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# Cáncer de pulmón microcítico y otros tumores

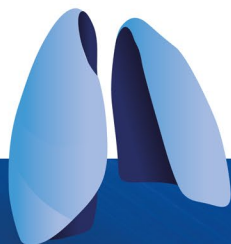
Andrés Barba Joaquín

*Hospital de la Santa Creu i Sant Pau*

# CONFLICTO DE INTERESES

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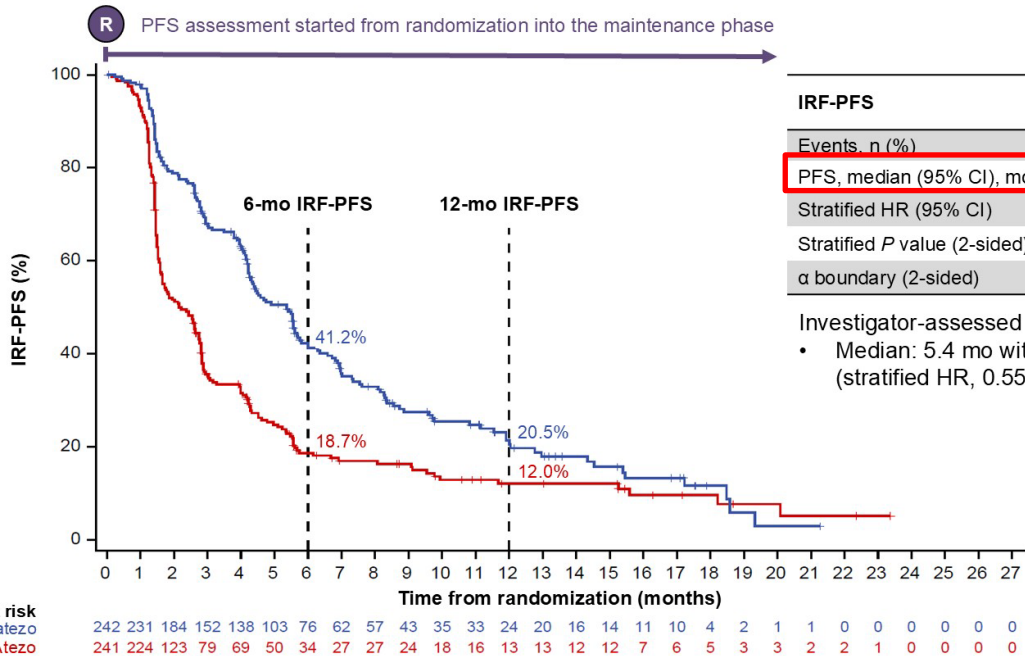
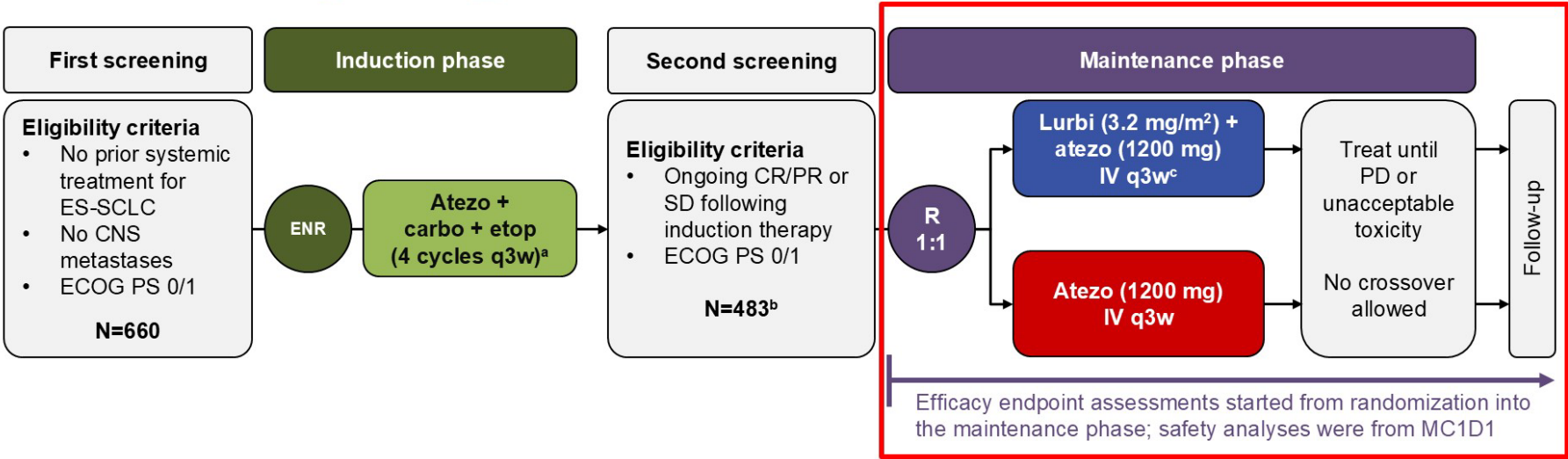
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# CPCP

## Tratamiento de mantenimiento en 1L

# MANTENIMIENTO LURBINECTEDINA-ATEZOLIZUMAB – Ph3 IMforte



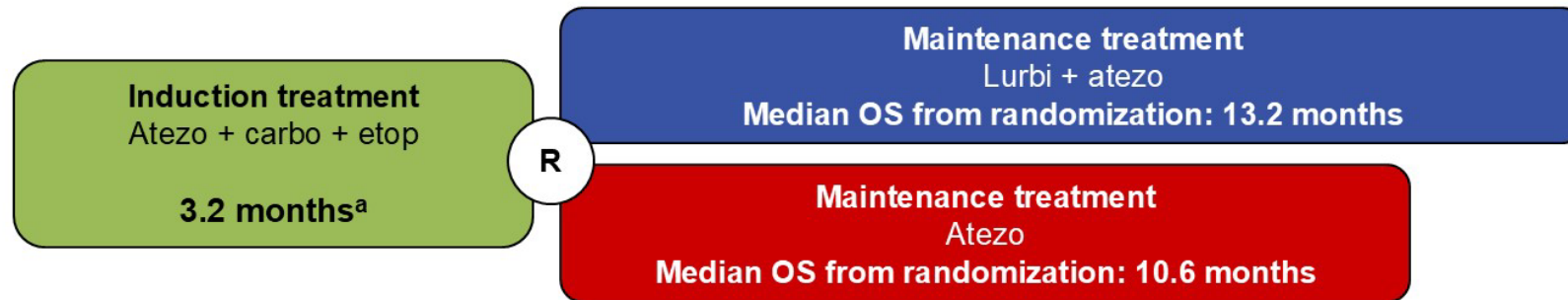
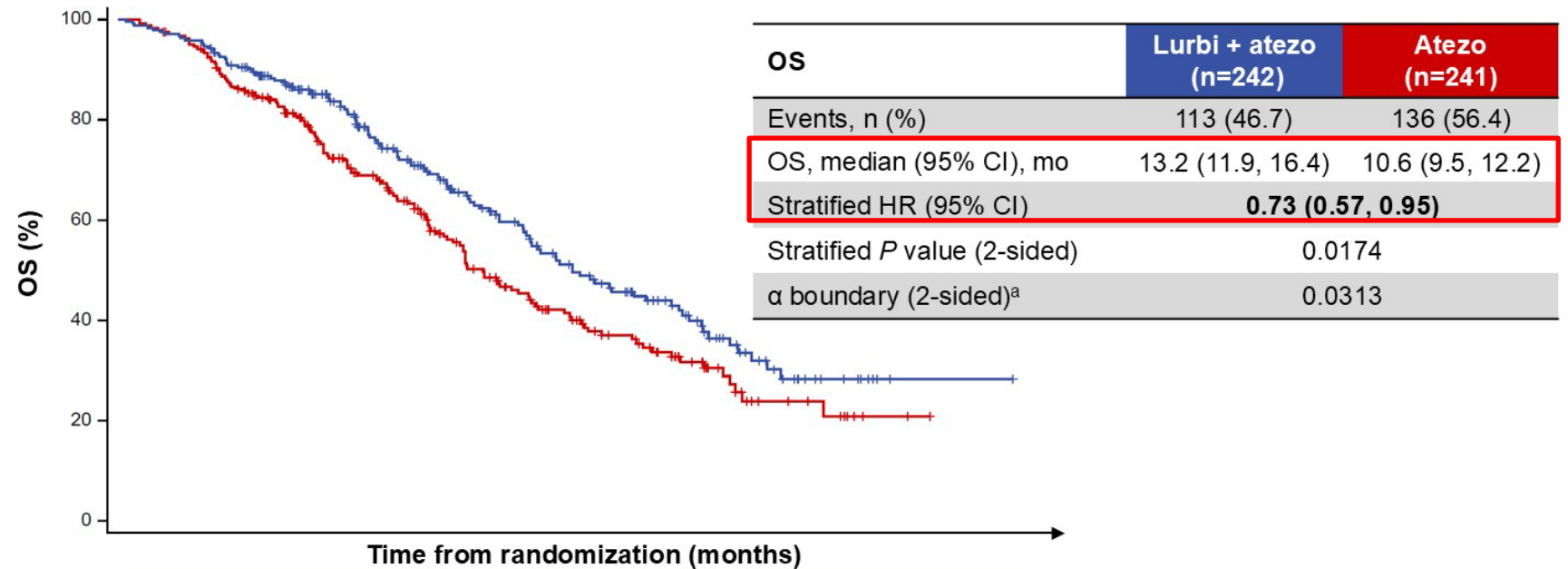
**ORR: 19.4% vs. 10.4%**  
**DoR: 9.0 m vs. 5.6m**

**PFS a 6m: +22.5%**  
**PFS a 1 año: +8.5%**

L.Paz-Ares, ASCO 2025  
L.Paz-Ares, ASCO 2025

# MANTENIMIENTO LURBINECTEDINA-ATEZOLIZUMAB – Ph3 IMforte

## OS from randomization into maintenance phase

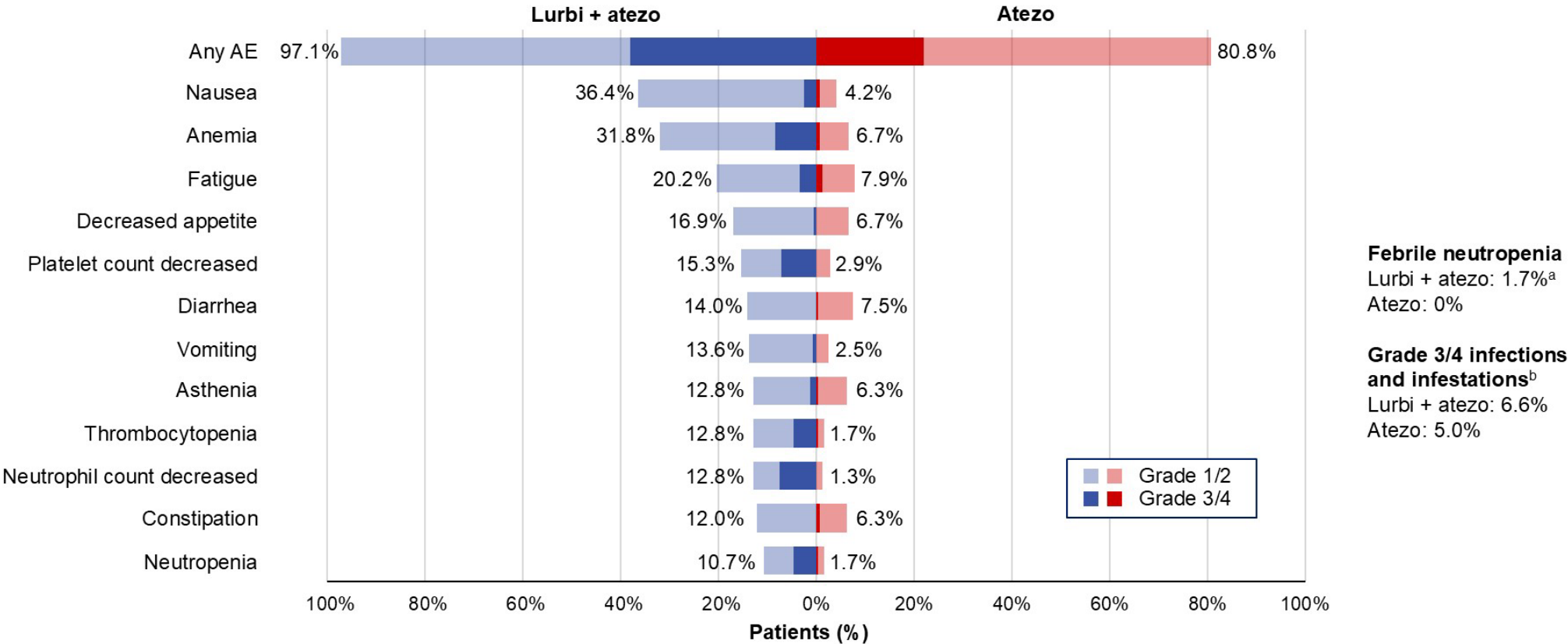


**IMforte results do not include time on induction treatment**



# MANTENIMIENTO LURBINECTEDINA-ATEZOLIZUMAB – Ph3 IMforte

## All-cause AEs with incidence $\geq 10\%$ in either arm



Clinical cutoff: July 29, 2024. Percentage labels represent all-grade AEs, including Grade 5 AEs. Grade 5 AEs occurred in 12 (5.0%) patients in the lurbi + atezo arm and 6 (2.5%) patients in the atezo arm.  
<sup>a</sup> Includes 1 Grade 5 AE. <sup>b</sup> Grade 5 infections: lurbi + atezo arm (n=6 [2.5%]): COVID-19 pneumonia, pneumonia, pneumonia viral, sepsis, septic shock, and vascular device infection (n=1 each); atezo arm (n=4 [1.7%]): pneumonia (n=2), abscess intestinal, and sepsis (n=1 each).

An illustration of two human lungs, rendered in a light blue color with a darker blue outline. They are positioned on the left side of the slide, set against a dark blue background with a subtle pattern of diagonal lines.

# T-cell engagers en CPCP

# TARLATAMAB EN 2L - Ph3 DeLLphi-304

## Key inclusion criteria

- Histologically or cytologically confirmed SCLC
- Progression after 1L platinum-based chemotherapy +/- anti-PD-(L)1
- ECOG PS 0 or 1
- Asymptomatic, treated or untreated brain metastases

## Randomization stratified by

- Prior anti-PD-(L)1 exposure (yes/no)
- Chemotherapy-free interval (< 90 days vs ≥ 90 to < 180 days vs ≥ 180 days)
- Presence of (previous/current) brain metastases (yes/no)
- Intended chemotherapy (topotecan/amrubicin vs lurbinectedin)

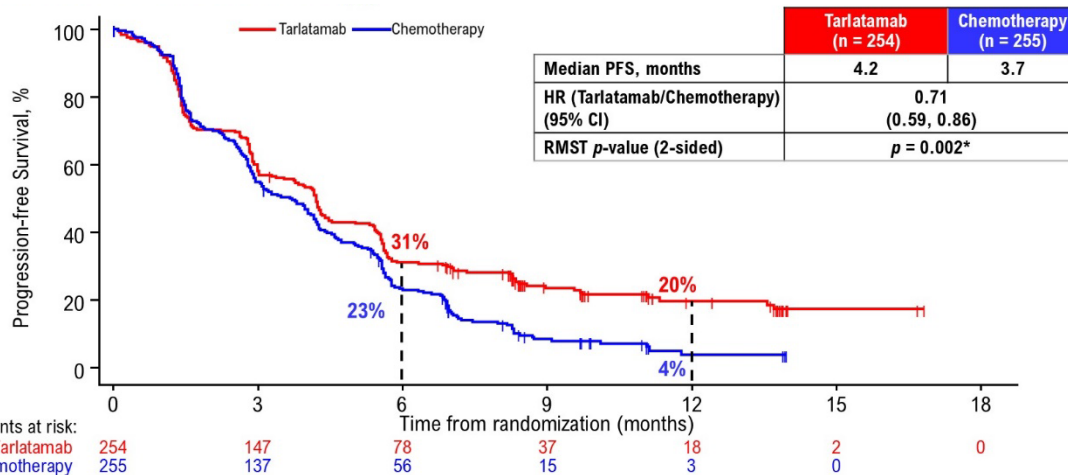
R  
1:1  
(N = 509)

**Tarlatamab (n = 254)**

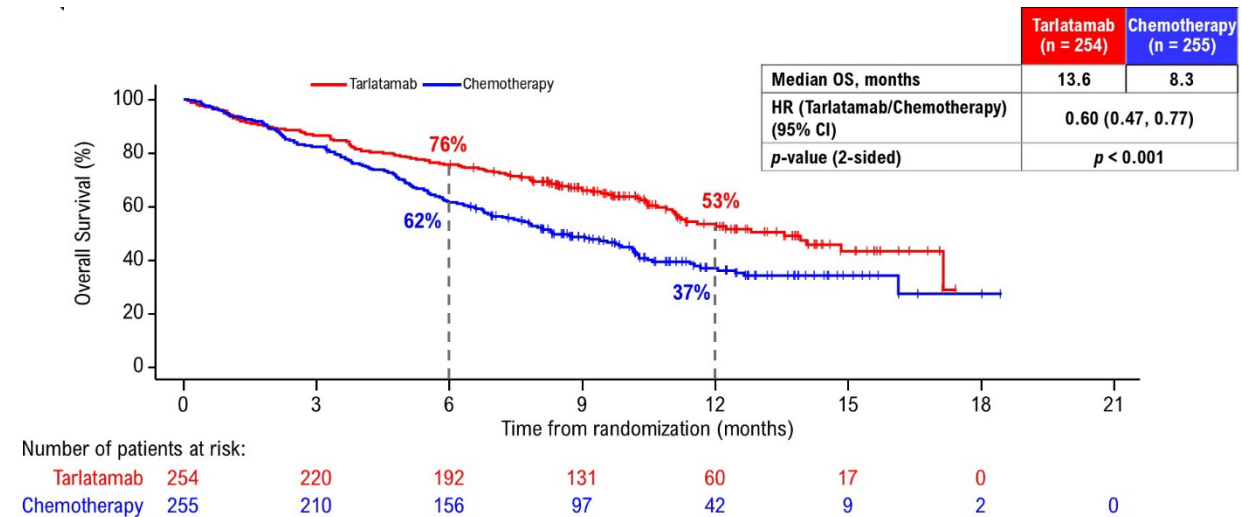
**Chemotherapy\* (n = 255)**

Topotecan (n = 185); Lurbinectedin (n = 47);  
Amrubicin (n = 23)

CFI <90d: T 43%; CT 45%  
Brain M1: T 44%; CT 45%



**PFS 1 año: +16%**

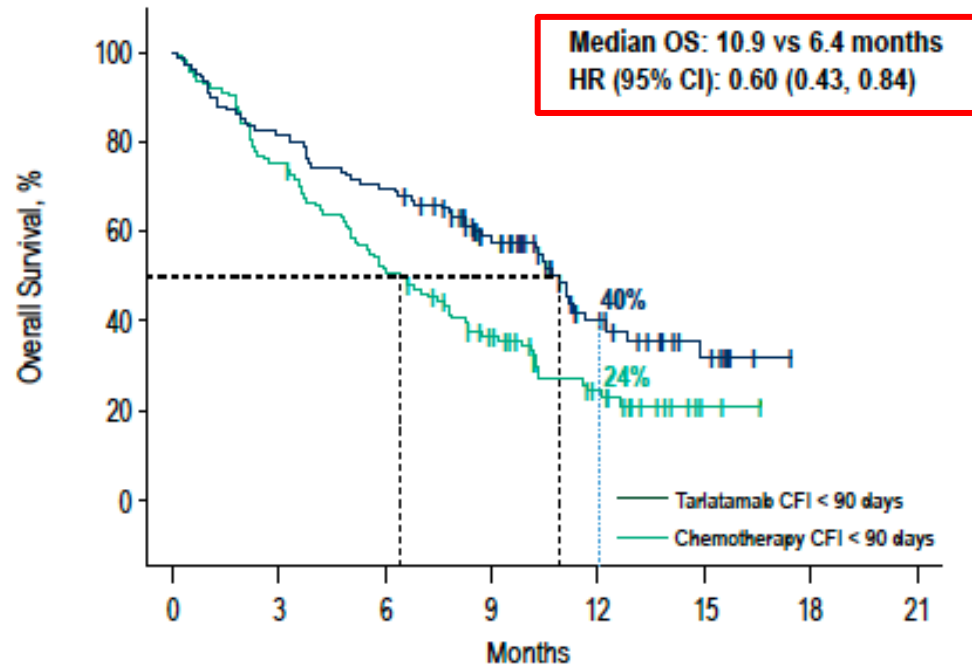


**OS 1 año: +16%**



# TARLATAMAB EN 2L - Ph3 DeLLphi-304

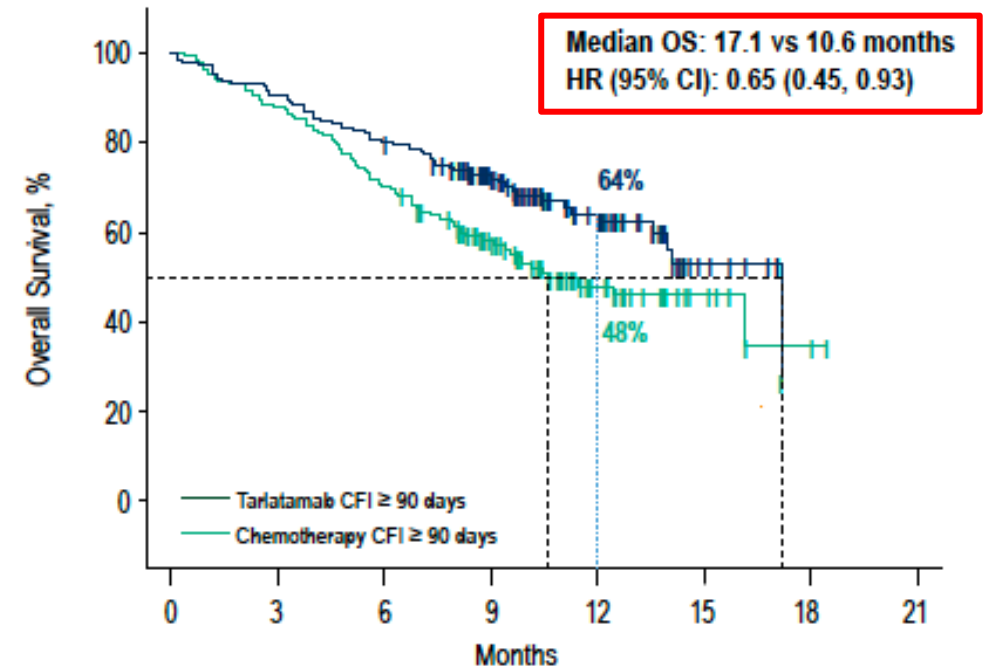
**CFI < 90 days (platinum-resistant disease)**



Number of Patients at Risk:  
Tarlatamab CFI < 90 days  
Chemotherapy CFI < 90 days

109	89	76	50	21	9	0
114	86	57	36	16	2	0

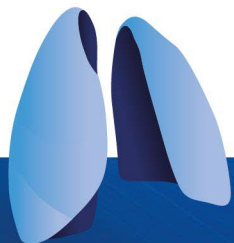
**CFI ≥ 90 days (platinum-sensitive disease)**



Number of Patients at Risk:  
Tarlatamab CFI ≥ 90 days  
Chemotherapy CFI ≥ 90 days

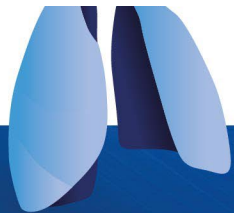
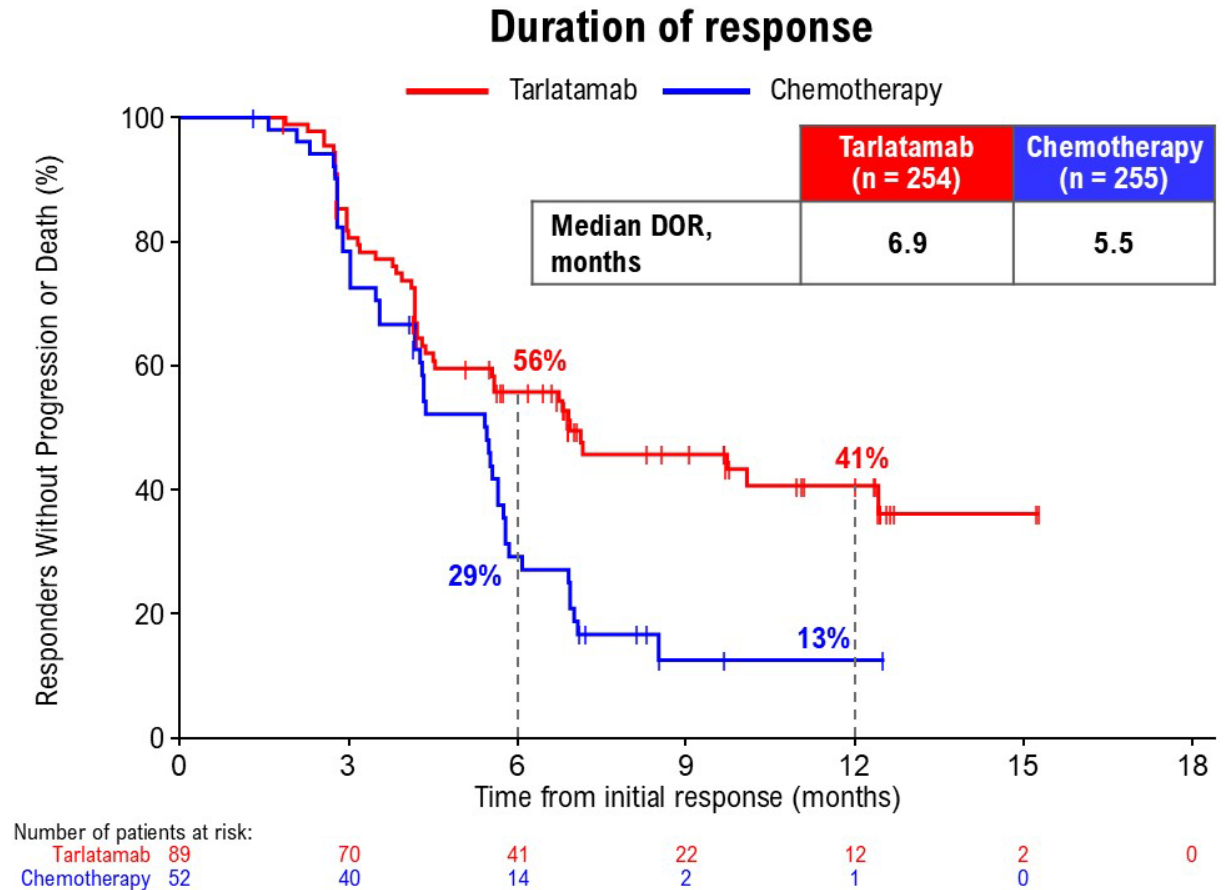
145	131	116	81	39	8	0	0
141	124	99	61	26	7	2	0

**OS - HR Brain M1: 0.45 (0.31-0.65)**



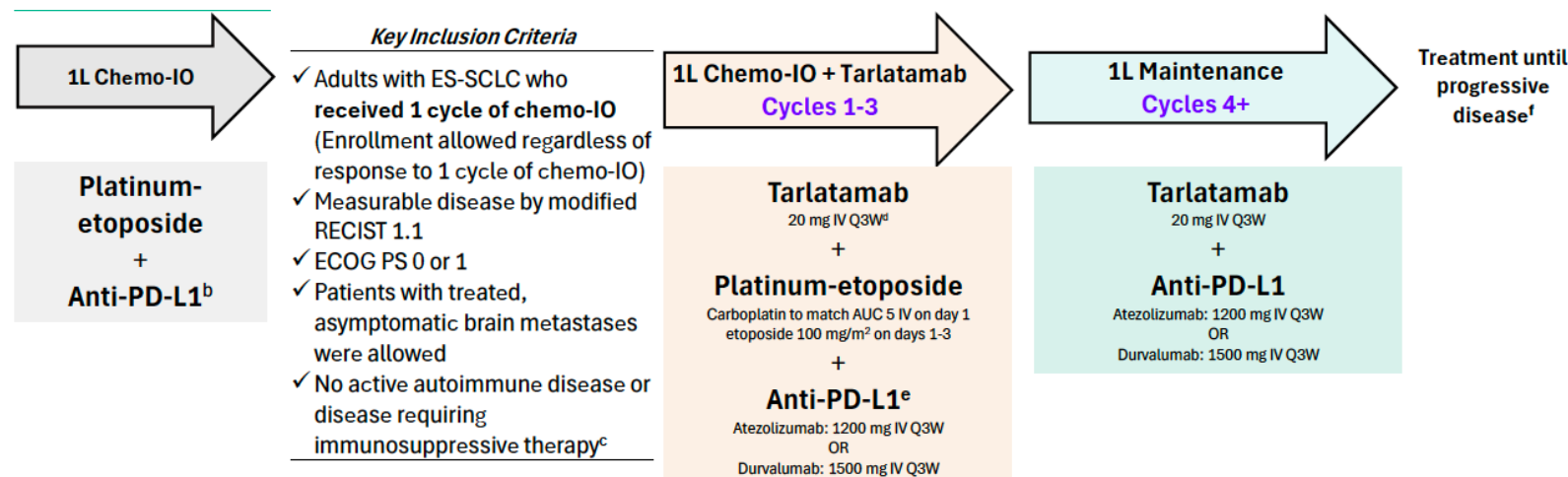
# TARLATAMAB EN 2L - Ph3 DeLLphi-304

	Tarlatamab (n = 254)	Chemotherapy (n = 255)
<b>Best overall response*†, n (%)</b>		
Complete response	3 (1)	0 (0)
Partial response	86 (34)	52 (20)
Stable disease	84 (33)	112 (44)
Progressive disease	56 (22)	50 (20)
Not evaluable/no post-baseline scan	25 (10)	41 (16)
<b>Objective response rate‡, % (95% CI)</b>	<b>35 (29–41)</b>	<b>20 (16–26)</b>
<b>Median duration of response, months</b>	<b>6.9</b>	<b>5.5</b>
<b>Median time to objective response, months</b>	<b>1.5</b>	<b>1.4</b>
<b>Ongoing response at data cutoff, n§ (%)</b>	<b>42 (47)</b>	<b>8 (15)</b>

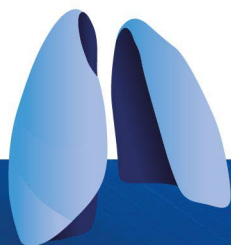
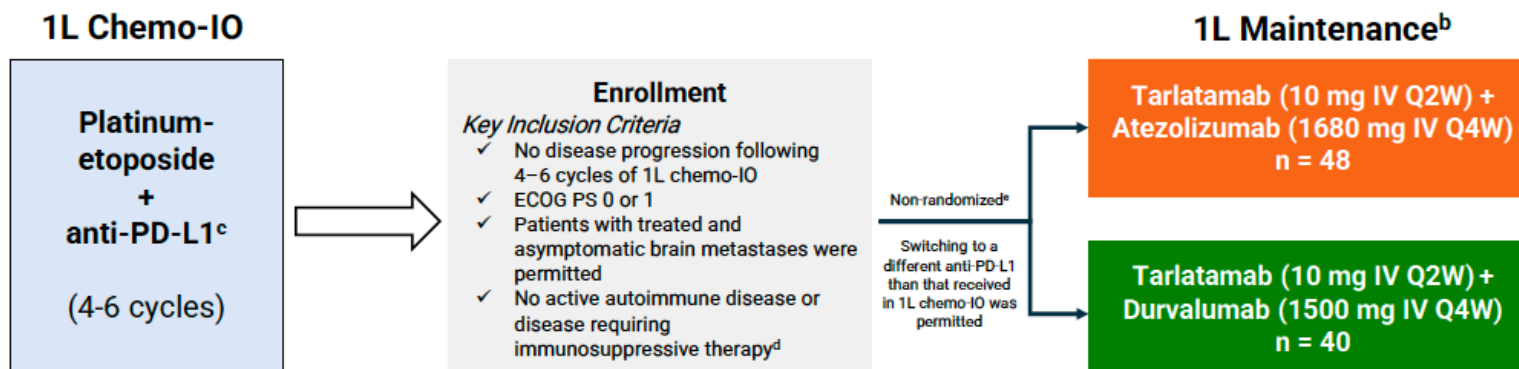


# TARLATAMAB EN 1L - Ph1b: DeLLphi-303

## Cohortes 2,4,7



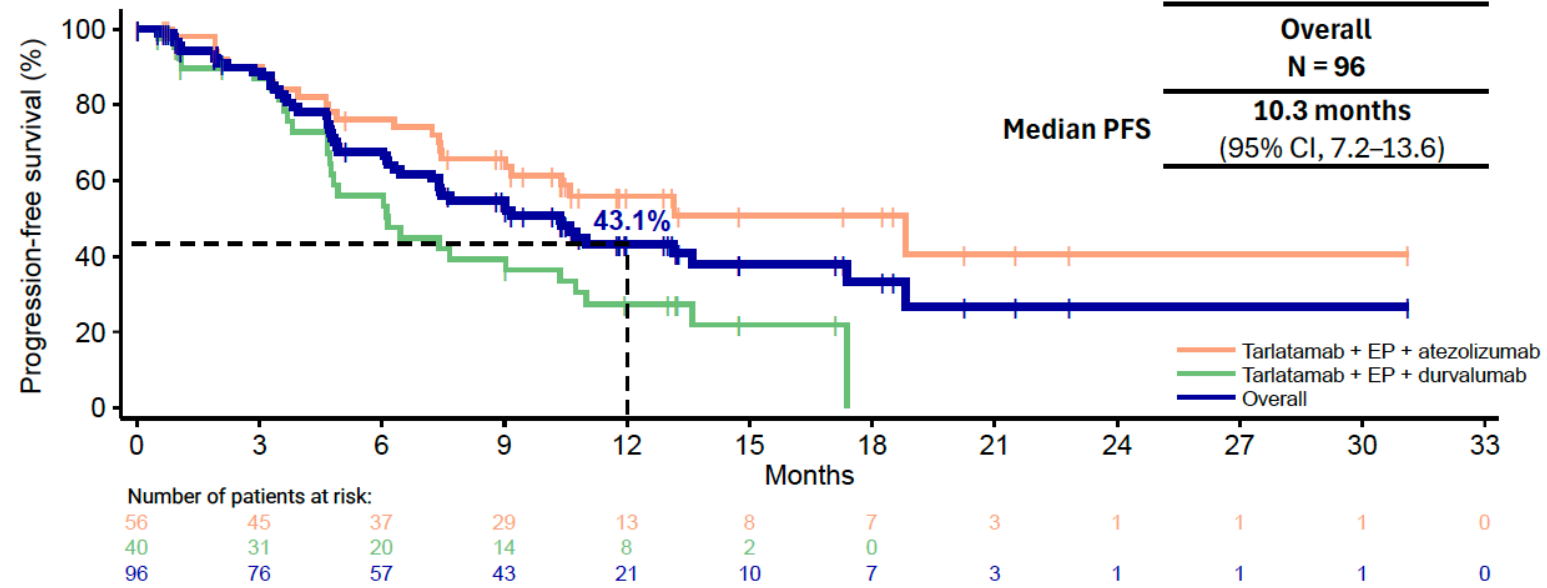
## Cohortes 5,6,8



# TARLATAMAB EN 1L - Ph1b: DeLLphi-303

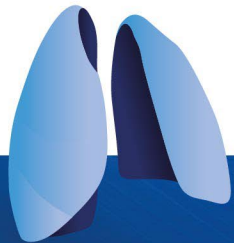
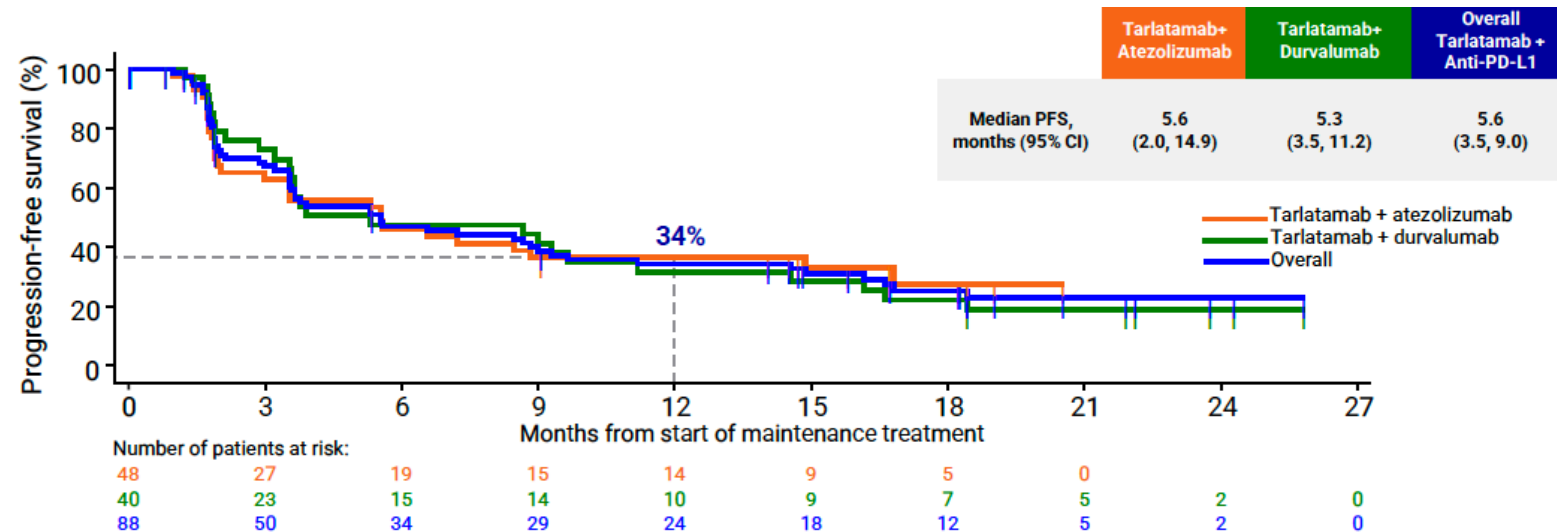
DESDE  
INDUCCIÓN

(Cohortes 2,4,7)



DESDE  
MANTENIMIENTO

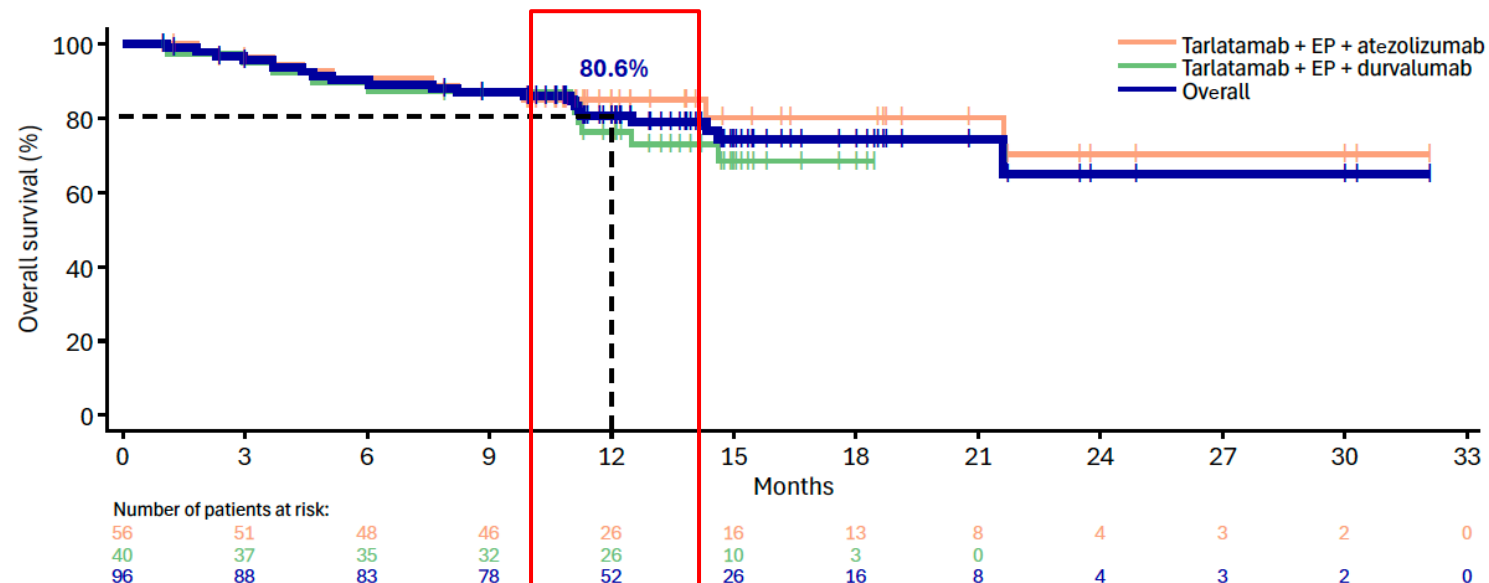
(Cohortes 5,6,8)



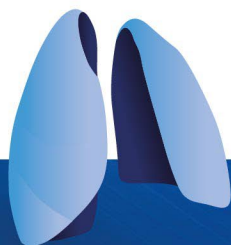
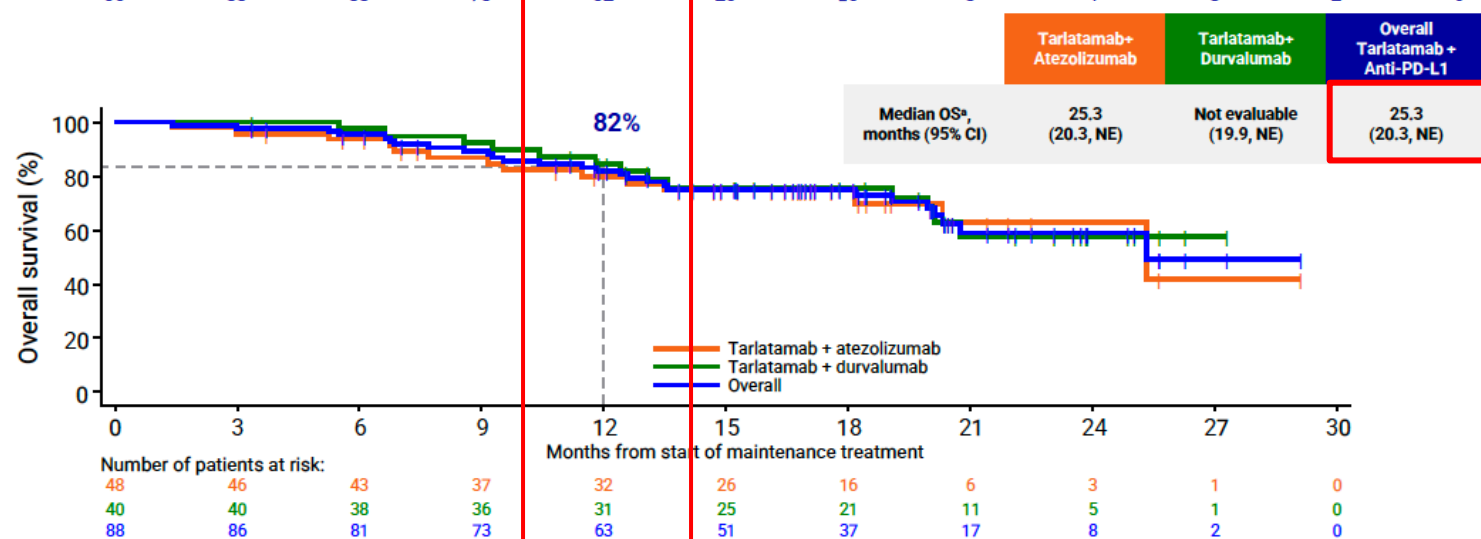


# TARLATAMAB EN 1L - Ph1b: DeLLphi-303

**DESDE  
INDUCCIÓN  
(Cohortes 2,4,7)**



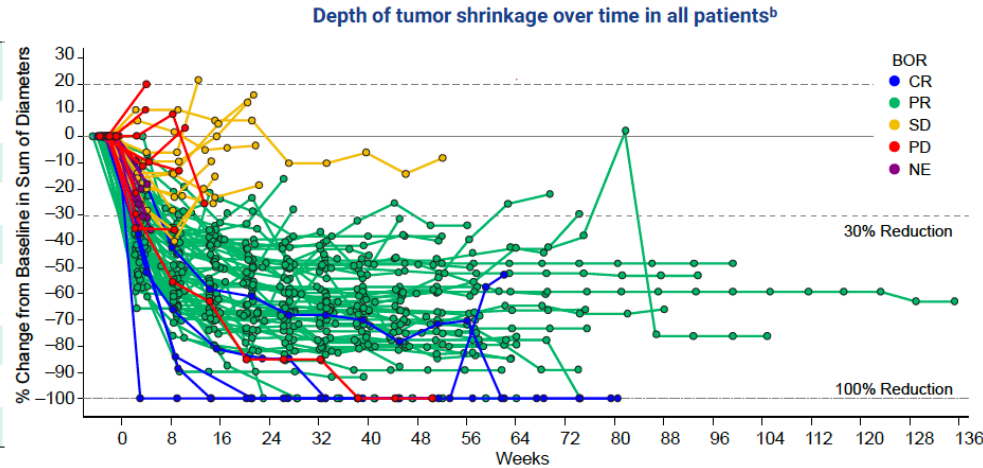
**DESDE  
MANTENIMIENTO  
(Cohortes 5,6,8)**



# TARLATAMAB EN 1L - Ph1b: DeLLphi-303

Overall N = 96	
ORR <sup>a</sup> , n (%)	68 (71)
95% CI	61-80
CR	5 (5)
PR	63 (66)
SD	11 (11)
PD	8 (8)
NE/no post-baseline scan	9 (9)
mDOR, mos (95% CI)	11.0 (8.5, NE)
DCR, % (95% CI)	82 (73-89)
mDoDC, mos (95% CI)	10.7 (7.7-18.8)

Median study follow-up: 13.8 months

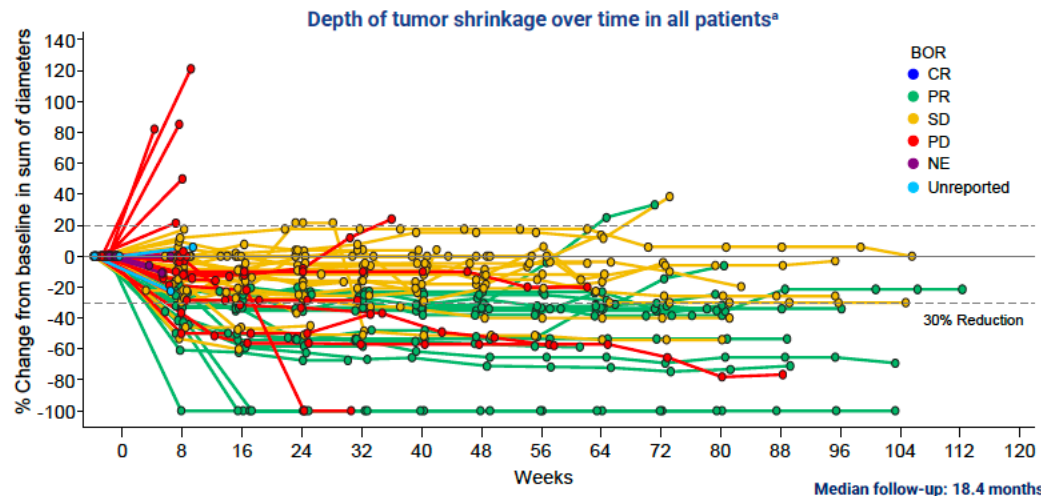


## DESDE INDUCCIÓN (Cohortes 2,4,7)

Control de enfermedad sostenida  $\geq 52$   
semanas: 39%

From a baseline obtained after completion of 1L chemo-IO:

- Overall ORR: 24% (2 CR, 19 PR); median duration of response: 16.6 months (7.1, NE)
- Overall DCR: 60% (2 CR, 19 PR, 32 SD); median duration of disease control: 14.6 months (95% CI 7.2, 18.4)
- 24% of patients remained on treatment at data cutoff and 36% of patients showed sustained DC  $\geq 52$  weeks



Median follow-up: 18.4 months

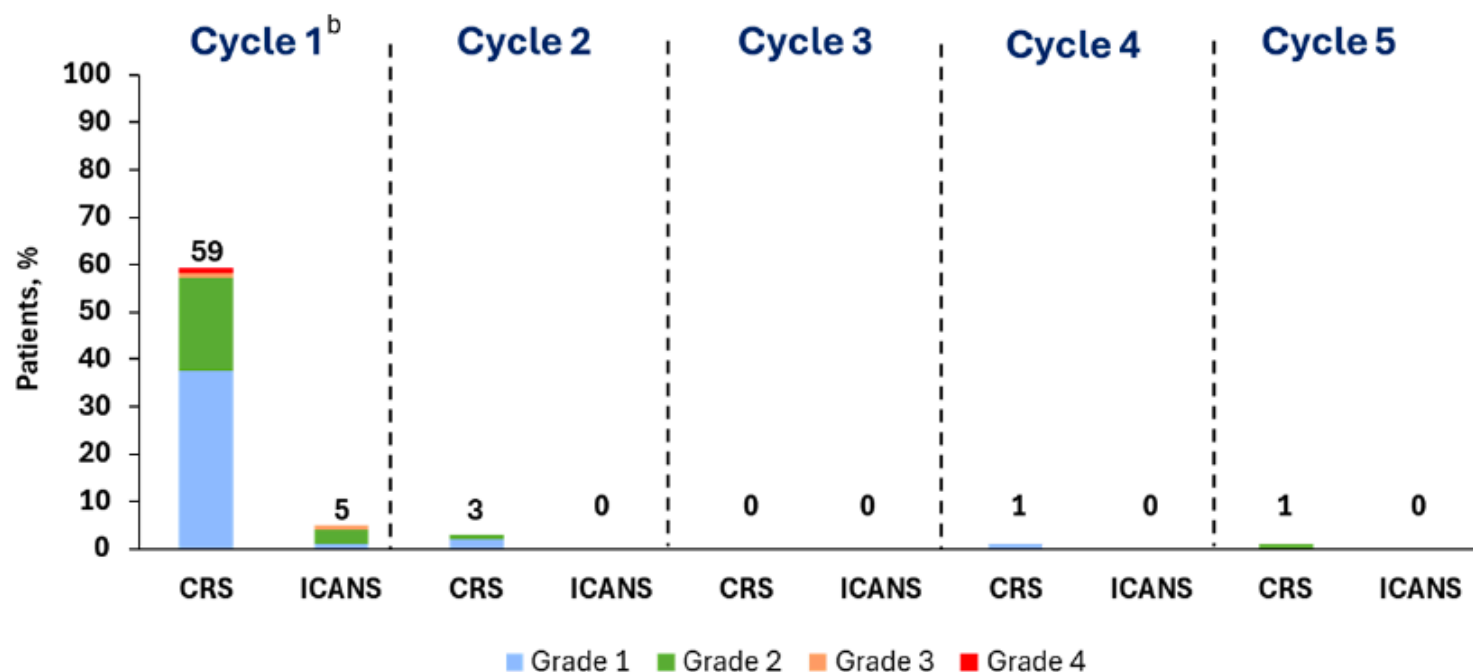
## DESDE MANTENIMIENTO (Cohortes 5,6,8)

Control de enfermedad sostenida  $\geq 52$   
semanas: 36%

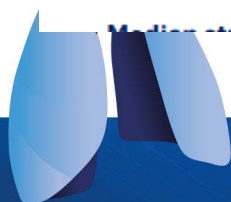
# TARLATAMAB EN 1L - Ph1b: DeLLphi-303

DESDE INDUCCIÓN (Cohortes 2,4,7)

## Treatment-emergent CRS and ICANS by cycle<sup>a</sup>

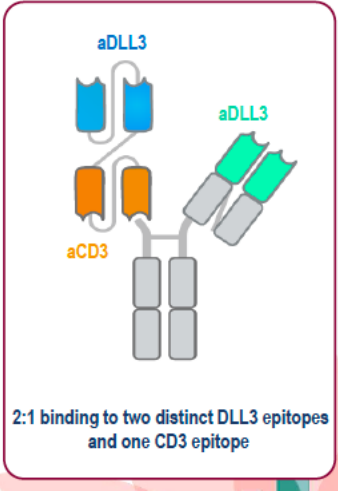


- CRS and ICANS were associated with a low rate of
  - tarlatamab dose interruptions (CRS and ICANS: 1% each)
  - tarlatamab discontinuations (CRS and ICANS: 1% each)
- There were no fatalities related to CRS or ICANS
- The median time to onset of CRS from last prior dose of tarlatamab was 13.3 hours (IQR: 8.0–19.3)
- The median time to onset of ICANS from last prior dose of tarlatamab was 5 days (IQR: 3.0–5.0)

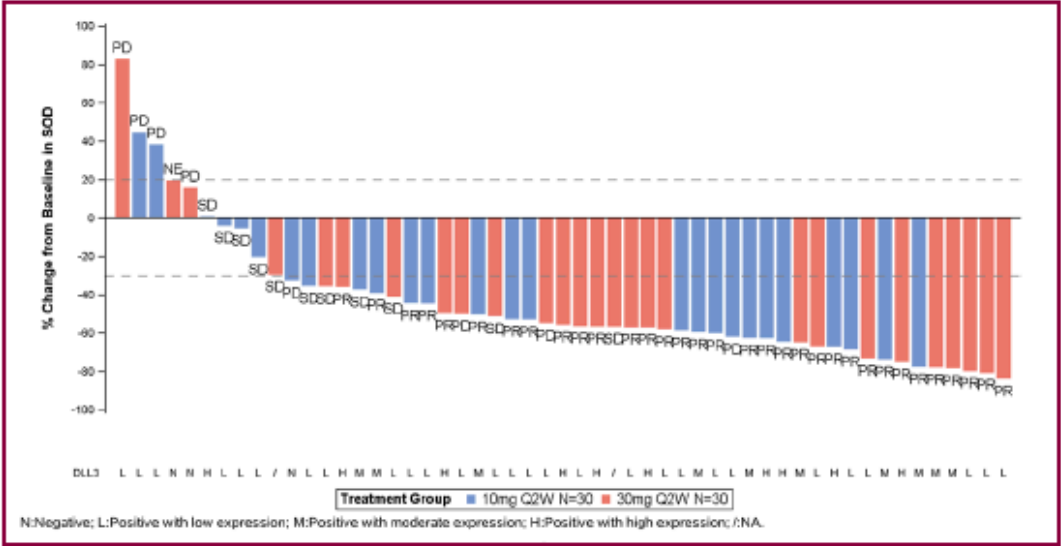


# ALVELTAMIG EN PREVIAMENTE TRATADOS (>2L) - Ph II

(ZG006)



	10mg Q2W (N=30)	30mg Q2W (N=30)
BOR		
PR, n (%)	18 (60.0)	20 (66.7)
SD, n (%)	4 (13.3)	2 (6.7)
PD, n (%)	7 (23.3)	6 (20.0)
NE, n (%)	1 (3.3)	2 (6.7)
uORR, n (%)	18 (60.0)	20 (66.7)
95% CI	(40.60, 77.34)	(47.19, 82.71)
cORR, n (%)	16 (53.3)	17 (56.7)
95% CI	(34.33, 71.66)	(37.43, 74.54)
DCR, n (%)	22 (73.3)	22 (73.3)
95% CI	(54.11, 87.72)	(54.11, 87.72)



	10mg Q2W (N=30)	30mg Q2W (N=30)
PFS (Months)		
Median (95% CI)	7.03 (2.92, NE)	5.59 (2.79, NE)
DoR (Months)		
Median (95% CI)	NR (3.78, NE)	NR (4.60, NE)
9m (95% CI)	61.6 (29.51, 82.46)	55.6 (22.67, 79.30)

OS 12m  
69.1% (10mg); 58.2% (30mg)

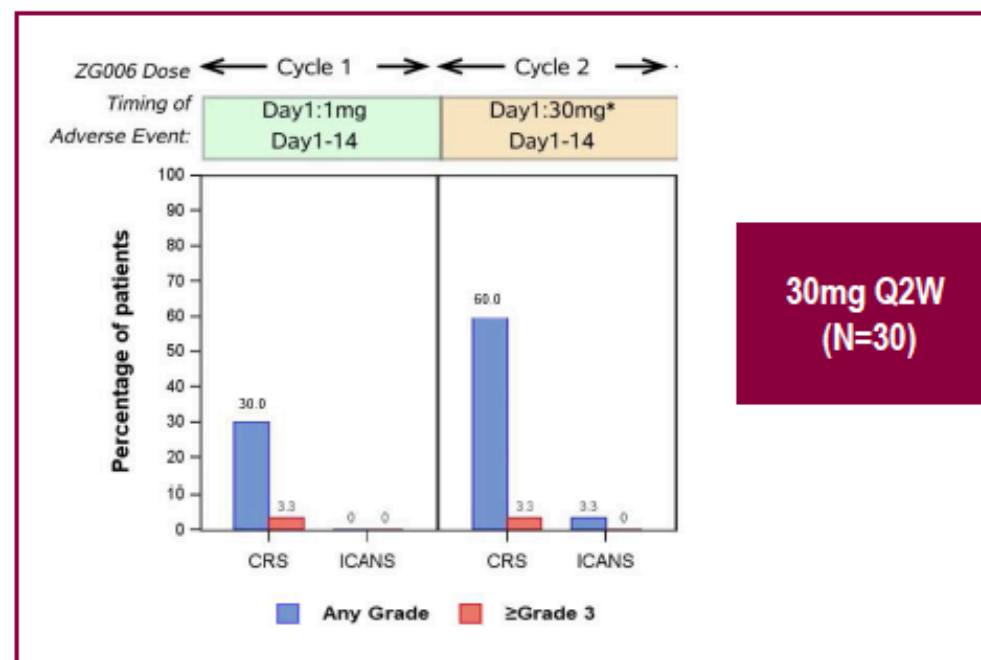
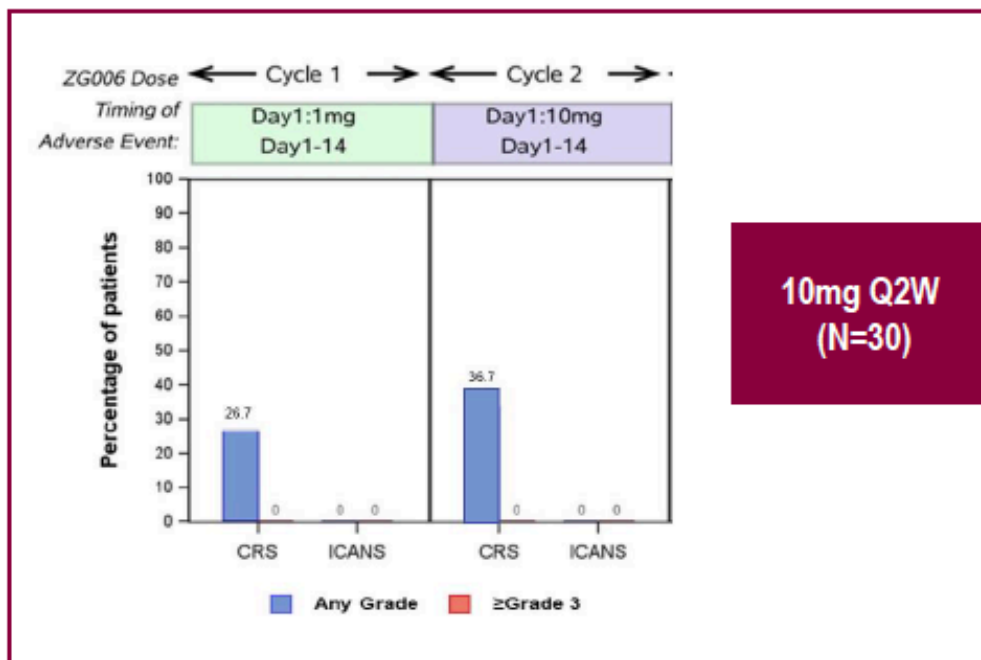




# ALVELTAMIG EN PREVIAMENTE TRATADOS (>2L) - Ph II: DeLLphi-303

## CRS and ICANS on Treatment

- Most CRS events were Grade 1-2 in severity, and resolved quickly after symptomatic treatment
- The incidence of CRS and ICANS according to severity by the treatment cycle showed most CRS events occurred during the 1st to 2nd dosing cycle
- Grade  $\geq 3$  CRS occurred in only 2 patients at 30 mg dose group





# ADCs y anti-VEGF en CPCP

# ADCs EN CPCP

DLL3

**Rova-T**  
SHR-4849 (IDE819)

SEZ-6

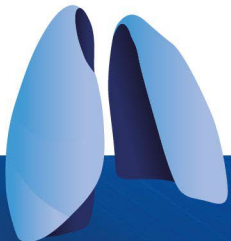
**ABBV-706**  
ABBV-011

B7-H3

HMB088C  
HS-20093 (GSK5764227)  
**I-DXd (DS7300)**  
YL201

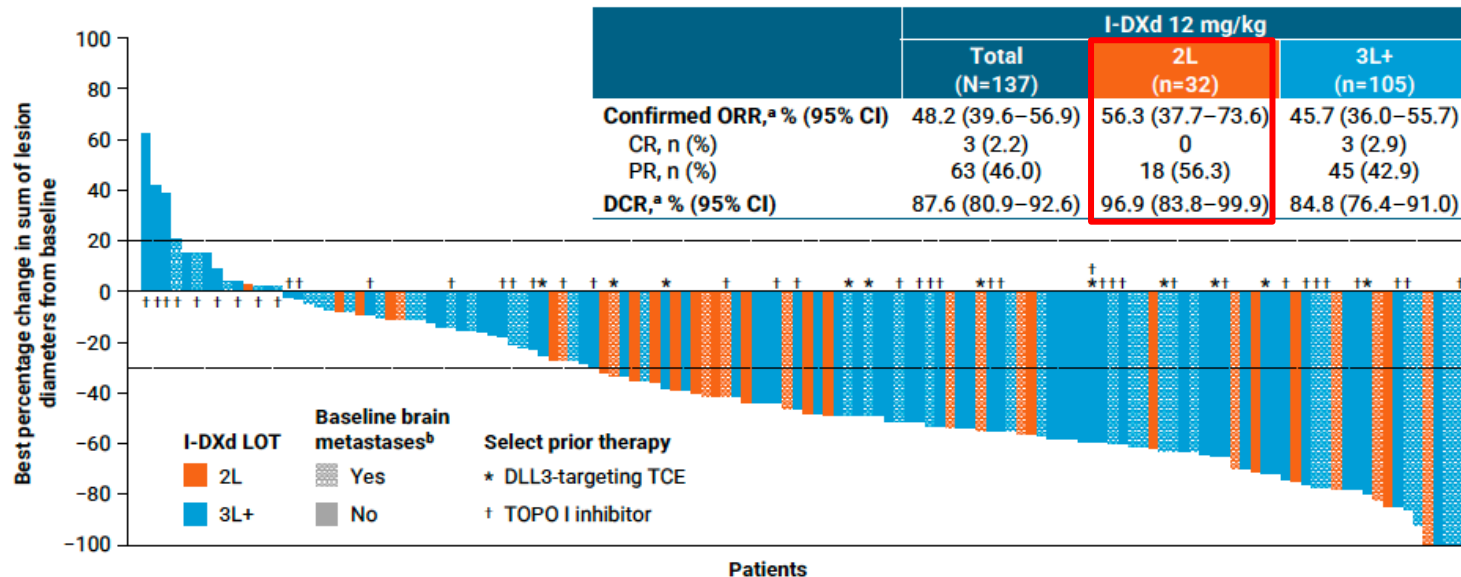
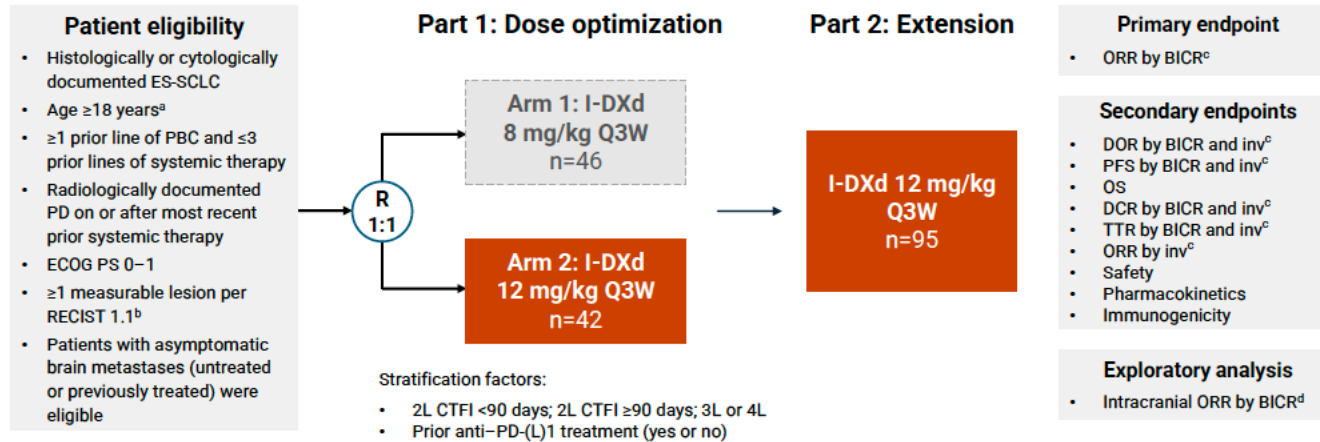
TROP-2

SACI-GOVITECAN  
SHR-A1921



# IFINATAMAB-DXd EN PRETADOS - Ph2: Ideate-Lung01

## ADC anti-B7-H3



ORR CFI  $\leq 30$  d: 11.1%

ORR CFI 30-90 d: 50.0%

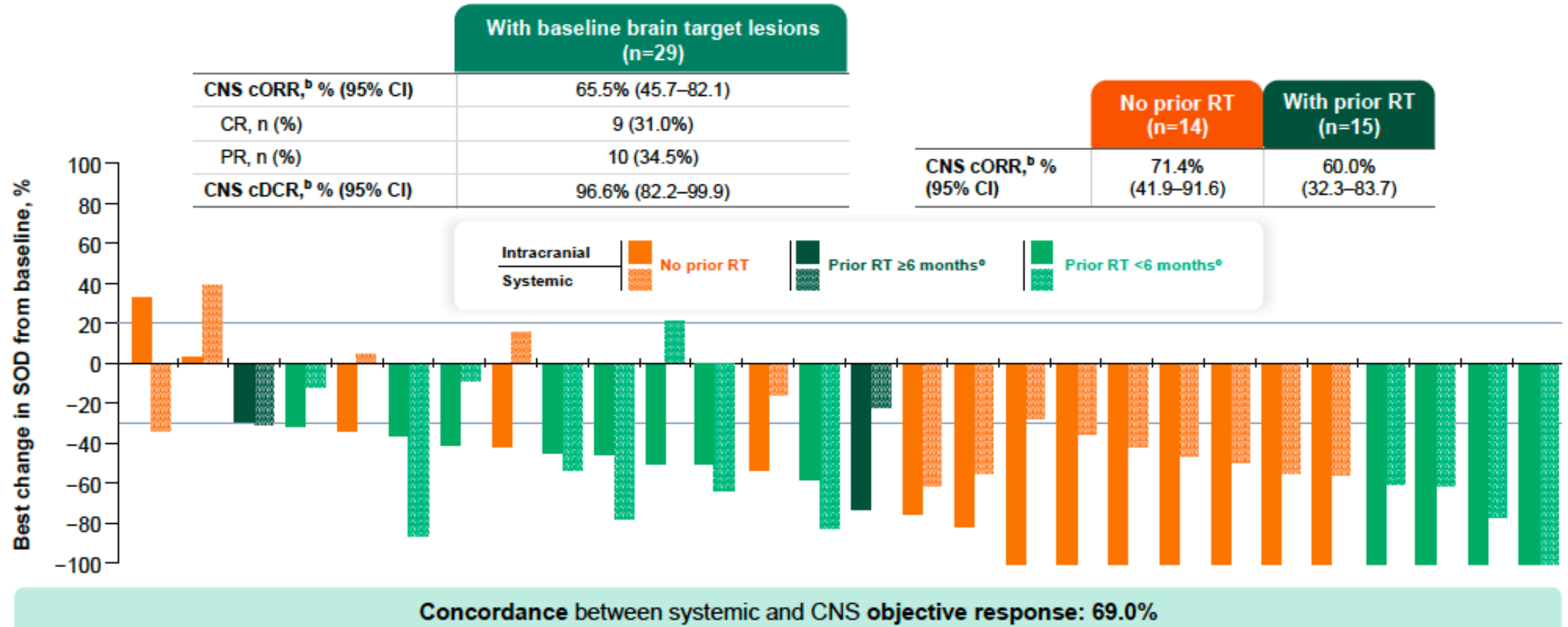
ORR CFI  $\geq 90$  d: 55.6%

DoR: 5.3m (2L: 7.2m)



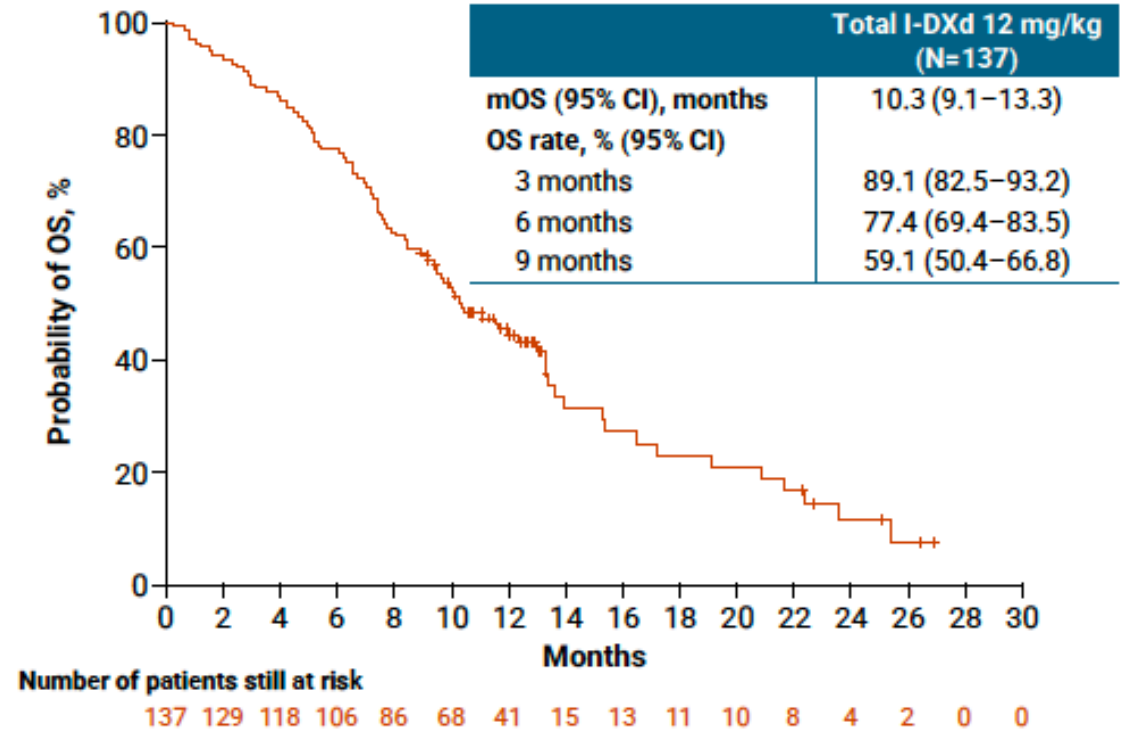
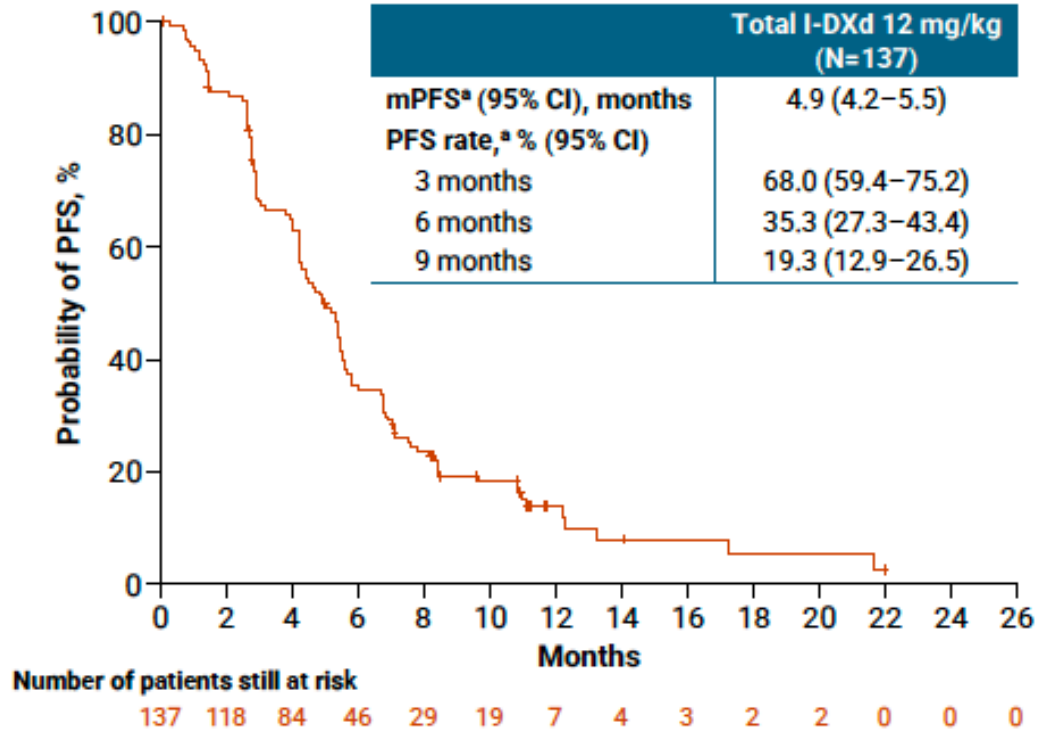
# IFINATAMAB-DXd EN PRETADOS - Ph2: Ideate-Lung01

## ADC anti-B7-H3



# IFINATAMAB-DXd EN PRETADOS - Ph2: Ideate-Lung01

## ADC anti-B7-H3



**I-DXd 12mg/Kg 2L (n=32)**

mPFS: 5.6m

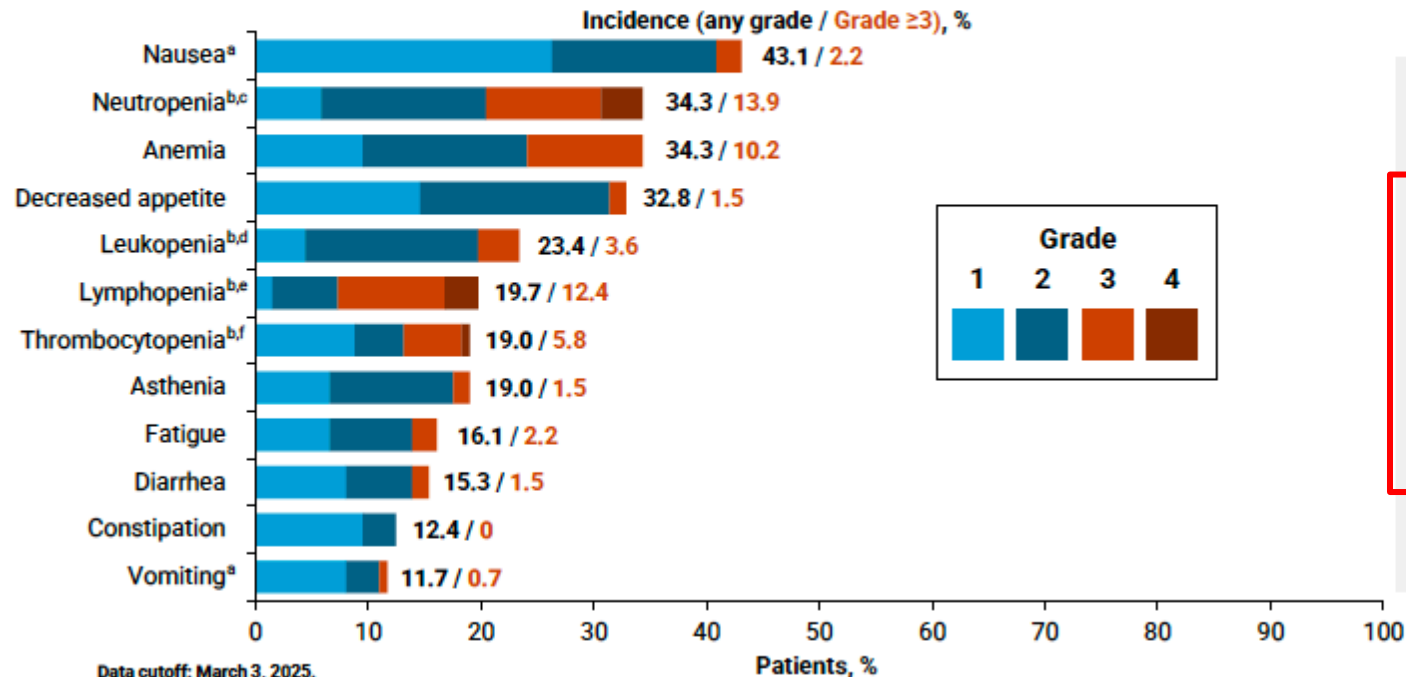
mOS: 12.0m

# IFINATAMAB-DXd EN PRETADOS - Ph2: Ideate-Lung01

## ADC anti-B7-H3

### The most common TRAEs were hematologic or gastrointestinal in nature, and fatigue

TRAEs reported in  $\geq 10\%$  of patients in the total I-DXd 12-mg/kg group (N=137)



- Among the most common TRAEs, the majority were Grade 1 or 2

- Adjudicated treatment-related ILD/pneumonitis was reported in 17 (12.4%) patients:
  - Grade 1 or 2, n=11 (8.0%)
  - Grade 3, n=4 (2.9%)
  - Grade 5, n=2 (1.5%)<sup>g</sup>

- No ILD events were pending adjudication at data cutoff

Data cutoff: March 3, 2025.

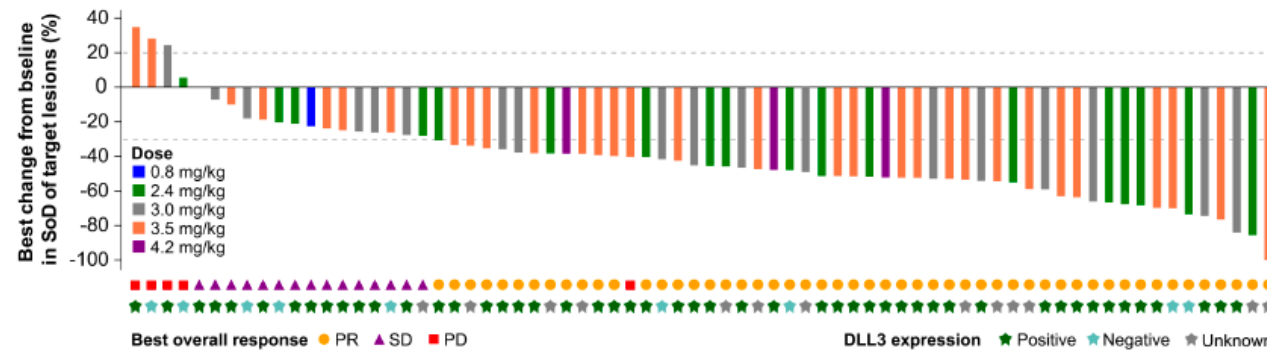
<sup>a</sup>Prior to each I-DXd dose, antiemetic premedication with a 2- or 3-drug combination was mandatory across both study parts. <sup>b</sup>For prophylaxis or treatment of hematologic toxicity, trilaciclib, hematopoietic growth factors, or transfusion of blood, red blood cells, and platelets could be administered. <sup>c</sup>Includes the preferred terms "neutrophil count decreased" and "neutropenia." <sup>d</sup>Includes the preferred terms "white blood cell count decreased" and "leukopenia." <sup>e</sup>Includes the preferred terms "lymphocyte count decreased" and "lymphopenia." <sup>f</sup>Includes the preferred terms "platelet count decreased" and "thrombocytopenia." <sup>g</sup>Both patients were deemed to have adjudicated Grade 5 treatment-related ILD by the ILD adjudication committee; however, only 1 of these patients also had treatment-related ILD associated with death per investigator. ILD, interstitial lung disease; TRAE, treatment-related adverse event.

# SHR-4849 (IDE849) en pretratados - Ph1 first-in-human

## ADC anti-DLL3

Líneas de tratamiento previas (%):  
1L 51%, 2L 33.0%, 3L 15.0%

M1 cerebrales: 24.0%

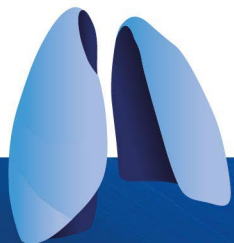


ORR en M1 SNC: 83.3%

DCR en M1 SNC: 100%

PFS 6m: **59% 2L**; 55.3% Todos

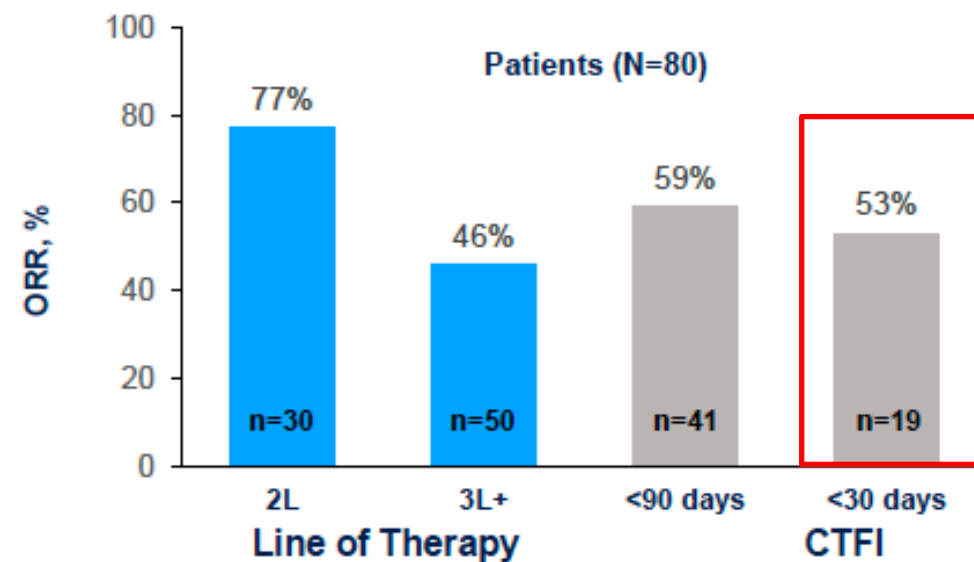
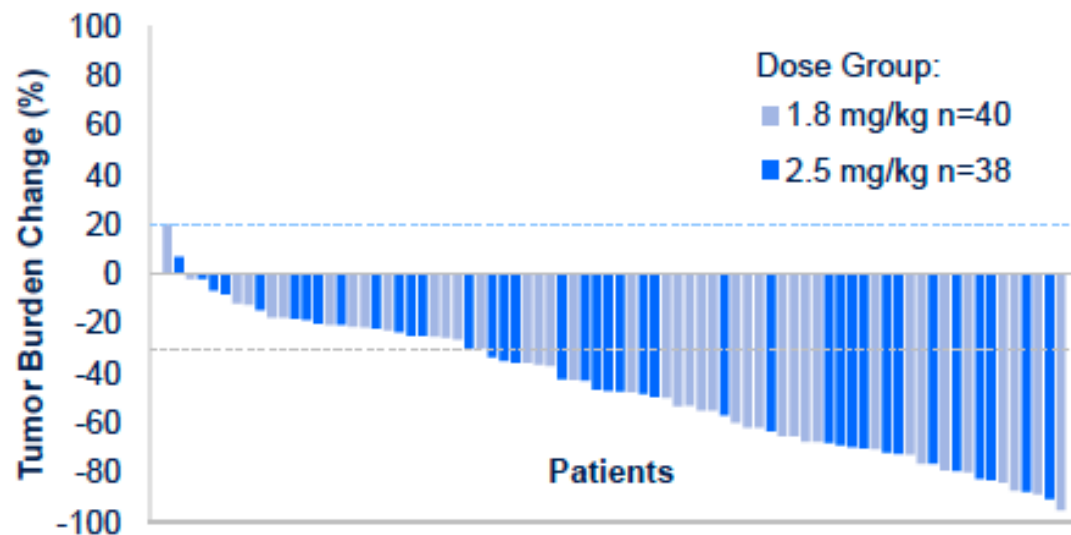
	Total ( $\geq 2.4$ mg/kg)	
	2L Setting (n=35)	All (n=71)
<b>ORR, n (%; 95% CI)</b>	27 (77.1%; 59.9-89.6)	52 (73.2%; 61.4-83.1)
<b>Confirmed ORR, n (%; 95% CI)</b>	21 (60.0%; 42.1-76.1)	34 (47.9%; 35.9-60.1)
<b>Response pending confirmation, n (%)</b>	4 (11.4%)	10 (14.1%)
<b>DCR, n (%; 95% CI)</b>	34 (97.1%; 85.1-99.9)	66 (93.0%; 84.3-97.7)





# ABBV-706 en pretratados - Ph1 M2-385

## ADC anti-SEZ6 (*Seizure-related homolog protein 6*)



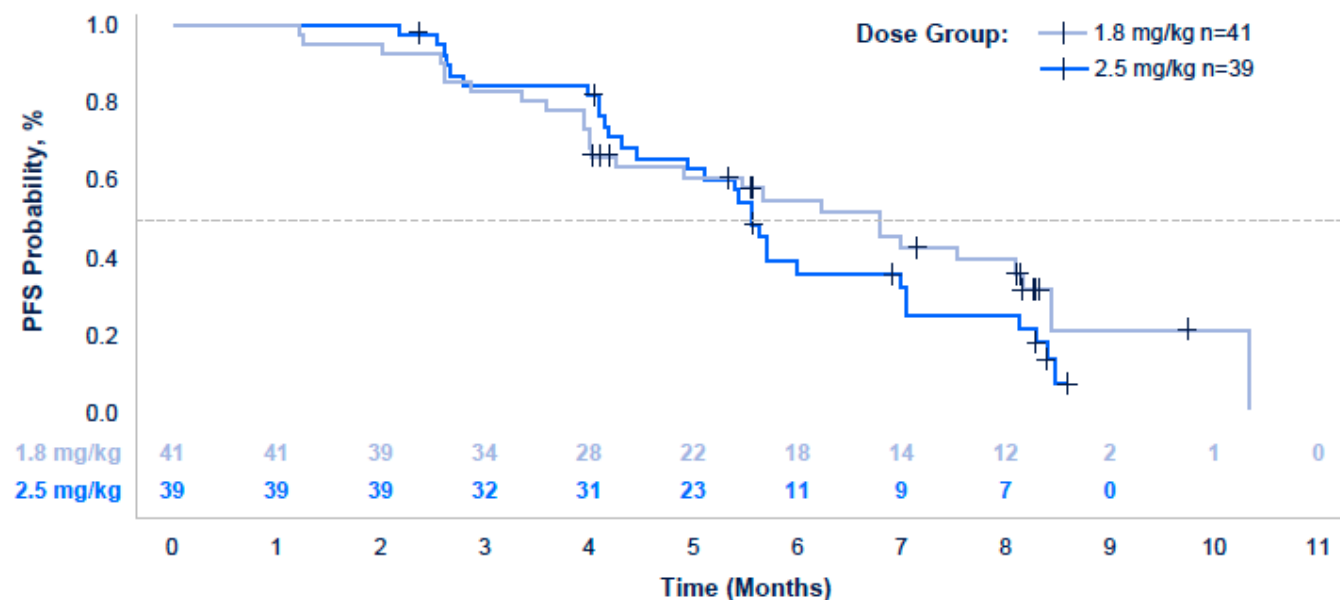
	1.8 mg/kg (n=41)	2.5 mg/kg (n=39)	Total (N=80)
<b>Median (95% CI) DOR, months</b>	<b>6.2 (4.2–NE)</b>	<b>4.4 (3.5–6.9)</b>	<b>5.6 (4.2–6.9)</b>
2L	6.9 (3.0–NE)	5.2 (2.7–NE)	6.9 (3.2–7.1)
CTFI <30 days	6.2 (2.8–NE)	5.0 (3.2–7.0)	5.7 (2.8–7.5)

# ABBV-706 en pretratados - Ph1 M2-385

## ADC anti-SEZ6 (*Seizure-related homolog protein 6*)

**PFS was longer at 1.8 mg/kg trending toward further improvement in 2L**

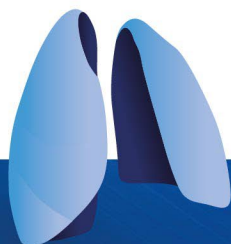
**Progression-Free Survival**



	1.8 mg/kg (n=41)	2.5 mg/kg (n=39)	Total (N=80)
<b>Median (95% CI) PFS, months</b>	<b>6.8 (4.0–8.2)</b>	5.6 (4.4–7.0)	5.7 (4.9–7.0)
2L	7.5 (4.0–8.4)	5.4 (2.8–NE)	6.8 (4.4–8.4)
CTFI <30 days	7.0 (2.0–NE)	4.4 (2.2–7.0)	5.7 (3.9–7.5)
<b>9-month OS, probability (95% CI)</b>	<b>0.6 (0.4–0.7)</b>	0.6 (0.4–0.7)	0.6 (0.5–0.7)

Encouraging PFS observed in CTFI <30 days subgroup

OS data remain immature at time of reporting



# ADCs EN CPCP PREVIAMENTE TRATADOS

	<b>ABBV-706</b> <b>1.8mg/kg</b>  <b>(N=41)</b>	<b>QLC5508</b> <b>(MHB088C)</b> <b>1.6-2.4mg/kg</b> <b>(N=103)</b>	<b>I-DXd</b> <b>12mg/kg</b>  <b>(N=137)</b>	<b>HS20093</b> <b>(GSK5764227)</b> <b>8mg/kg</b> <b>(N=31)<sup>1</sup></b>	<b>YL201</b> <b>1.6-2.8mg/kg</b>  <b>(N=72)<sup>2</sup></b>	<b>Sacituzumab</b> <b>Govitecan</b>  <b>(N=43)<sup>3</sup></b>
Target	SEZ6	B7-H3	B7-H3	B7-H3	B7-H3	TROP2
Payload	Top1 inhibitor	Top1 inhibitor (SuperTopoi™)	Top1 inhibitor (DXd)	Top1 inhibitor (HS-9265)	Top1 inhibitor	Top1 inhibitor (SN-38)
DAR	6	4	4	4	8	7.6
Linker	Cleavable	Cleavable	Cleavable	Cleavable	Cleavable	Cleavable
ORR/DCR	56%	36.9/90.3%	48.2/87.6%	61.3/80.6%	63.9/91.7%	41.9/83.7%
mPFS/OS	6.8 months/ 60%(9month OS)	5.72/11.50 months	4.9/10.3 months	5.9/9.8 months	6.3/- months	4.4/13.6 months
Main AEs	Hematological toxicity	Hematological toxicity	Hematological toxicity/GI toxicity	Hematological toxicity	Hematological toxicity	Diarrhea/Hematol ogical toxicity

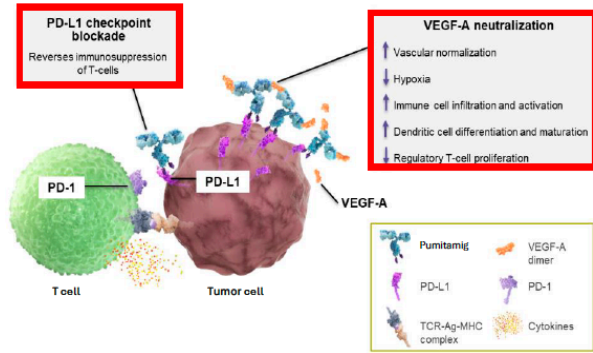
1) WCLC 2024, 2) Ma Y, et al. Nat Med. 2025 Jun;31(6):1949-1957. 3) Dowlati A et al. J Thorac Oncol. 2025 Jun;20(6):799-808.



# PUMITAMIG (BNT327) en pretratados - Ph2 BNT327-01

## Biespecífico anti-PD-L1 y anti-VEGF

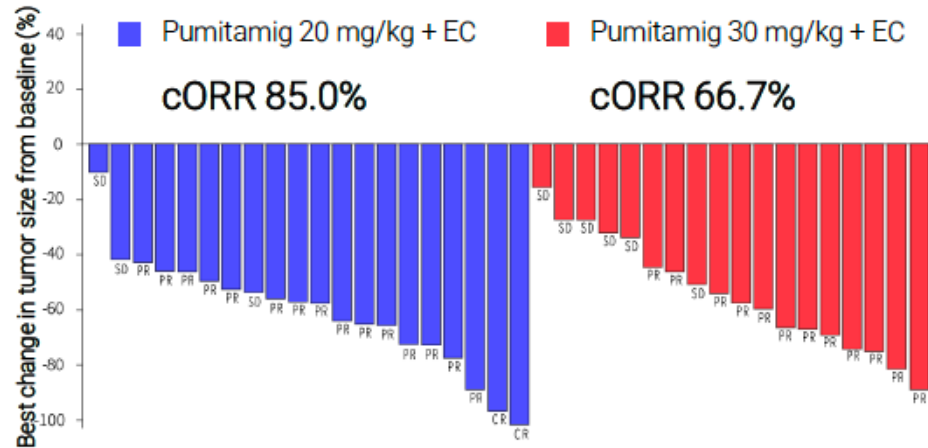
### Pumitamig (BNT327) MOA



### Inclusion criteria:

- Cohort 1**
- Untreated ES-SCLC\*
  - TFI ≥6 months since last CTx/CRTx/RTx for LS-SCLC

1:1 N=40	Arm 1	Pumitamig 20 mg/kg IV Q3W + etoposide 100 mg/m <sup>2</sup> IV + carboplatin AUC5 x 4 cycles	Pumitamig 20 mg/kg IV Q3W
	Arm 2	Pumitamig 30 mg/kg IV Q3W + etoposide 100 mg/m <sup>2</sup> IV + carboplatin AUC5 x 4 cycles	Pumitamig 30 mg/kg IV Q3W

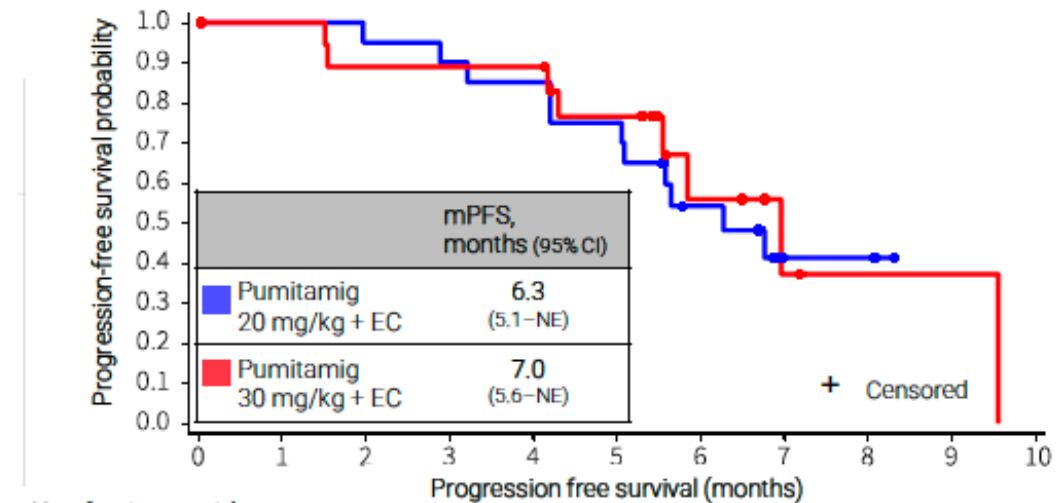


**mDoR: 4.9m**

**mDoR 20mg/Kg: 4.9m**

**mDoR 30mg/Kg: 5.4m**

**mPFS in months (95% CI): 6.8 (5.6–NE) overall.**



	0	1	2	3	4	5	6	7	8	9	10
Pumitamig 20 mg/kg + EC	22	20	19	18	17	15	9	2	2	0	0
Pumitamig 30 mg/kg + EC	21	18	16	16	16	12	5	2	1	1	0

# PUMITAMIG (BNT327) en pretratados - Ph2 BNT327-01

## Biespecífico anti-PD-L1 y anti-VEGF

### Pumitamig-related TEAEs (occurring in $\geq 5\%$ of overall population)

Patients, n (%)	All (N=43)		Pumitamig 20 mg/kg + EC (N=22)		Pumitamig 30 mg/kg + EC (N=21)	
	Any grade	Grade $\geq 3$	Any grade	Grade $\geq 3$	Any grade	Grade $\geq 3$
Any	18 (41.9)	6 (14.0)	9 (40.9)	1 (4.5)	9 (42.9)	5 (23.8)
Nausea	4 (9.3)	0	4 (18.2)	0	0	0
Fatigue	4 (9.3)	0	3 (13.6)	0	1 (4.8)	0
Constipation	3 (7.0)	0	2 (9.1)	0	1 (4.8)	0
Hypertension	3 (7.0)	2 (4.7)	1 (4.5)	0	2 (9.5)	2 (9.5)
Epistaxis	2 (4.7)	0	1 (4.5)	0	1 (4.8)	0
Hemoptysis	2 (4.7)	1 (2.3)	1 (4.5)	1 (4.5)	1 (4.8)	0
Decreased platelet count	2 (4.7)	1 (2.3)	1 (4.5)	0	1 (4.8)	1 (4.8)
Alopecia	2 (4.7)	0	1 (4.5)	0	1 (4.8)	0
Proteinuria	2 (4.7)	1 (2.3)	0	0	2 (9.5)	1 (4.8)

- Pumitamig-related Grade  $\geq 3$  AEs were reported in:

1 patient with 20 mg/kg

- Hemoptysis (n=1)

5 patients with 30 mg/kg

- Hypertension (n=2)
- Decreased platelet count (n=1)
- Proteinuria (n=1)
- Pulmonary embolism (n=1)

- TEAEs leading to discontinuation of pumitamig:

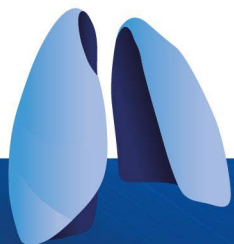
2 patients with 20 mg/kg (9.1%)

- hemoptysis and sepsis; both n=1

4 patients with 30 mg/kg (19.0%)

- hemoptysis, sepsis, pulmonary embolism, and pulmonary hemorrhage; all n=1

- No treatment-related deaths.**





# AVANCES EN 2025 EN CPCP

## POTENCIALES NUEVOS ESTÁNDARES:

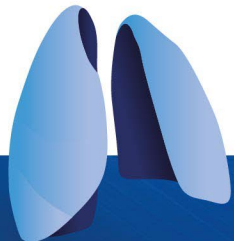
- MANTENIMIENTO CON LURBINECTEDINA-ATEZOLIZUMAB (No datos en pacientes con M1 cerebrales)
- TARLATAMAB EN 2L (Eficacia en enfermedad sensible y resistente y en M1 cerebrales)

## DATOS PROMETEDORES....

- TARLAMATAB EN 1L (Pendiente de resultados de los Ph3 DeLLphi-305 y DeLLphi-312)
- ADCs en 2L y posteriores (anti-B7-H3, anti-DLL3, anti-SEZ6, anti-TROP2)

## EC. EN CURSO

- 2L- Ph3 ADCs vs. SoC: IDeate-Lung02, EVOKE-SCLC-04,...
- 2L - COMBOs ADCs + BiTEs (MK-6070-02, DeLLphi-310); Topotecan + BiTE (DAREON-9)
- 1L - COMBOs ADCs + anti-PD-L1 +/- BiTEs (DeLLphi-310, SEZanne)

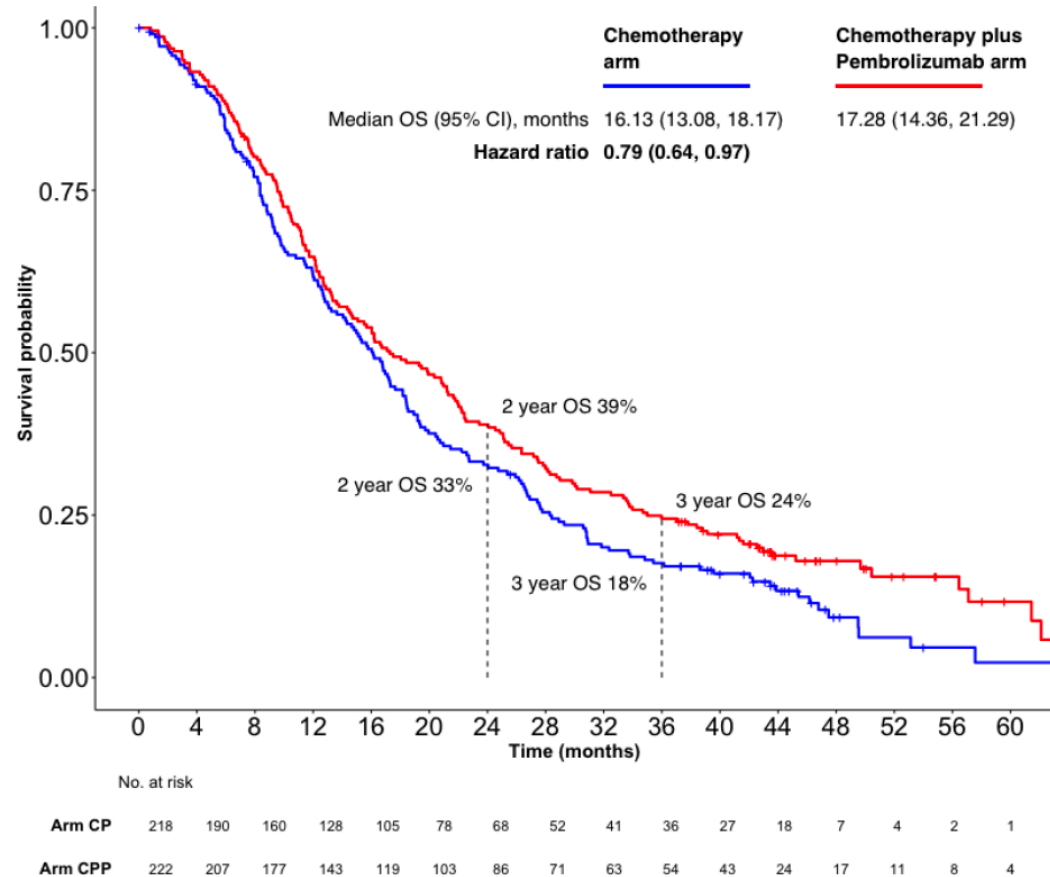




# Mesotelioma

# Update + Análisis exploratorio del CCTG IND227

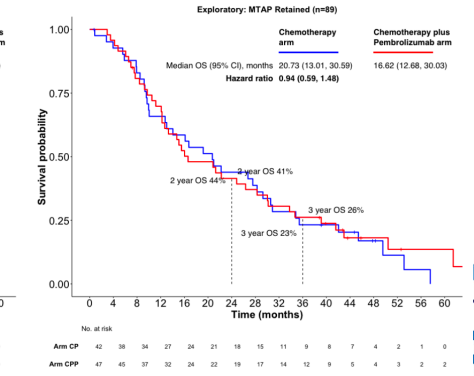
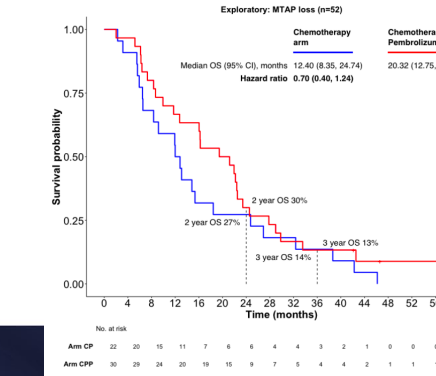
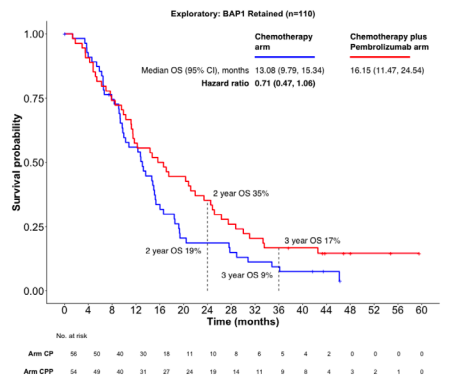
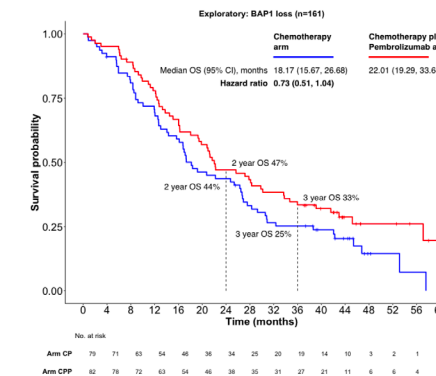
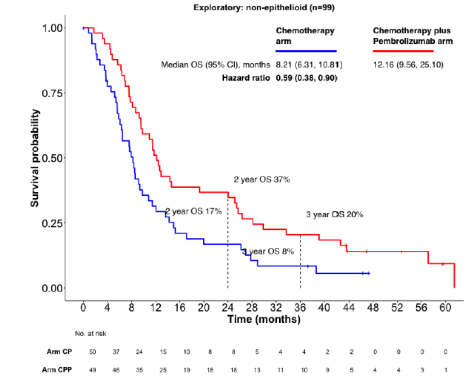
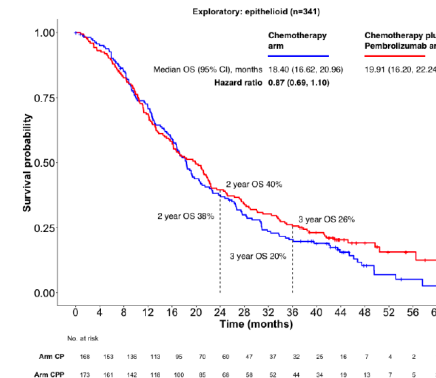
## 1L: QT +/- Pembrolizumab



Histología

BAP1

MTAP





# Tumores epiteliales tímicos

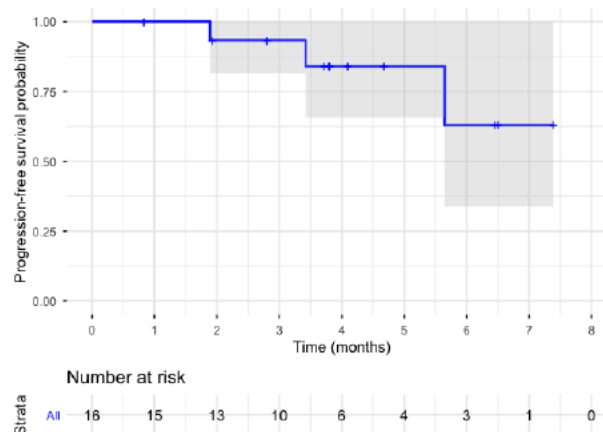
# RIVOCERANIB – Ph2 KCSG LU23-09 (THRIVE)

	Overall cohort (N=40)
<b>Age, years</b>	60 (29-82)
<b>Sex</b>	
Male	25 (62.5%)
Female	15 (37.5%)
<b>ECOG</b>	
0	13 (32.5%)
1	27 (67.5%)
<b>Line of therapy</b>	
2L	14 (35.0%)
3L	9 (22.5%)
4L +	17 (42.5%)
<b>Histological subtype</b>	
Thymoma A	1 (2.5%)
Thymoma B1	3 (7.5%)
Thymoma B2	8 (20.0%)
Thymoma B3	4 (10.0%)
Thymic Carcinoma	24 (60.0%)
<b>Sites of metastases</b>	
Pleura	25 (62.5%)
Lung	20 (50.0%)
Lymph node	14 (35.0%)
Liver	10 (25.0%)
Bone	8 (20.0%)
Pericardium	3 (7.5%)

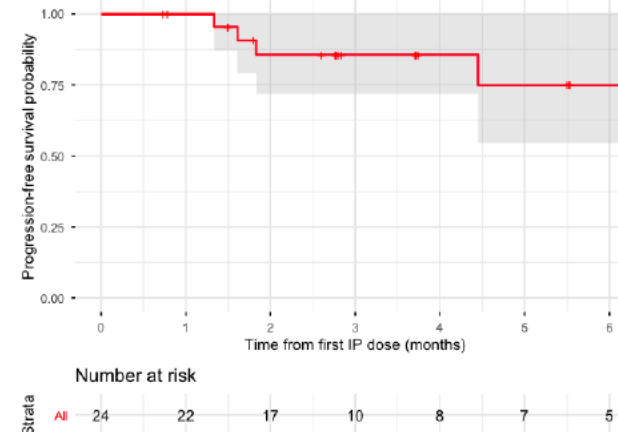
Variables	Total (N=40)	Thymoma (N=16)	Thymic carcinoma (N=24)
<b>Best response (RECIST 1.1)</b>			
Complete response	0 (0.0%)	0 (0.0%)	0 (0.0%)
Partial response	14 (35.0%)	5 (31.2%)	9 (37.5%)
Stable disease	20 (50.0%)	10 (62.5%)	10 (41.7%)
Progressive disease	2 (5.0%)	0 (0.0%)	2 (8.3%)
Not evaluable	4 (10.0%)	1 (6.3%)	3 (12.5%)

ORR 31.2%    ORR 37.5%  
 DCR 93.7%    DCR 79.2%

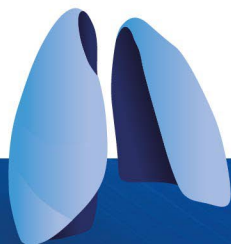
Thymoma



Thymic Carcinoma



mFU 3.7m: PFS NR







# GRACIAS