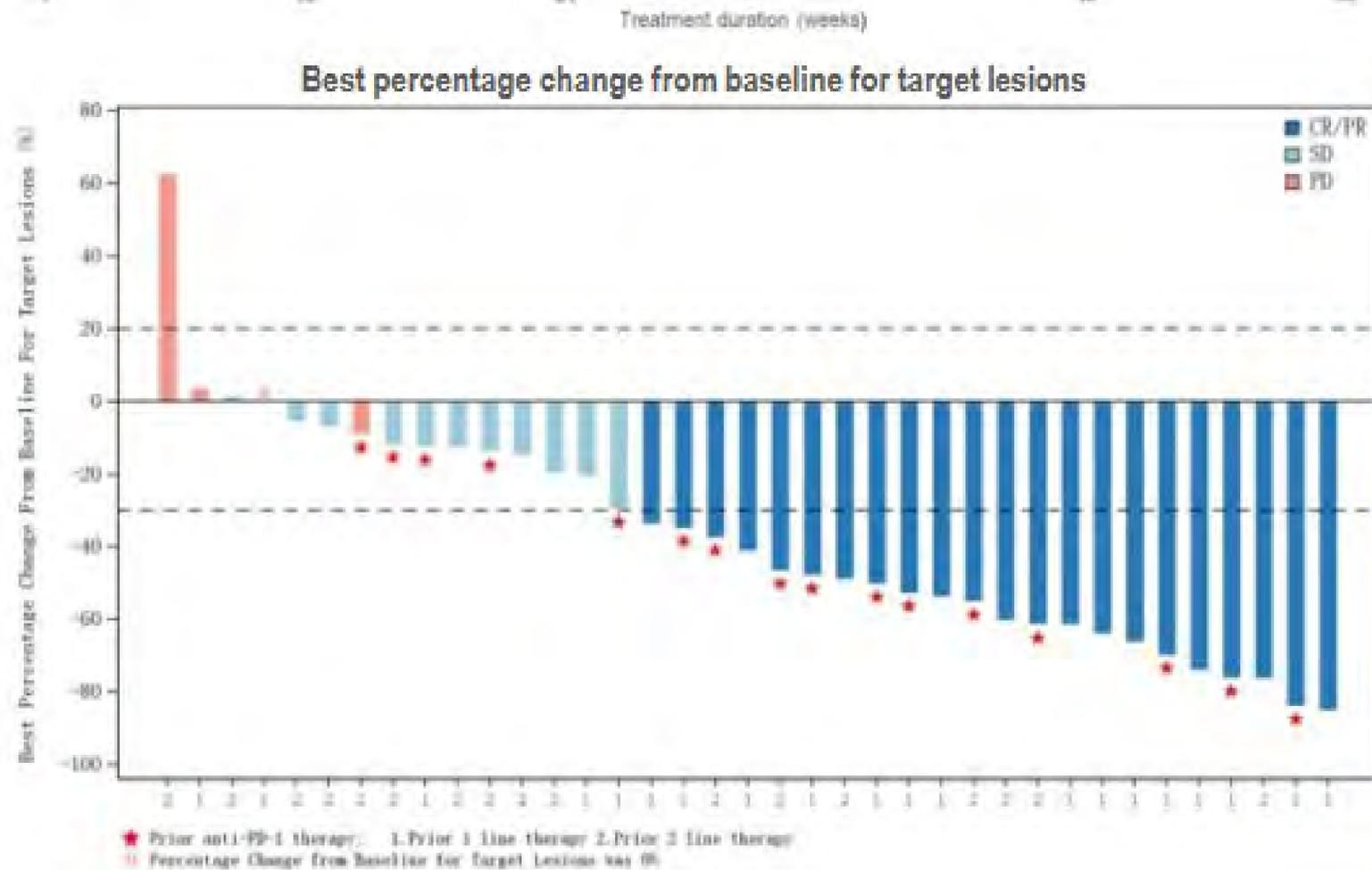
1. Tisotumab vedotin
2. Cemiplimab
3. Trastuzumab Deruxtecan
4. Nivolumab
5. Chemotherapy
6. Other



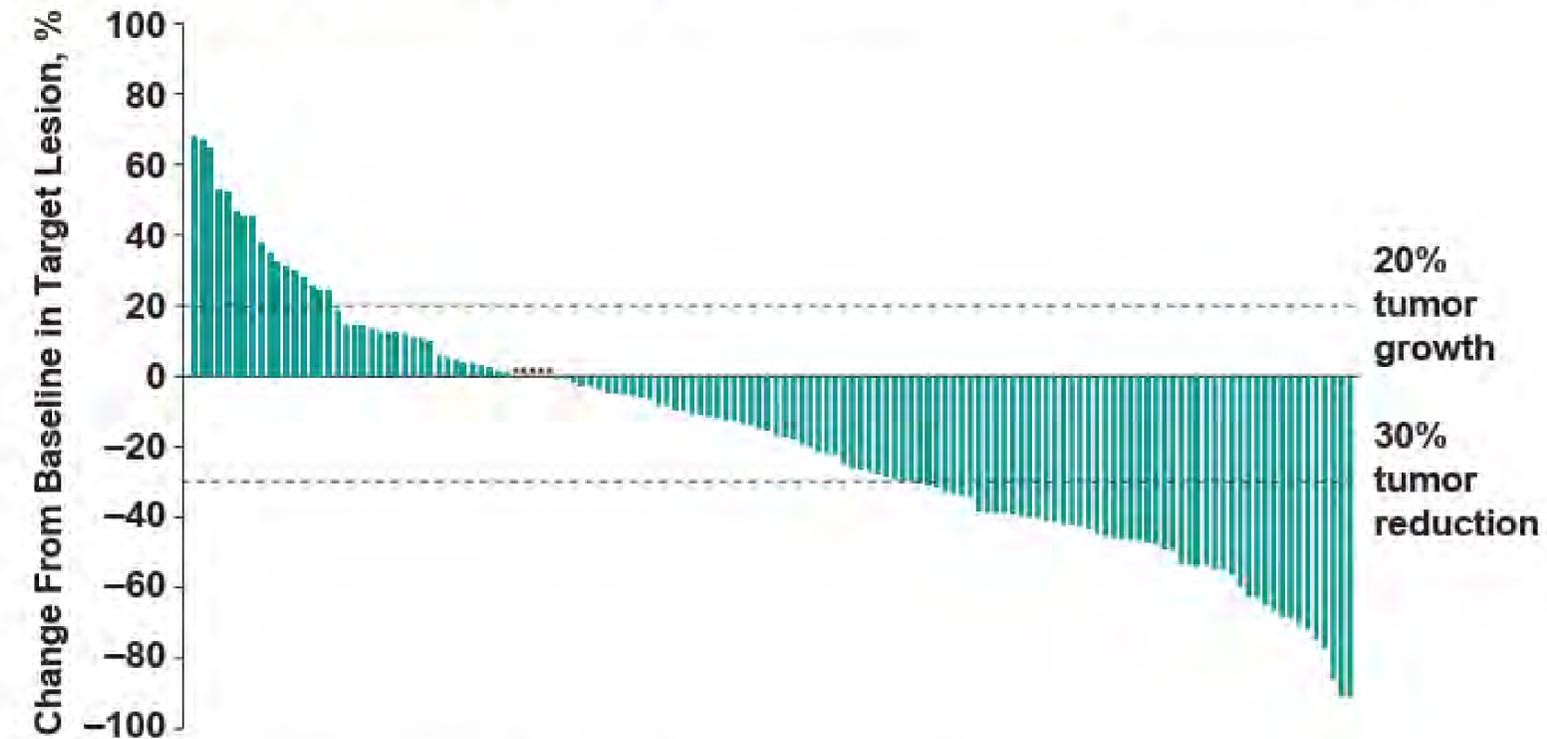
# TROP 2 ADC: Sacituzumab Tirumotecan

**SacTMT + Pembrolizumab**  
**ORR 57.9%**

**Sac TMT**  
**ORR 24.2%**  
**DCR 71.9%**  
**DOR 7.5 m**



**Figure 4. Best percentage change from baseline in target lesion size**



\*Percentage change from baseline for target lesions was 0%.

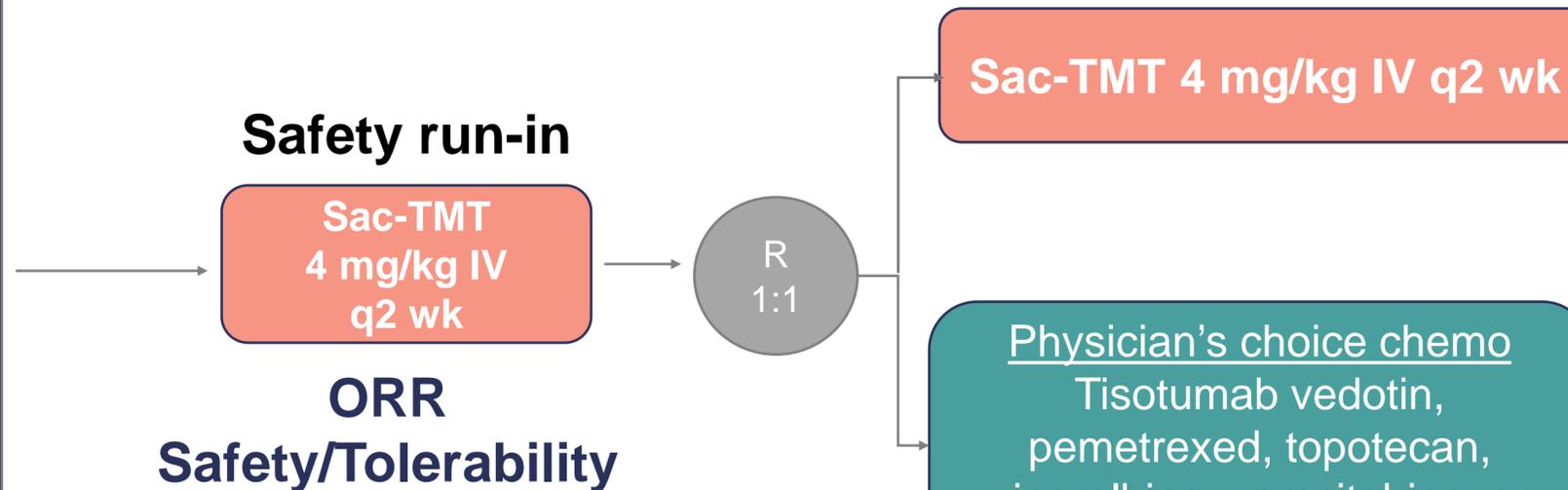
# GOG-3101/TroFuse-020/ENGOT-cx20

A Phase 3 Randomized, Active-controlled, Open-label, Multicenter Study to Compare the Efficacy and Safety of MK-2870 Monotherapy Versus Treatment of Physician's Choice as Second-line Treatment for Participants with Recurrent or Metastatic Cervical Cancer

PI: Ritu Salani, MD

## Eligibility

- Squamous, adenosquamous, adenocarcinoma cervical cancer
  - Recurrent or metastatic:
    - Progressed on or after treatment with 1 prior line of systemic platinum doublet chemotherapy (with or without bevacizumab) NOTE: may have also received prior chemoradiotherapy in the LACC setting
- AND**
- Received anti-PD-1/anti-PD-L1 therapy as part of prior cervical cancer regimens
  - Measurable disease per RECIST 1.1
  - ECOG PS 0-1
  - <Grade 2 PN



## Primary Endpoint

- OS

## Secondary Endpoints

- PFS
- ORR
- DOR
- Safety/Tolerability
- PROs
  - Time to first deterioration EORTC-QLQ-C30
  - Change baseline C30
    - Health status
    - QOL
    - Physical functioning
    - Role functioning

# ADCs in Development

	n	Target	Payload	DAR	ORR	mDOR
Disitamab vedotin	17	Her2	MMAE	4	43.8%	5.52 mo
SacTMT +pembrolizumab	38	Trop2	Belinotecan derivative	7.4	57.9%	NR
SacTMT	153	"	"	"	24.2%	7.5
Sacituzumab govitecan	18	Trop2	SN-38	7.6	50%	9.2
9MW2821	53	Nectin4	MMAE	4	35.8%	Not rep.

# Key Points

- ADCs are emerging as a key strategy in the management of recurrent cervical cancer
  - Tisotumab vedotin has shown an overall survival advantage
  - Trastuzumab deruxtecan has shown impressive responses in HER2 2-3+
- Toxicity management is key
  - Ocular toxicities
  - Interstitial Lung disease
- Clinical trials are ongoing
  - TROP2 targets
  - NECTIN4
  - Combinations

**ENROLL YOUR PATIENTS!**



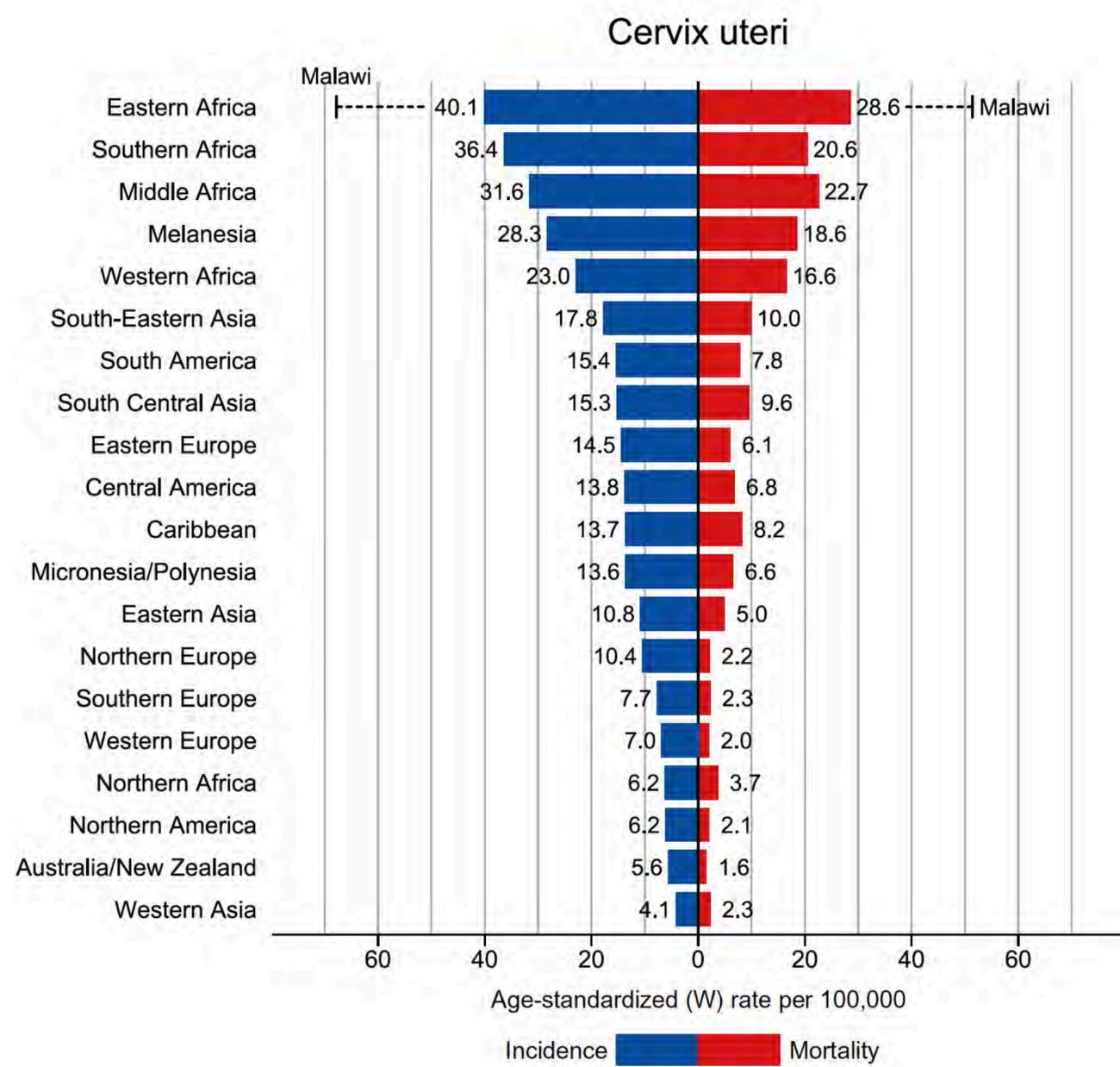
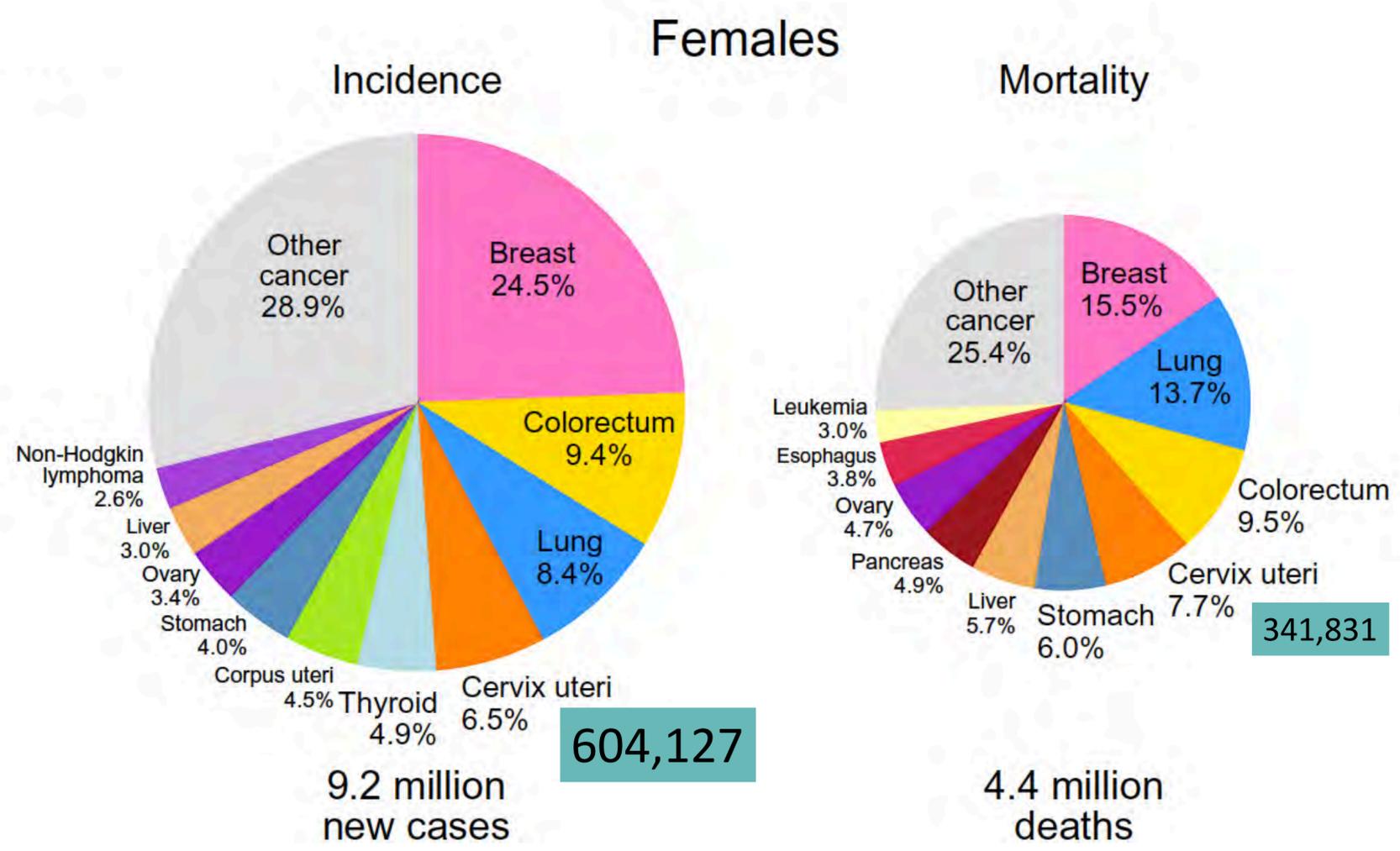
# Global Epidemiology & Prevention Strategies

**Leslie Randall, MD, MAS**

Inova Health System  
Washington, DC, USA

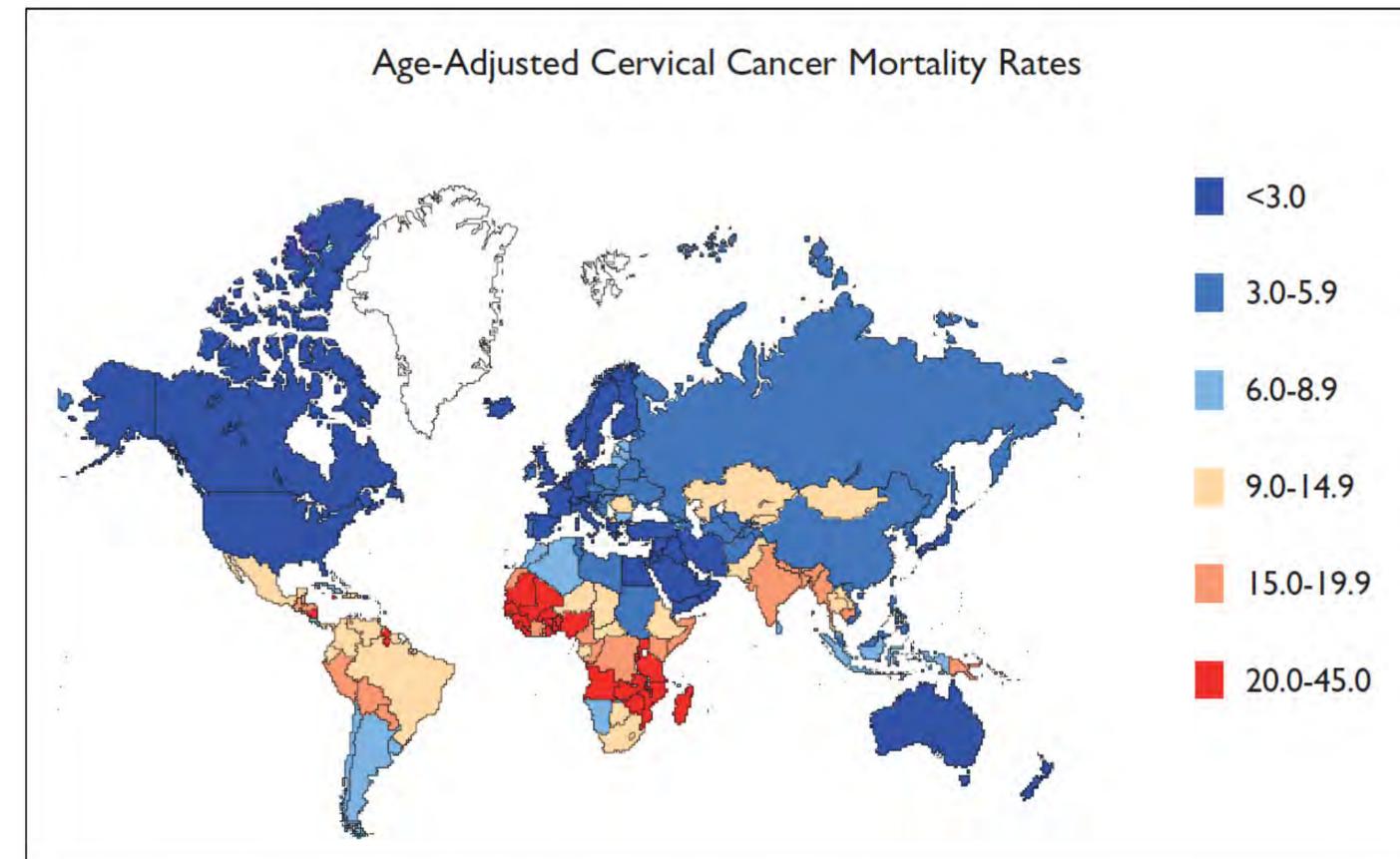
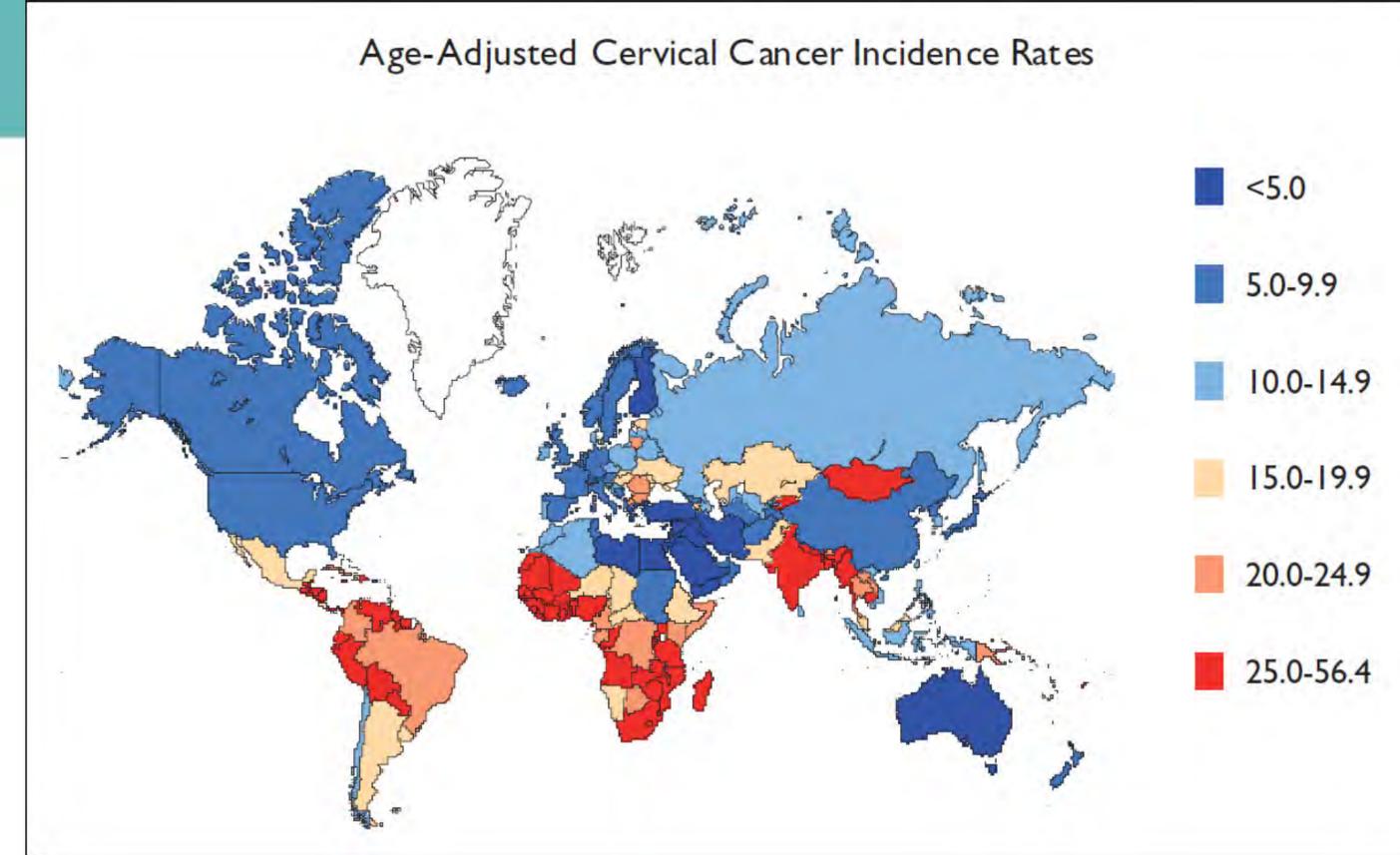


# Region-Specific Incidence and Mortality Rates for Cervical Cancer in 2020



Global disparities in cervical cancer treatment access are significant, with nearly 94% of deaths occurring in low- and middle-income countries due to limited access to screening, vaccination, and advanced treatment like surgery and radiotherapy. These disparities are driven by systemic issues including poverty, inadequate healthcare infrastructure, shortages of specialists and equipment, and lack of health insurance. Addressing this requires global efforts to expand prevention, improve diagnostic and treatment capabilities, and strengthen healthcare systems in affected regions.

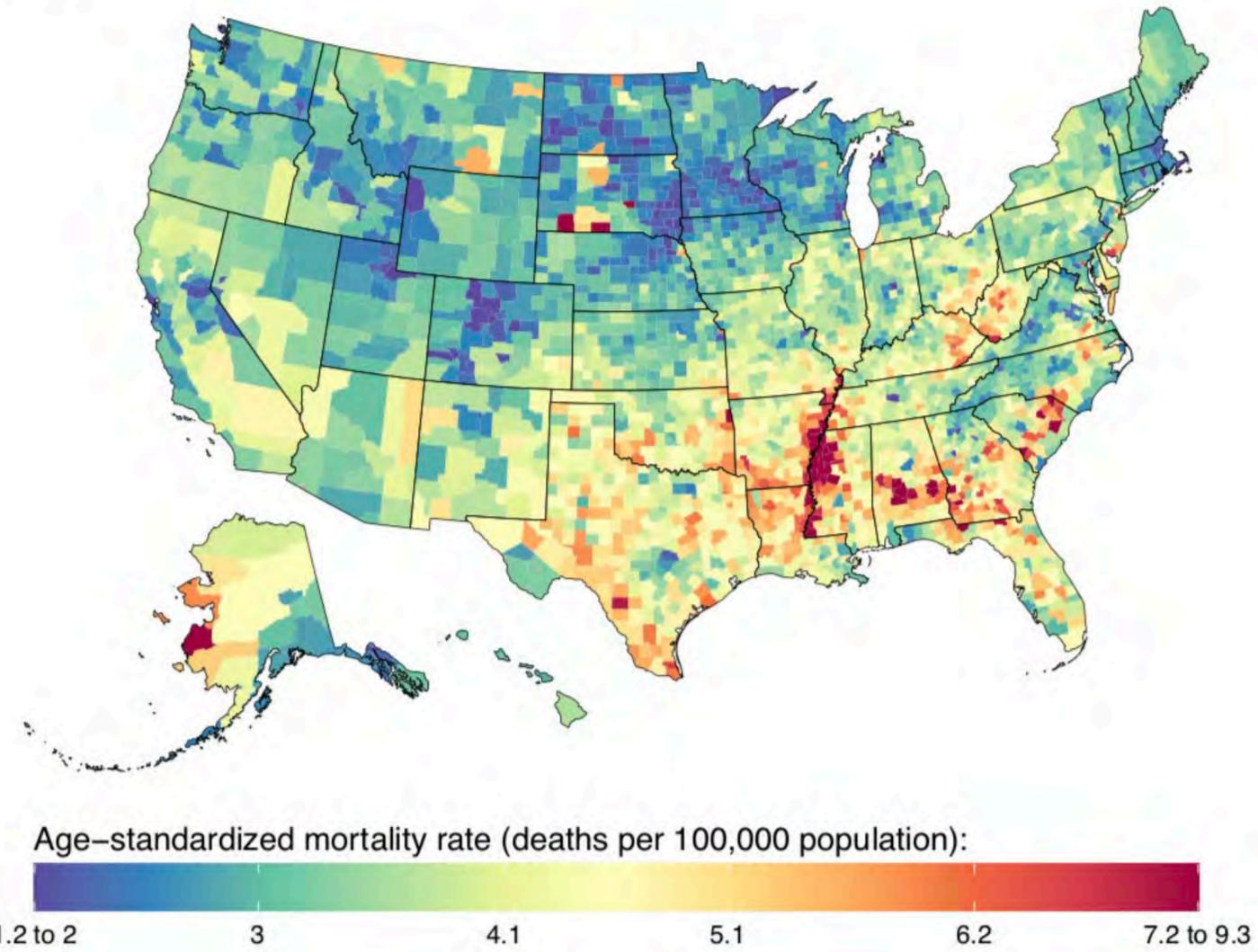
[https://www.who.int/news-room/fact-sheets/detail/cervical-cancer#:~:text=Globally%2C%20cervical%20cancer%20is%20the,to%20cervical%20cancer%20\(2\).](https://www.who.int/news-room/fact-sheets/detail/cervical-cancer#:~:text=Globally%2C%20cervical%20cancer%20is%20the,to%20cervical%20cancer%20(2).)



# Cervical cancer is relevant in the United States



[A]



In 2017, there were an estimated 291,704 women living with cervical cancer in the United States.

Mokdad AH et al. *JAMA*. 2017;317(4):388–406.



Credits +



# Launch of the Global Strategy to Accelerate the Elimination of Cervical Cancer

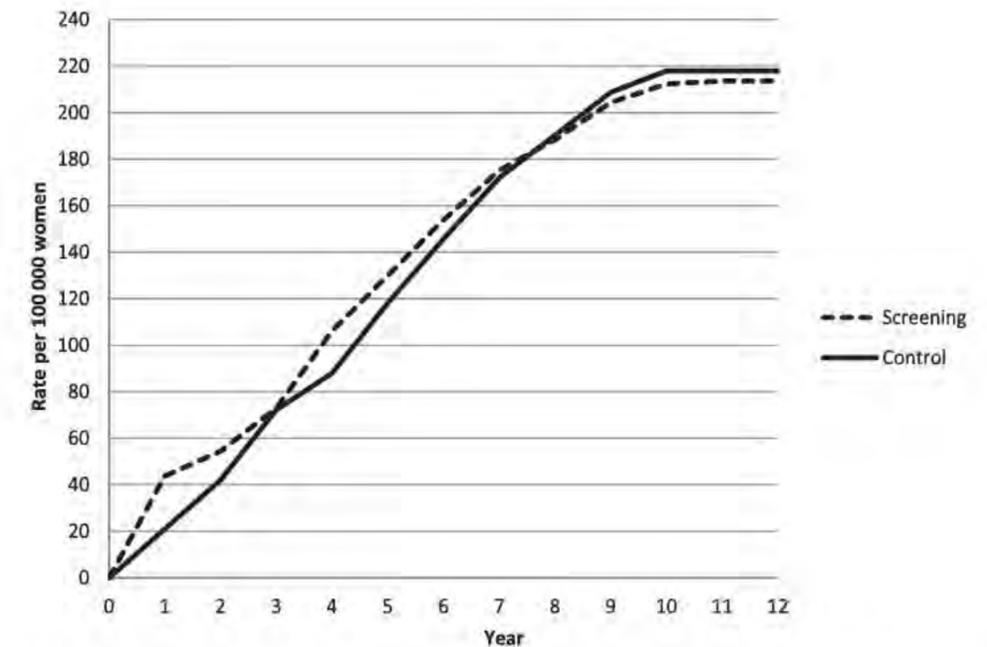
- العربية
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17 November 2020 14:30 – 17:45 CET

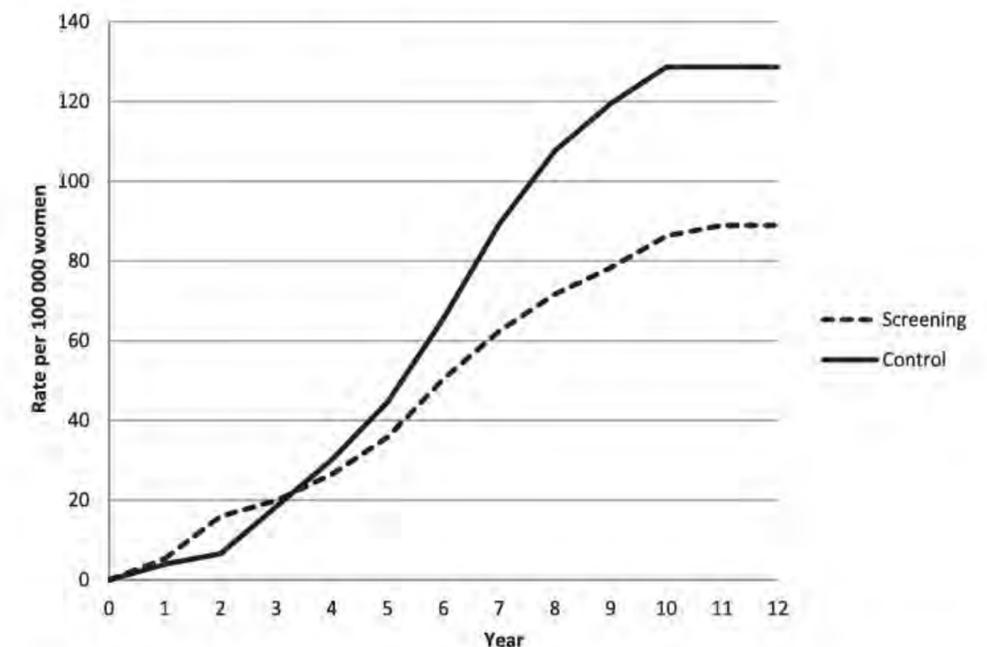
Related

# VIA in Mumbai, India

- A cluster-randomized controlled trial
- Visual inspection with acetic acid (VIA)
- Primary health workers
- 75360 women screening group
- 76178 women control group.
- In the screening group, we achieved 89% participation for screening and 79.4% compliance for diagnosis confirmation.
- The incidence of invasive cervical cancer was 26.74 per 100000 (95% confidence interval [CI] = 23.41 to 30.74) in the screening group and 27.49 per 100000 (95% CI = 23.66 to 32.09) in the control group.
- Compliance to treatment for invasive cancer was 86.3% in the screening group and 72.3% in the control group.
- The screening group showed a statistically significant 31% reduction in cervical cancer mortality (RR = 0.69; 95% CI = 0.54 to 0.88;  $P = .003$ ).

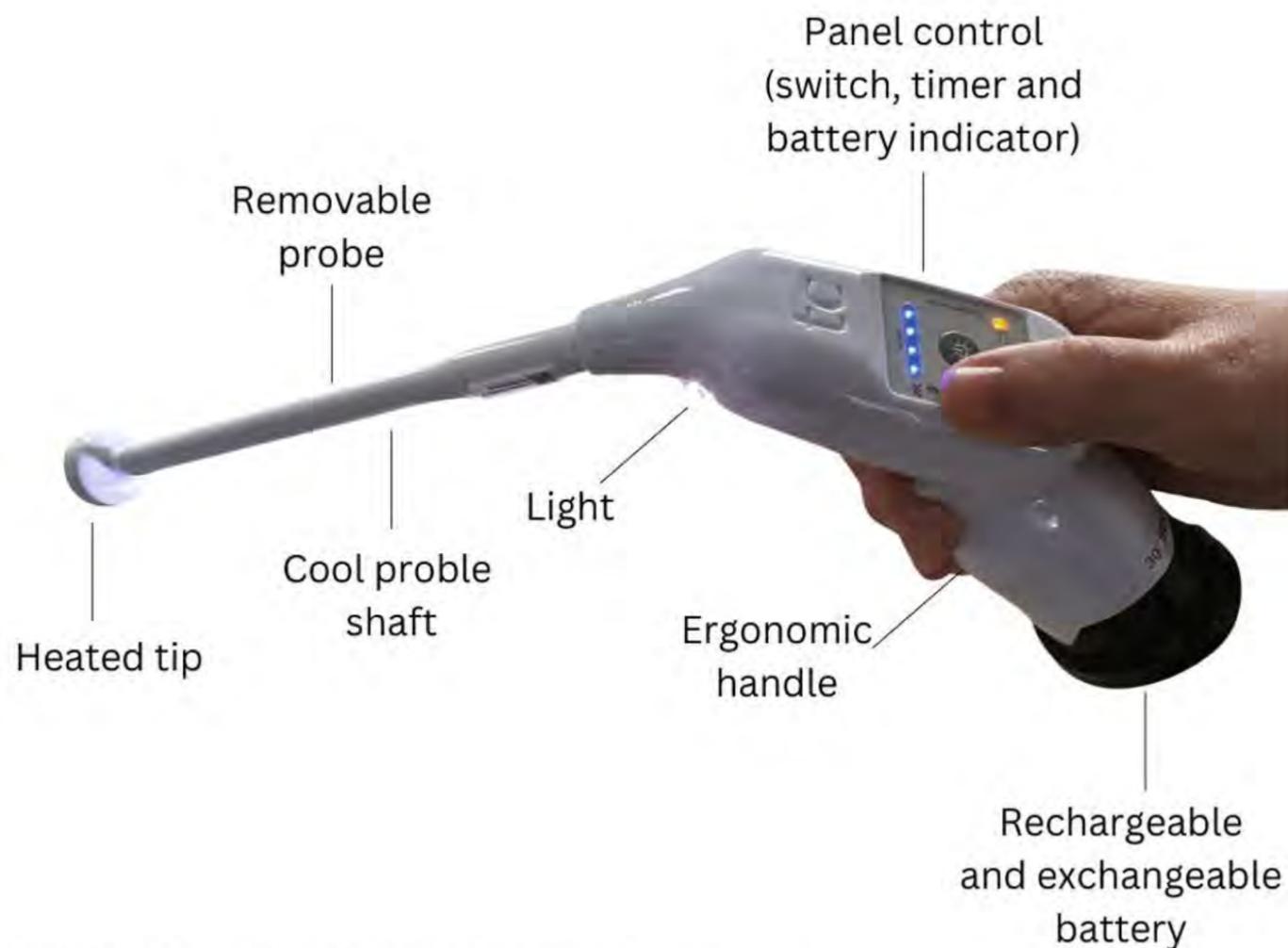


**Figure 2.** Cumulative incidence curve for cervix cancer. Cumulative incidence of cervical cancer in the screening and control groups after 12 years of follow-up. The screening group recruited 75360 women aged 35 to 64 years and had an annual attrition of 2.2%, whereas the control group recruited 76178 women also aged 35 to 64 years and had an annual attrition of 2.3%.



**Figure 3.** Cumulative mortality curve for cervix cancer. Cumulative mortality from cervical cancer in the screening and control groups after 12 years of follow-up. The screening group recruited 75360 women aged 35 to 64 years and had an annual attrition of 2.2%, whereas the control group recruited 76178 women also aged 35 to 64 years and had an annual attrition of 2.3%.

# A portable thermal ablation device for cervical cancer prevention in a screen-and-treat setting: a randomized, noninferiority trial



Extended Data Fig. 1 | A portable battery-driven thermal ablation device showing different components.

- Randomized controlled noninferiority trial in Zambia
- Portable, battery-driven thermal ablation (TA) device compared to cryotherapy and electrosurgical excision (large loop excision of transformation zone (LLETZ))
- 3,124 women VIA +
- HPV clearance or a negative VIA
- After a median follow-up of 12 months, treatment success rates were 74.0%, 71.1% and 71.4% for the TA, cryotherapy and LLETZ arms, respectively, thus demonstrating noninferiority (P = 0.83).
- Only 3.6% of those randomized to TA reported moderate-to-severe pain, compared to 6.5% and 1.9% for the cryotherapy and LLETZ arms, respectively.

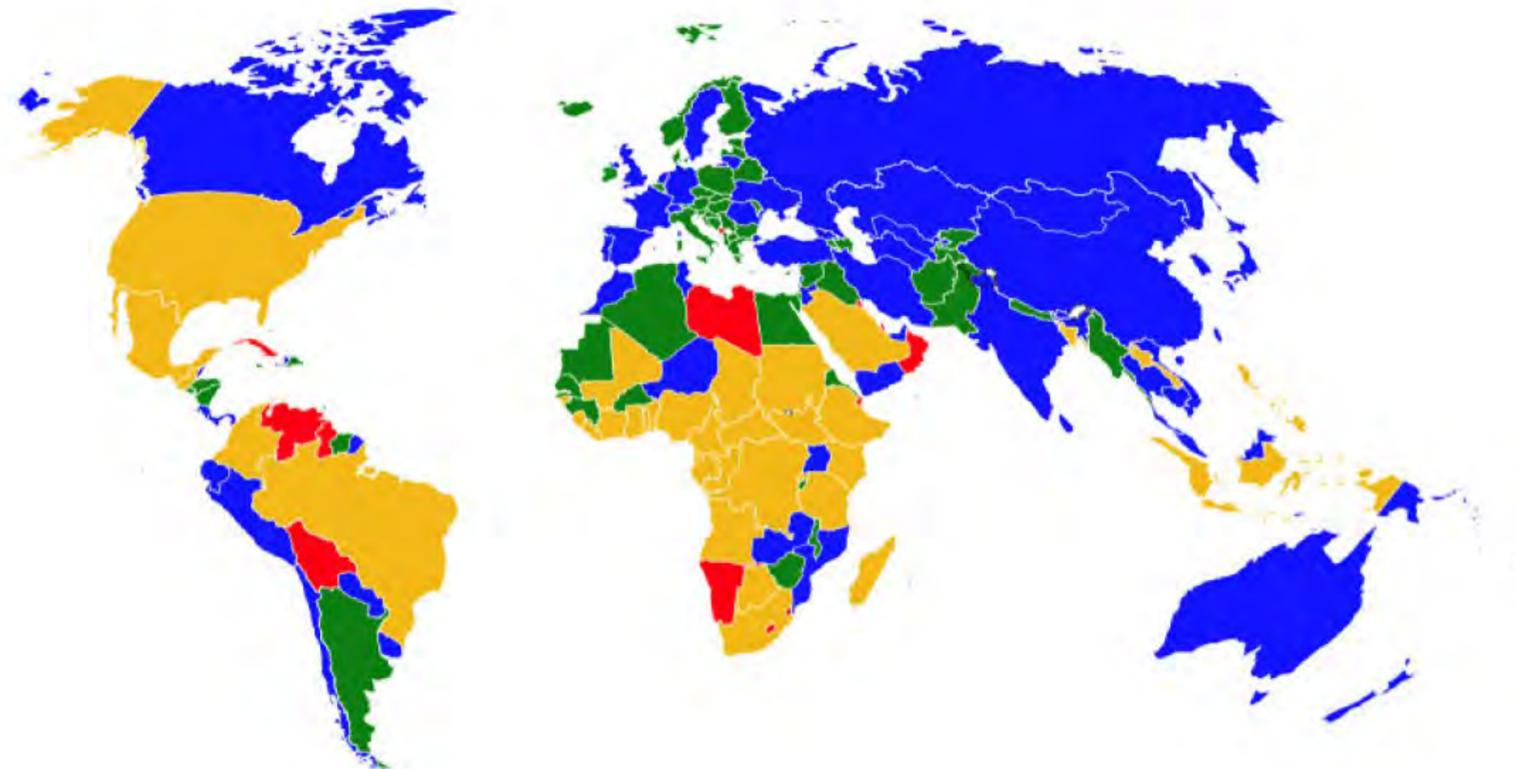
Risk ratio for TA versus cryotherapy (95% CI) <sup>b</sup>	1.01 (0.96–1.07)
Risk ratio for TA versus LLETZ (95% CI) <sup>c</sup>	1.02 (0.95–1.08)

Basu P, Mwanahamuntu M, Pinder LF, Muwonge R, Lucas E, Nyambe N, Chisele S, Shibemba AL, Sauvaget C, Sankaranarayanan R, Prendiville W, Parham GP. A portable thermal ablation device for cervical cancer prevention in a screen-and-treat setting: a randomized, noninferiority trial. *Nat Med.* 2024 Sep;30(9):2596-2604.

# Elimination can be geographically complex

## Global Feasibility Landscape for Cervical Cancer Elimination: Four Distinct Implementation Clusters

- High Readiness and Advanced Implementation
- Transitional Feasibility and Growing Momentum
- Complex Sociocultural and Geopolitical Barriers
- Low Resilience, High Vulnerability Settings



*Introducing*  
**IGCS ACCESS  
Series**

**Accelerating Cervical  
Cancer Elimination  
Strategies Symposia**

*A bold 5-year plan*



Supported by the  
IGCS World of Hope  
Development Fund



TOGETHER, WE CAN  
**ELIMINATE  
CERVICAL  
CANCER**

**Inaugural Symposium  
Friday, Nov. 7th here in Cape  
Town to focus on Africa's  
elimination strategy**

**Region-focused symposium to be  
held at each IGCS Annual Global  
Meeting through 2030**

**Development of an online resource  
hub and navigational tool to  
centralize knowledge and guidance**

# IGCS Vision and Purpose Statement

## VISION:

**“All high burden regions will have a national cervical cancer elimination strategy that is being implemented - shaping locally adapted programme scale-up and unlocking domestic and innovative funding mechanisms”**

## PURPOSE:

**IGCS commits to 5-year focus on building a *Cervical Cancer Elimination Resource Hub and Navigational Tool* and leveraging these resources for regional and national deployment and impact in LMICs with a focus on high burden countries**

**Phase I: Announce the *IGCS ACCESS series*, set up hub and conduct regional deep dives**

**Phase II: Nurture and implement IGCS and other partner responses to gaps and needs; advocate for and feed into Global CCE Forum dialogues**

**Phase III: IGCS regional and 5-year summary reports to drive national action and shape CCEI targets and actions post-2030**

# IGCS ACCESS Leadership & Advisory Board



**Mary Eiken,  
MS IGCS CEO**



**Julie Torode, PhD  
IGCS ACCESS Consultant**



**Dr. Alexander  
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**Dr. Heleen van  
Beekhuizen**

# ACCESS Landscaping & Analysis Team

## University of Sydney



Led by  
**Dr. Karen Canfell**



**Dr. Deborah Bateson**



**Dr. Michael Caruna**



**Dr. Telma Costa**



**Dr. Diep Nguyen**

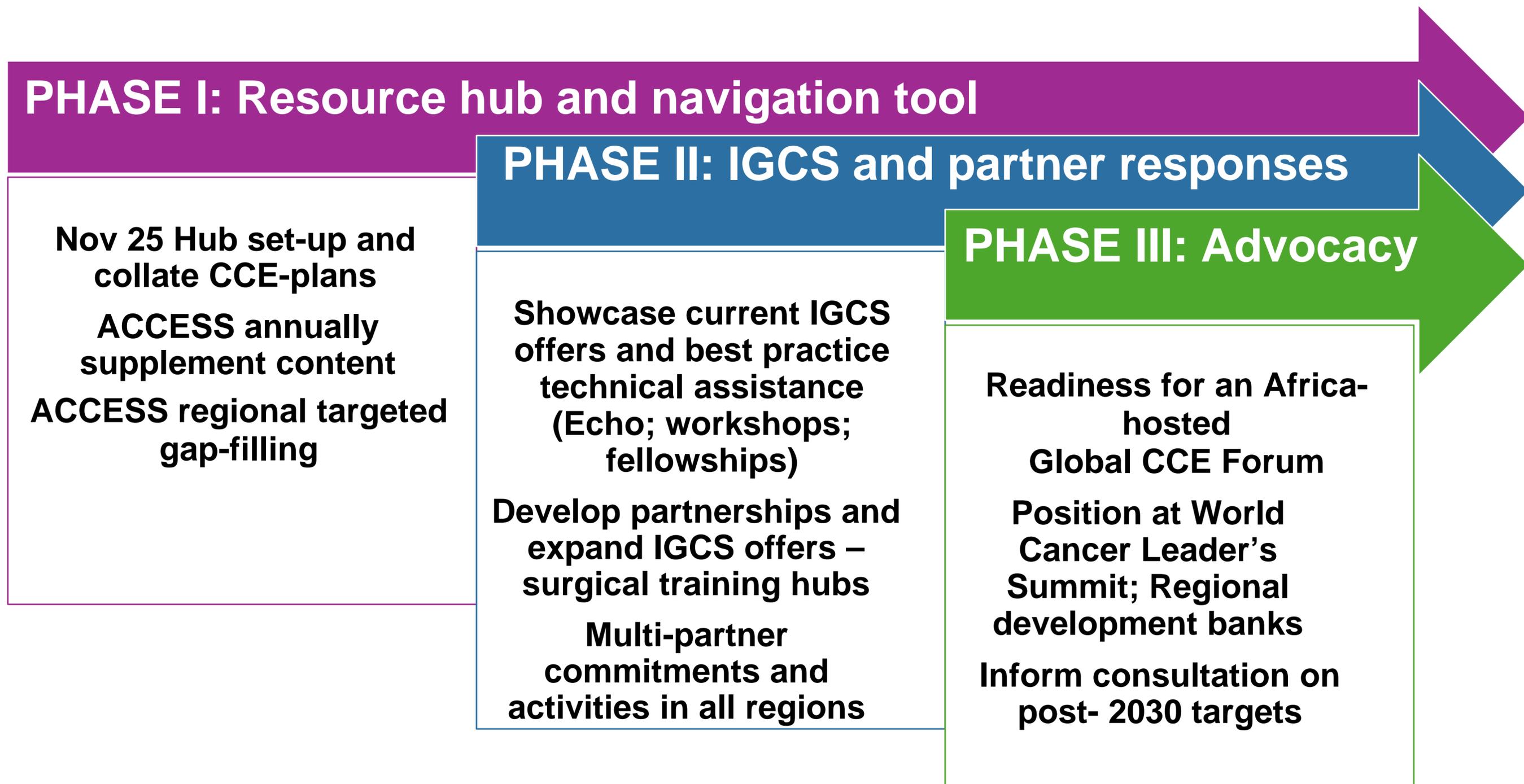


**Dr. Ai Ling  
Tan**



**Dr. Kate Simms**

# Building more than a resource hub



# ACCESS: Africa

Accelerating Cervical Cancer  
Elimination Strategies Symposia

TOGETHER, WE CAN  
**ELIMINATE**  
**CERVICAL**  
**CANCER**



Cape Town, South Africa  
Nov. 7-8



**IGCS**



INTERNATIONAL  
GYNECOLOGIC  
CANCER SOCIETY

Supported by the IGCS World of Hope Development Fund



**Pillar 1**  
**HPV Vaccination & Public Awareness**

**Pillar 2**  
**Screening & Treatment of Precancerous Lesions**

**Pillar 3**  
**Treatment & Palliative Care of Invasive Cancers**

**ACCESS Series Pillar Leads**



**Dr. Suzanne Garland**



**Prof. Karen Canfell**



**Dr. Kathleen Schmeler**

**ACCESS: Africa Pillar Leads 2025**



**Dr. Nelly Mugo**



**Dr. Deborah Watson-Jones**



**Dr. Laura Muzingwani**



**Dr. Rakiya Saidu**



**Dr. Surbi Grover**



**Dr. Anisa Mburu**



**Dr. Eve Namisango**

# Make your commitment

Visit the IGCS Booth in the exhibit hall

- Sign the pledge board to display your commitment
- Take a photo and post online with #IGCSAccessAfrica & #IGCS2025



**IGCS ACCESS Africa**

Accelerating Cervical Cancer  
Elimination Strategies Symposia



# *Please join us* **ACCESS Africa** Reception

- **Urban Umami Restaurant  
Century City Hotel Urban  
Square**
- **Friday, November 7, 2025**
- **4:30 – 6:00 PM**
- **Immediately after the  
Closing Ceremony**



# Panel Discussion and Q&A, and Closing Comments

All Faculty





# Thank You

View this symposium as part of the IGCS  
on-demand program following the meeting

