# *innate* pharma

# Lacutamab KOL event

12 December 2023

EURONEXT : IPH.PA NASDAQ : IPHA

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# Speakers on Today's Call



**Sonia Quaratino** MD, PhD

Chief Medical Officer of Innate Pharma

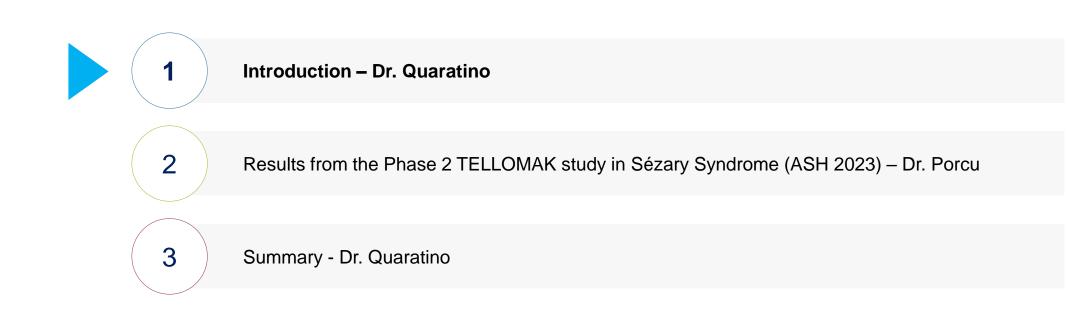


**Pierluigi Porcu** MD

Director, Division of Hematologic Malignancies and Hematopoietic Stem Cell Transplantation, Sidney Kimmel Cancer Center, Jefferson Health, Philadelphia



#### Agenda





# Scientific innovation drives our strategy

#### Our ambition -

leverage our scientific know-how in innate immunity and antibody engineering to develop cancer drugs that improve the lives of patients.

Drive near-term value with lacutamab

Advance our innovative pipeline Build a sustainable business through partnerships

#### **Our robust pipeline of proprietary & partnered assets**

|           | Program  | Target                                       | Indication                            | Pre-Clinical                           | Phase 1            | Phase 2     | Phase 3 | Upcoming Milestone      |
|-----------|--|--|---------------------------------------|--|--------------------|-------------|---------|-------------------------|
|           | Leeutemek  | KIR3DL2                                      | Sezary syndrome / MF                  | TELLOMAK (FDA FAS                      | ST TRACK/EMA PRIME | ESIGNATION) |         | ASH/ Final data H2 2023 |
|           | Lacutamab  | KIK3DL2                                      | PTCL                                  | KILT Phase 2 / Phase                   | 1b                 |             |         | IST with LYSA           |
| rietary   | IPH5301  | CD73   | Cancer (solid tumors)                 | CHANCES                                |                    |             |         | IST with IPC            |
| Prop      | IPH6501<br>Others  | CD20<br>Undisclosed                          |                                       | Phase 1 starting<br>Pre-clinical       |                    |             |         | IPH6501 Phase 1 start   |
|           | IPH45<br>IPH43<br>Other  | Nectin-4<br>MICA/B ADC<br>Undisclosed        |                                       | Pre-clinical                           |                    |             |         |                         |
|           |  |  |                                       |  |                    |             |         |                         |
| Partnered | Monalizumab AstraZeneca  | NKG2A  | Unresectable Stg III NSCLC            | PACIFIC-9                              |                    |             |         | Data readout > 2024     |
|           |  |  | Neoadjuvant NSCLC                     | NeoCOAST-2                             |                    |             |         | Data readout > 2024     |
|           | IPH5201 AstraZeneca  | CD39   | Neoadjuvant NSCLC                     | MATISSE                                |                    |             |         |                         |
|           | SAR443579 / IPH6101<br>SAR'514 / IPH6401<br>IPH62 <b>SONOFI</b><br>2 options | CD123<br>BCMA CONKET<br>B7-H3<br>Undisclosed | R/R AML, B-ALL, HR-MDS<br>R/R MM, LCA | PHASE 1 / 2<br>PHASE 1 / 2<br>Research |                    |             |         | ASH 2023                |
|           | Other Takeda   | Undisclosed ADC                              | Celiac disease                        | Pre-clinical                           |                    |             |         |                         |

ADC: antibody drug conjugate; MF: Mycosis Fungoides ; PTCL: Peripheral T Cell Lymphoma; NSCLC: Non-small cell lung cancer; AML: Acute Myeloid Leukemia; B-ALL: B-cell Acute Lymphoblastic Leukemia; HR-MDS: High Risk-myelodysplasia; MM: Multiple Myeloma; LCA: Light-chain Amyloidosis; IST: investigator-sponsored study; IND: Investigational new drug

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#### **Presentations at the ASH 2023 Annual Meeting**





#### What is presented?

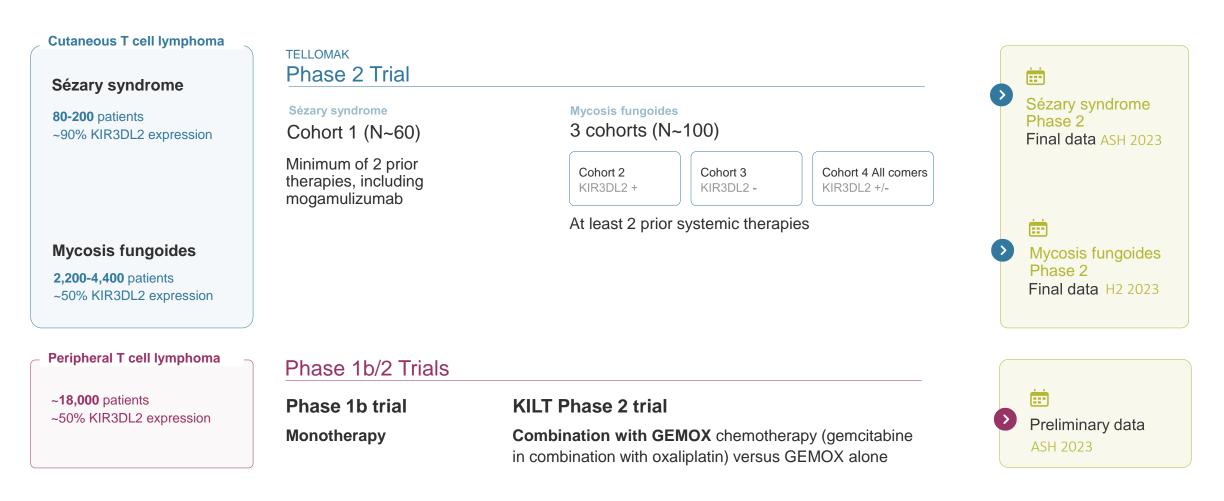
Lacutamab Phase 2 Sézary syndrome

Lacutamab Phase 1b PTCL initial data

SAR443579 (CD123 ANKET) (Sanofi)

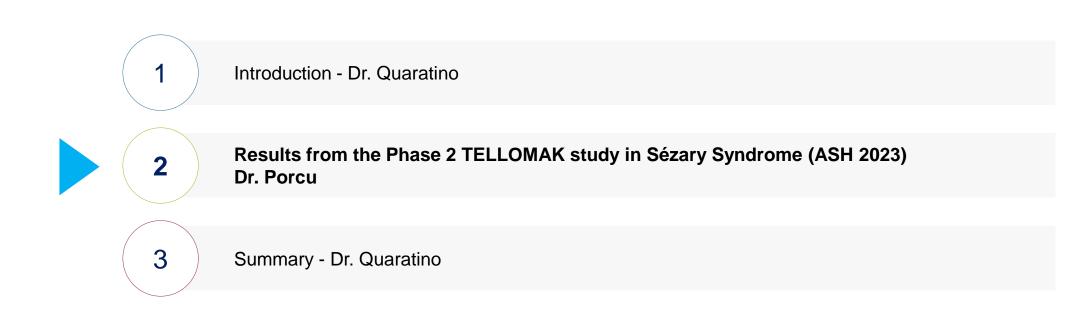
#### Lacutamab | Development

A potential new standard of care in the T-cell lymphomas expressing KIR3DL2



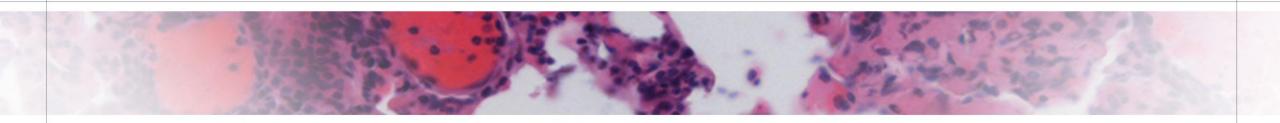


#### Agenda





#### American Society of Hematology Helping hematologists conquer blood diseases worldwide



# Lacutamab in Patients with Relapsed and Refractory Sézary Syndrome: Results from the Tellomak Phase 2 Trial

Publication Number: 185 Submission ID: 173806



# Lacutamab in Patients with Relapsed and Refractory Sézary Syndrome: Results from the Tellomak Phase 2 Trial

Martine Bagot<sup>1</sup>, Youn H. Kim<sup>2</sup>, Larisa J. Geskin<sup>3</sup>, Pablo L. Ortiz-Romero<sup>4</sup>, Ellen Kim<sup>5</sup>, Neha Mehta-Shah<sup>6</sup>, Olivier Dereure<sup>7</sup>, Saskia Oro<sup>8</sup>, Marie Beylot-Barry<sup>9</sup>, Stéphane Dalle<sup>10</sup>, Eric Jacobsen<sup>11</sup>, Frederick Lansigan<sup>12</sup>, Caroline Ram-Wolff<sup>1</sup>, Michael S Khodadoust<sup>2</sup>, Maxime Battistella<sup>13</sup>, Alejandro Gru<sup>14</sup>, Hélène Moins-Teisserenc<sup>15</sup>, Pier Luigi Zinzani<sup>16</sup>, Julien Viotti<sup>17</sup>, Christine Paiva<sup>17</sup>, Marianna Muller<sup>17</sup>, <u>Pierluigi Porcu<sup>18</sup></u>

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Clinicaltrials.gov: NCT03902184



#### Lacutamab

#### **KIR3DL2** targeted treatment

- In development in:
  - Cutaneous T-cell lymphoma (CTCL)
    - Sézary Syndrome (SS) and Mycosis Fungoides (MF)
  - Peripheral T-cell lymphoma (PTCL)
- Phase 1 data in SS patients after ≥ 2 prior systemic therapies<sup>1</sup>:
  - Median Prior Lines of Therapy in SS population: 2 (2-4)
  - Global Objective Response Rate (ORR): 42.9% (95%CI: 28.0-59.1)
  - Median duration of response (DoR):
     13.8 months (95%CI: 7.2-NA)
  - Median progression free survival (PFS):
     11.7 months (95%CI: 8.1-NA)

#### First-in-class humanized anti-KIR3DL2 cytotoxicity-inducing antibody

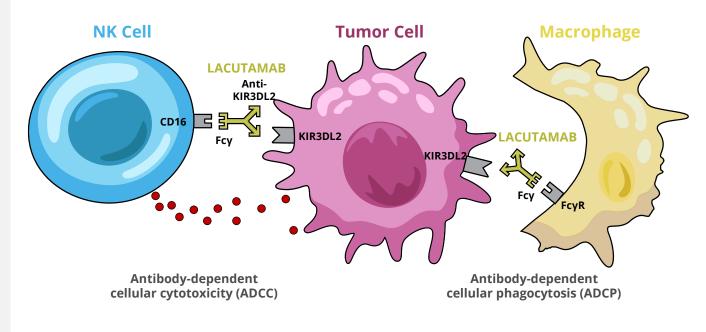


Figure 1: Lacutamab Mechanism of Action

- Orphan drug designation for the treatment of CTCL (EMA and FDA)
- PRIME (EMA) and Fast Track (FDA) designation for SS patients who have been treated by at least 2 prior systemic therapy



#### **TELLOMAK - NCT03902184** Phase 2 Study in Two CTCL Subtypes

#### Sézary Syndrome (N~60) ≥ 2 prior systemic therapies

#### Cohort 1

Sézary Syndrome ≥ 2 prior systemic therapies, Must include mogamulizumab as prior therapy

#### Administration

• Lacutamab is administered every week for 5 weeks then every 2 weeks for 10 administrations then every 4 weeks, by intravenous infusion, until disease progression or unacceptable toxicity

#### **Study Endpoints**

- Primary endpoint: global ORR
- Secondary endpoints: PFS, OS, DoR, quality of life, safety and tolerability, PK & immunogenicity

#### **Key Eligibility Criteria**

- Relapsed and/or refractory stage IVA, IVB SS (B2 blood in screening)
- No evidence of large cell transformation (LCT), based on central histologic evaluation at screening

#### Mycosis Fungoides (N~100) ≥ 2 prior systemic therapies

| <u>Cohort 2</u> | Cohort 3      | All Comers   |
|-----------------|---------------|--------------|
| KIR3DL2 ≥ 1%    | KIR3DL2 <1%   | KIR3DL2 ≥ 1% |
| Simon 2 Stage   | Simon 2 Stage | or <1%       |



#### **TELLOMAK - NCT03902184**

#### Patient baseline characteristics in SS patients (n=56)

| Patient Characteristics   | Cohort 1 N=56                                  |
|---|--|
| Age in years, Median (range)  | 69 (42-86)                                     |
| <ul> <li>Female, N (%)</li> <li>Male, N (%)</li> </ul>  | 22 (39.3)<br>34 (60.7)                         |
| <ul> <li>Stage of the disease at screening, N (%)</li> <li>Stage IVA1</li> <li>Stage IVA2</li> <li>Stage IVB</li> </ul> | 36 (64.3)<br>19 (33.9)<br>1 (1.8)              |
| B2 blood involvement at screening, N (%)  | 56 (100.0)                                     |
| Nodal involvement at screening, N* (%)<br>- N2<br>- N3 involvement<br>- Nx  | 6 (10.7)<br>20 (35.7)<br>14 (25.0)             |
| T4 confluence of erythema covering ≥ 80% BSA  | 38 (67.9%)                                     |
| N prior systemic lines, Median (range)<br>- 2, N (%)<br>- 3-4, N (%)<br>- > 4, N (%)                                    | 5 (2–15)<br>7 (12.5)<br>15 (26.8)<br>34 (60.7) |
| Follow-up (months), Median (95% CI)   | 14.4 (9.0-18.4)                                |

\*Nodal involvement at baseline: N2, N3 or Nx



#### **TELLOMAK - NCT03902184** Efficacy results in SS patients (*N*=56)

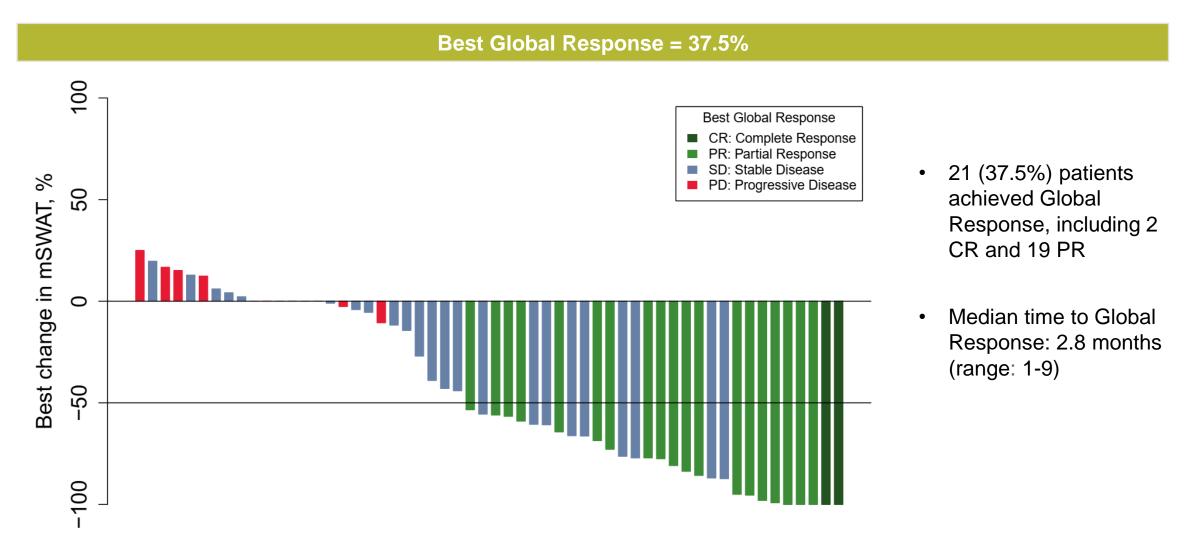
|                      | Best<br>Global<br>Response<br>N=56 | Best<br>Response<br>in Skin<br>N=56 | Best<br>Response<br>in Blood<br>N=56 | Best<br>Response<br>in LN<br>N=46* |
|----------------------|------------------------------------|-------------------------------------|--------------------------------------|------------------------------------|
| Best Response, N (%) |                                    |                                     |                                      |                                    |
| CR                   | 2 (3.6)                            | 5 (8.9)                             | 15 (26.8)                            | 3 (6.5)                            |
| PR                   | 19 (33.9)                          | 21 (37.5)                           | 12 (21.4)                            | 6 (13.0)                           |
| SD                   | 28 (50.0)                          | 27 (48.2)                           | 24 (42.9)                            | 28 (60.9)                          |
| PD                   | 7 (12.5)                           | 3 (5.4)                             | 5 (8.9)                              | 5 (10.9)                           |
| NE                   | 0                                  | 0                                   | 0                                    | 4 (8.7)                            |
| ORR%<br>[95%CI]      | <b>37.5%</b><br>[26.0-50.6]        | <b>46.4%</b><br>[34.0-59.3]         | <b>48.2%</b><br>[35.7-61.0]          | <b>19.6%</b><br>[10.7-33.2]        |

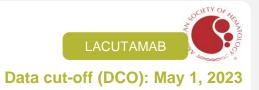
#### **Global Clinical Benefit Rate** (CR+PR+SD): **87.5%** (95% CI 76.4-93.8)

*CR: complete response; PR: partial response; SD: Stable Disease; PD progressive disease; NE: not evaluable; LN lymph nodes* \**includes patients not involved at baseline who progressed in the LN* 

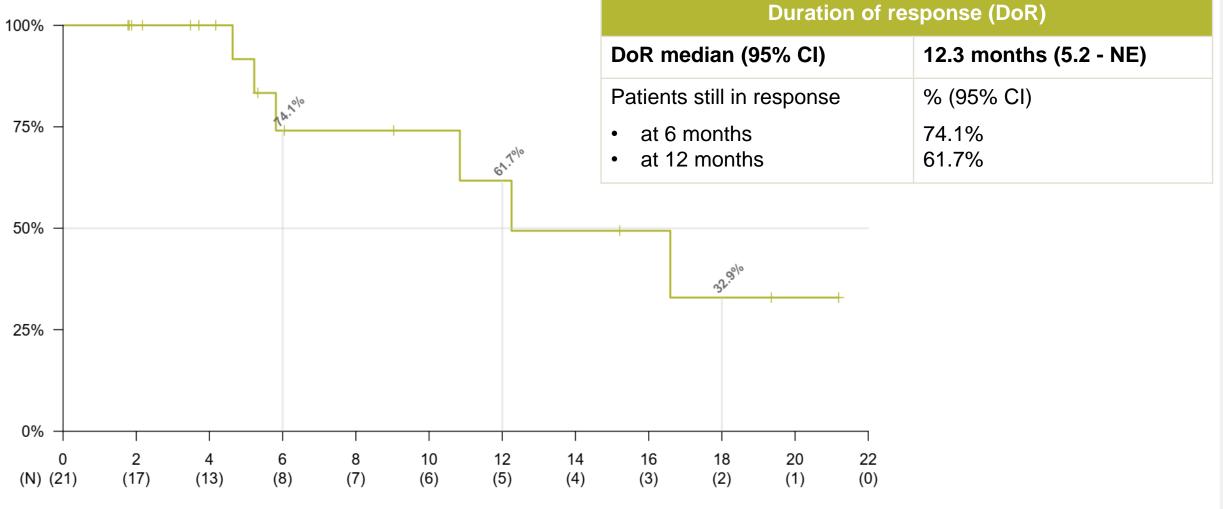


#### **TELLOMAK - NCT03902184** Best Global Response (*N*=56)





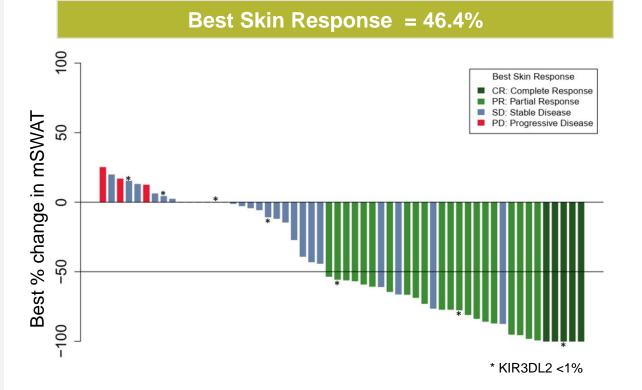
#### **TELLOMAK - NCT03902184** Duration of Response (*N*=56)



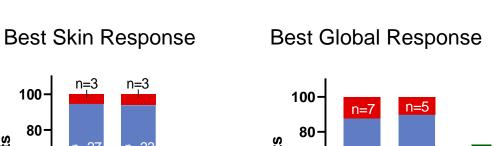


### TELLOMAK - NCT03902184

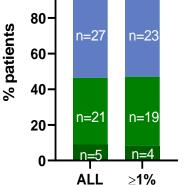
Best Skin response, Overall and According to KIR3DL2 status in skin (N=56)

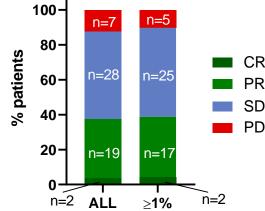


- 26 (46.4%) patients achieved a Skin Response;
   5 CR & 21 PR
- Median time to Skin Response: 2.8 months (range: 1-10)

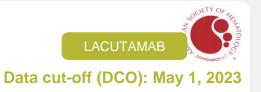


Best Skin/Global Response by KIR3DL2 status





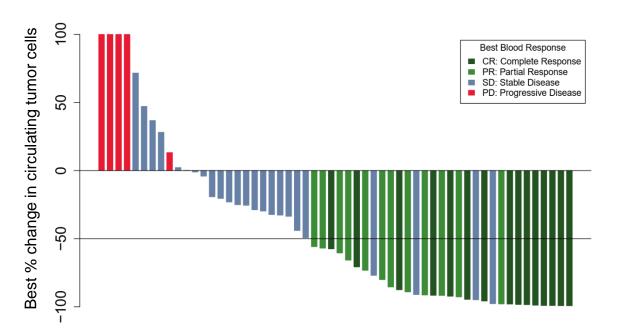
- With a threshold of 1%, KIR3DL2 is expressed in the skin of 87.5% of patients (49/56)
- Response in ≥1% subgroup is consistent with the overall population



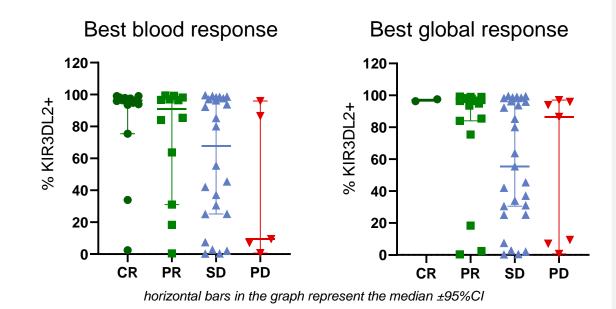
#### **TELLOMAK - NCT03902184**

Best Blood response & according to frequency of KIR3DL2+ circulating tumor cells (N=56)

#### Best Blood Response = 48.2%



- 27 (48.2%) patients achieved a Blood Response; 15 CR & 12 PR
- Median Time to Blood Response: 1.0 month (range 1-6)
- Note 1 unconfirmed CR confirmed after DCO

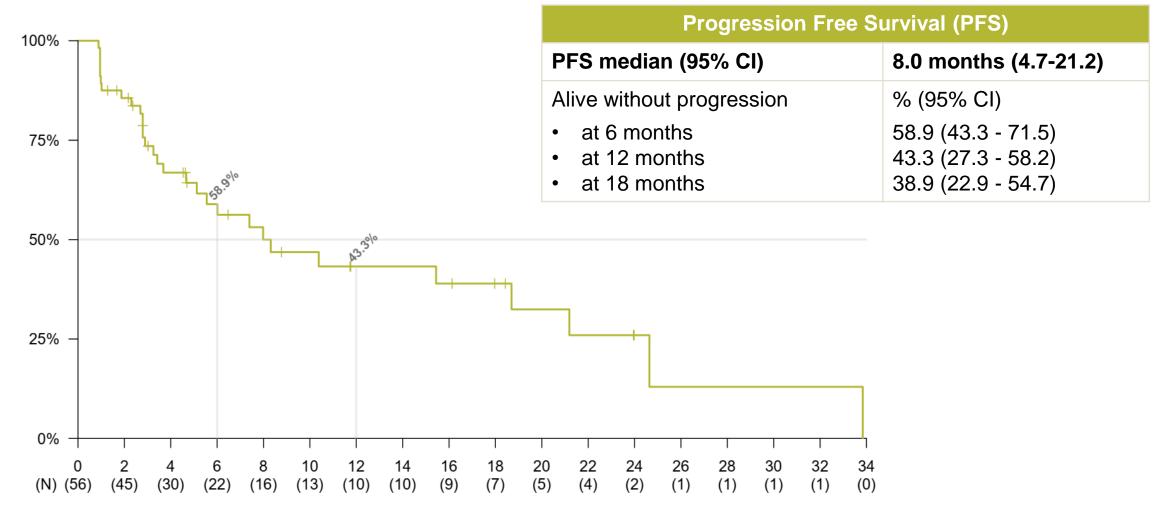


**Best Blood/Global Response by KIR3DL2 status** 

- KIR3DL2 expression on circulating tumor cells (CTCs) was found in all patients
- Median frequency of KIR3DL2+ CTCs: 92.3% [0.2-99.6]



#### **TELLOMAK - NCT03902184** Efficacy results in SS patients: PFS





#### **TELLOMAK - NCT03902184** Treatment Emergent related Adverse Events<sup>1</sup>

|  |   | Total N= 56<br>N (%) |
|--|---|----------------------|
| Any treatment-emergent AEs (TE                 | AEs)  | 54 (96.4)            |
| Any lacutamab-related TEAEs                    |   | 32 (57.1)            |
|  | <ul> <li>General disorders and administration site conditions*</li> </ul> | 15 (26.8)            |
| Nost frequent (>10%)<br>acutamab-related TEAEs | Skin and subcutaneous tissue disorders                                    | 7 (12.5)             |
|  | Gastrointestinal disorders  | 6 (10.7)             |
|  | Investigations  | 6 (10.7)             |
| Any Serious TEAEs                              | 13 (23.2)   |                      |
| Any Serious lacutamab-related TI               | 4 (7.1)   |                      |
| Any Grade <sup>2</sup> 3/4/5 lacutamab-relat   | 10 (17.9)   |                      |
| Any lacutamab-related TEAEs lea                | 3 (5.4)   |                      |
| Any death due to AEs***                        |   | 3 (5.4)              |
| Any death due to lacutamab-related AEs         |   | 0 (0)                |
|  |   |                      |

\* Fatigue 7 (12.5%), Asthenia 4 (7.1%), Peripheral edema 3 (5.4%); \*\* Toxic skin eruption, Skin fissures, Pruritus and AST elevation; \*\*\* Sepsis, Acute respiratory failure, Infection, Grade 5 all Not related to lacutamab. Of note, post DCO, one patient died with transformed cell lymphoma/HLH.

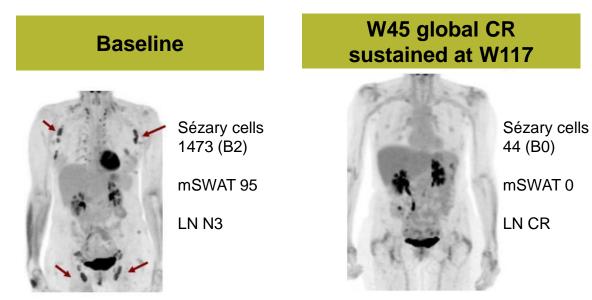
1. Event / as defined by the treating investigator 2. NCI Common Terminology Criteria for Adverse Events (CTCAE)



### TELLOMAK - NCT03902184

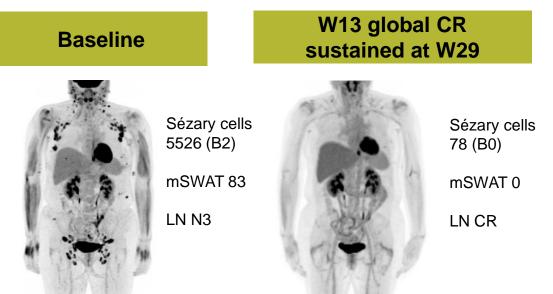
#### Patient Case #1, ongoing

- 58-year-old female
- 10 previous lines of therapy
- Stage IVA2 (N3) at baseline
- Response sustained at W117:
  - Skin: PR at W13, CR at W45
  - Blood: CR at W5
  - LN: PR at W5, CR at W13
  - Global: PR at W13, CR at W45



#### Patient Case #2, ongoing

- 51-year-old female
- 6 previous systemic lines of therapy
- Stage IVA2 (N3) at baseline
- Response sustained at W29:
  - Skin: PR at W5, CR at W13
  - Blood: CR at W5
  - LN: CR at W5
  - Global: PR at W5, CR at W13







#### **TELLOMAK - NCT03902184** Conclusions

TELLOMAK is a Phase 2 study evaluating lacutamab monotherapy in CTCL.

- Cohort 1 enrolls relapsed and/or refractory SS patients with ≥ 2 prior systemic therapies including mogamulizumab, a <u>high unmet</u> medical need population with no approved therapy.
- This analysis (56 patients), confirms robust clinical activity of lacutamab with favorable safety profile.
  - Patients were heavily pretreated (median 5 prior systemic therapies) and had highly refractory disease
  - Responses, including CRs, were observed in multiple compartments:
    - Overall ORR 37.5% [26.0-50.6]
    - Blood ORR 48.2% [35.7-61.0]
    - Skin ORR 46.4% [34.0-59.3]
  - In patients who achieved a global response,
    - Median DoR: 12.3 months (95% CI: 5.2-NE)
    - Median time to global response: 2.8 months (range: 1-9)
    - Median time to blood & skin response: 1.0 month (range 1-6) & 2.8 months (range: 1-10) respectively

Enrollment to TELLOMAK is completed

Long-term follow-up will provide more mature data on the key study endpoints.

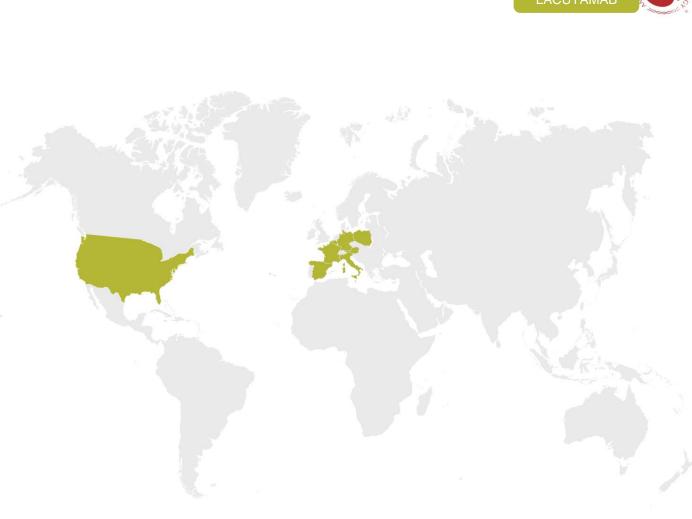


#### **TELLOMAK - NCT03902184** With Thanks

53 active sites

- USA (17)
- France (10)
- Germany (8)
- Spain (6)
- Italy (4)
- Belgium (3)
- Poland (3)
- Austria (2)

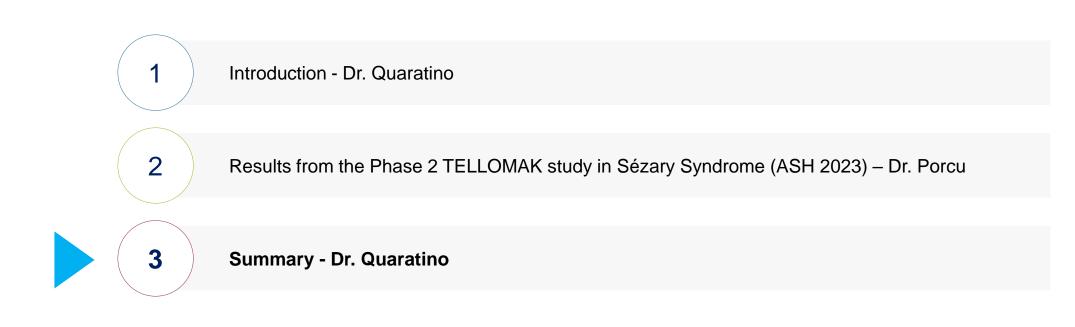
Study sponsored by Innate Pharma



Thank you to all the patients and their families, our investigators, experts, and site staff



#### Agenda



|                             | vsflow and upcoming catalysts<br>ering across our strategic objectives  | <ul> <li>Monalizumab PACIFIC-9   Phase 3 readout (AZ)</li> <li>Monalizumab NeoCOAST-2   Phase 2 readout (AZ)</li> <li>IPH5201(CD39) MATISSE   Phase 2 readout (AZ)</li> </ul>   |  |  |  |
|-----------------------------|---|---|--|--|--|
| CO23 ASCO<br>ANNUAL MEETING | <ul> <li>SAR'579 / IPH6101 ANKET<sup>®</sup> (CD123)   Phase 1<br/>(Sanofi)</li> <li>SAR'514 / IPH6401 ANKET<sup>®</sup> (BCMA)   Phase 1<br/>(Sanofi)</li> <li>IPH6501 ANKET<sup>®</sup> (CD20)   Phase 1 start</li> </ul> | <ul> <li>SAR'579 / IPH6101 ANKET® (CD123)   Next steps (Sanofi)</li> <li>SAR'514 / IPH6401 ANKET® (BCMA)   Next steps (Sanofi)</li> <li>IPH62 ANKET® (B7-H3)   Next Steps (Sanofi)</li> <li>2 ANKET options   (Sanofi)</li> </ul> |  |  |  |
| CLETTA ON WIERCAN           | <ul> <li>Lacutamab Phase 2 TELLOMAK   Final data<br/>SS &amp; MF</li> <li>Lacutamab Phase 1b PTCL   preliminary data</li> </ul>   | <ul> <li>IPH6501 ANKET<sup>®</sup> (CD20)   Phase 1 data</li> <li>IPH45 ADC (Nectin-4)   Phase 1 start</li> <li>Lacutamab CTCL   Next steps</li> <li>Lacutamab Phase 1b PTCL   Final data</li> </ul>                              |  |  |  |

2023

### 2024+



#### Summary

Create value for patients and shareholders

#### **Strategic Priorities**

Early R&D focus to drive value through later stage partnerships Robust pipeline of Antibody engineering capabilities Advancing lacutamab Maximizing ANKET<sup>®</sup> platform Continuing to explore ADC

Track record of strategic partnerships

AstraZeneca (monalizumab, IPH5201), Sanofi (SAR443579 / IPH6101, SAR'514 / IPH6401, IPH62, 2 options), Takeda (ADC)

Strong cash position €121.9m\* as of September 30, 2023 Sufficient to fund operations into H2 2025

## ) **innate** pharma

# Thank you

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