

Milan, Italy
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10th

International Meeting on
PULMONARY RARE DISEASES
AND ORPHAN DRUGS

EXTRACELLULAR VESICLES PROFILE IN BRONCHOALVEOLAR LAVAGE FROM IPF, HP AND SARCOIDOSIS PATIENTS: A MULTICENTER STUDY

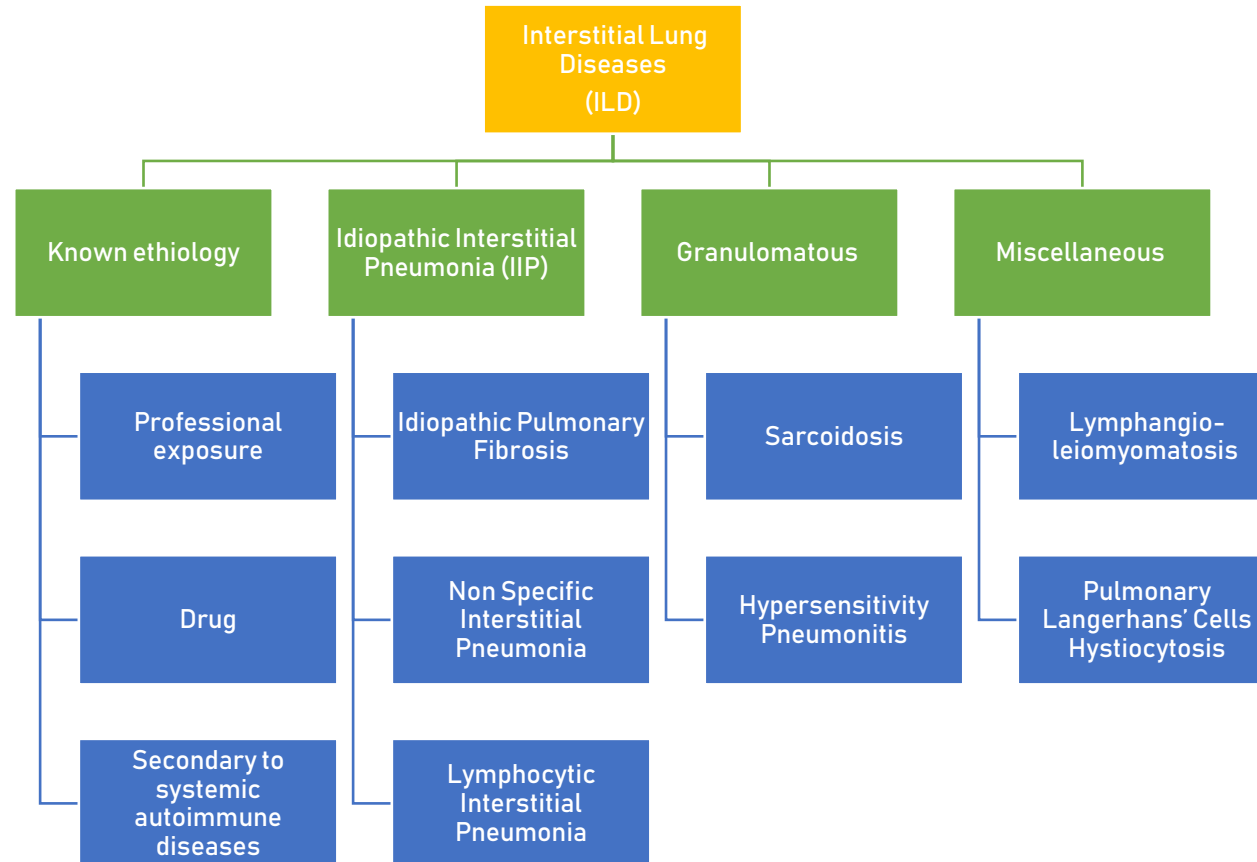
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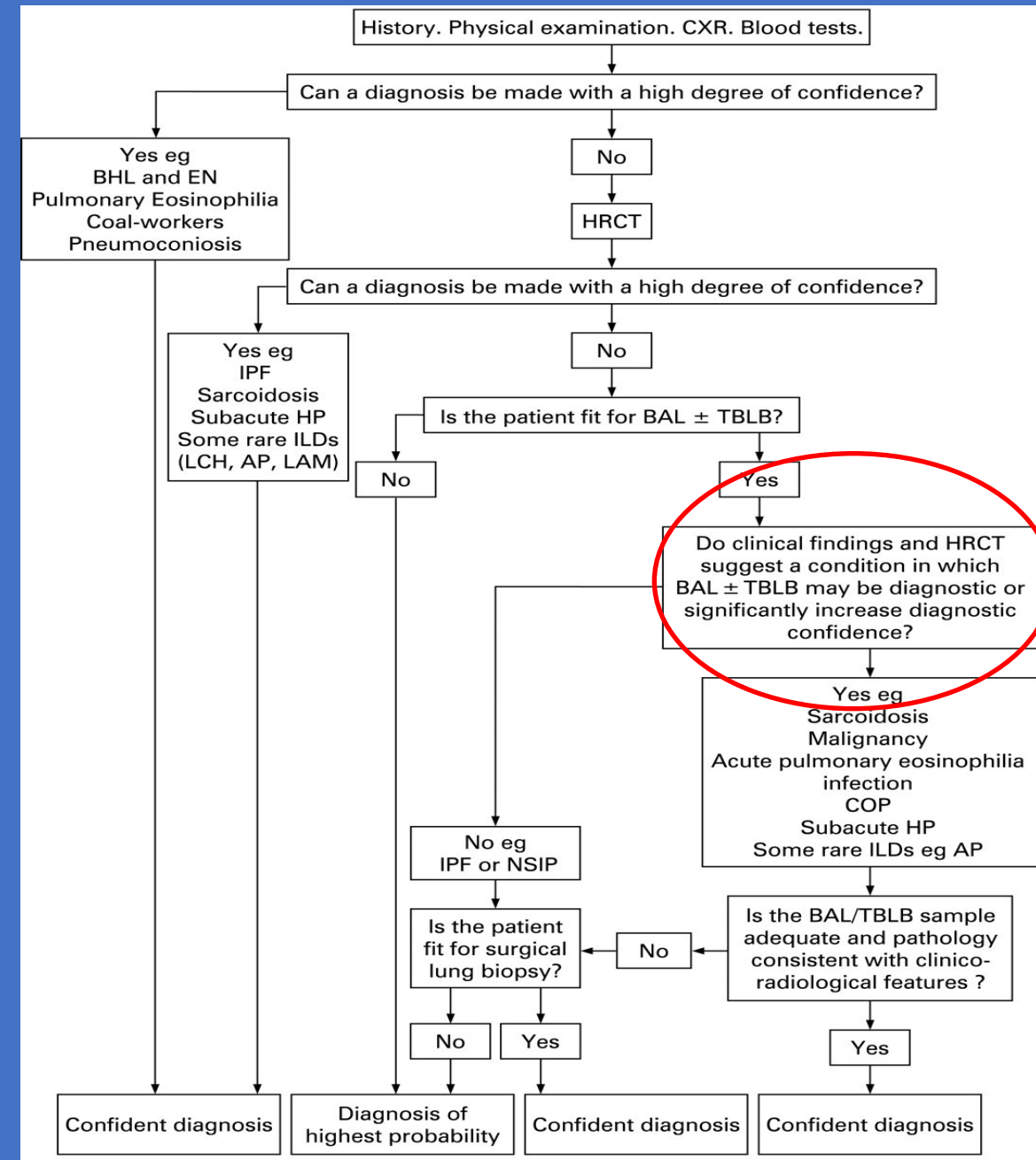
INTERSTITIAL LUNG DISEASES- CLASSIFICATION



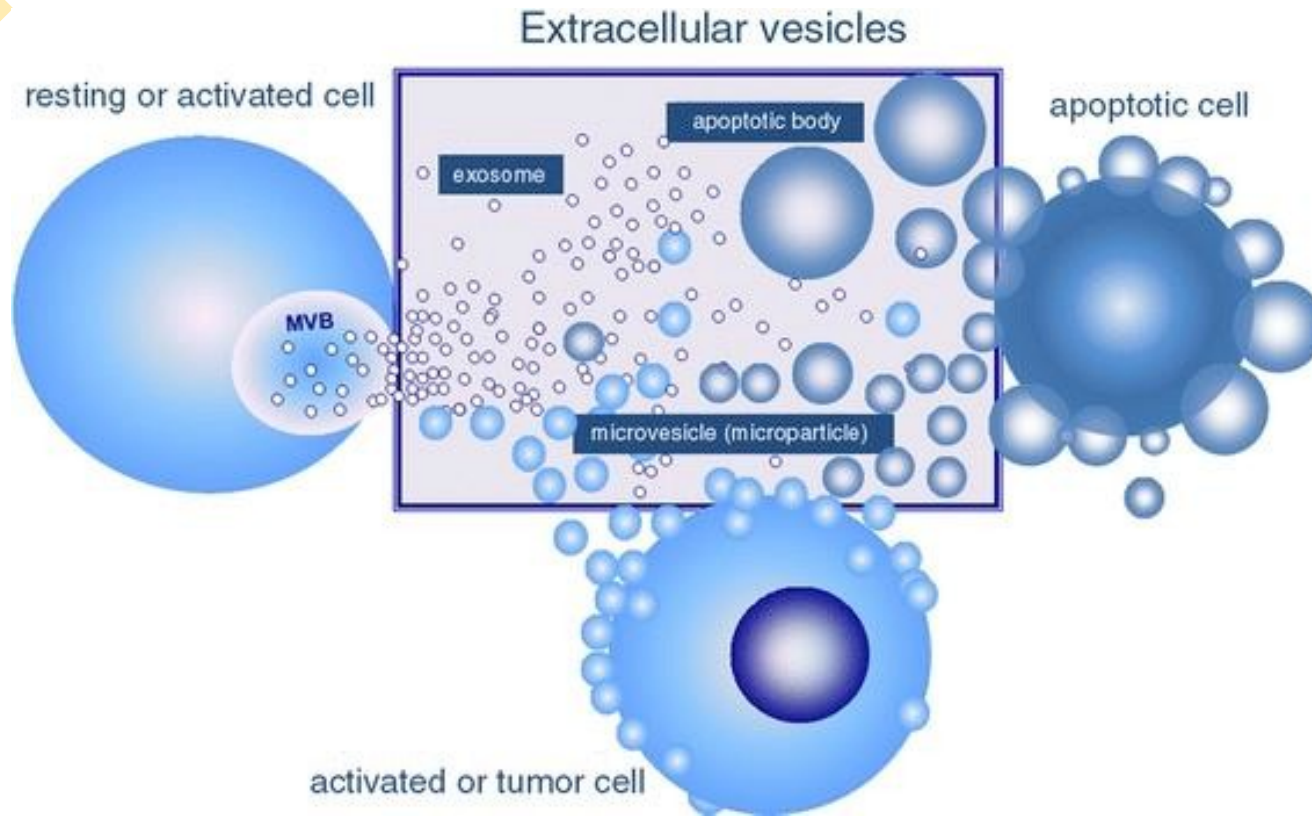
- *Heterogeneous group*
- *Non-neoplastic*
- *Destruction of the lung parenchyma*
- **FIBROSIS**

WHEN DOES THE LABORATORY COME INTO PLAY?

- Bronchoalveolar lavage is a biological fluid representative of the alveolar compartment useful for searching inflammatory cellular infiltrates and as a source of biomarkers in interstitial lung diseases



EXTRACELLULAR VESICLES

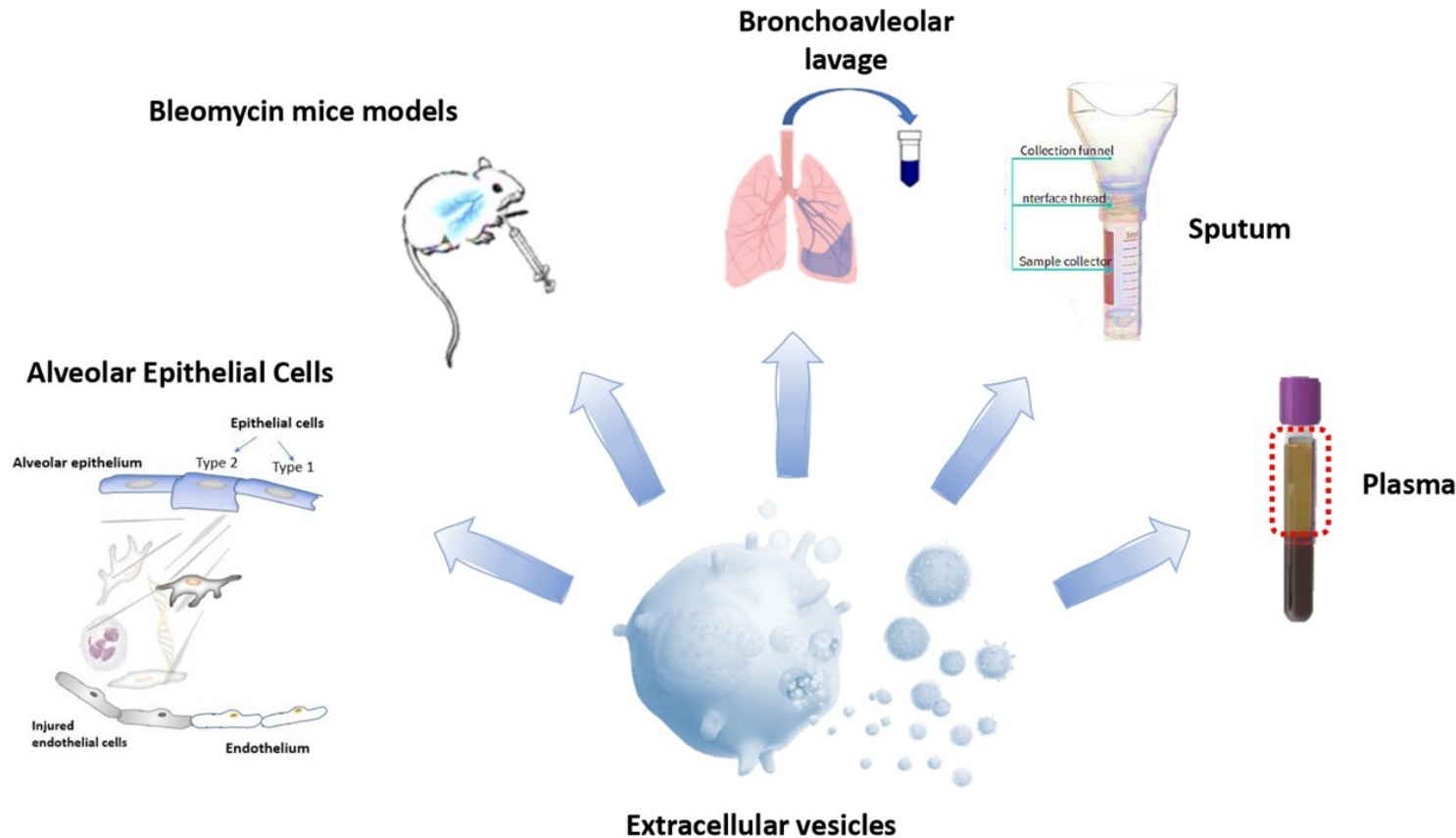


Generic term for particles released naturally by cells

Enclosed in a lipid bilayer and cannot replicate

No consensus has yet emerged on specific markers of EV subtypes with particular biogenetic pathways

EXTRACELLULAR VESICLES IN PULMONARY FIBROSIS MODELS AND BIOLOGICAL FLUIDS



pivotal role as
mediators of cell-cell
communication

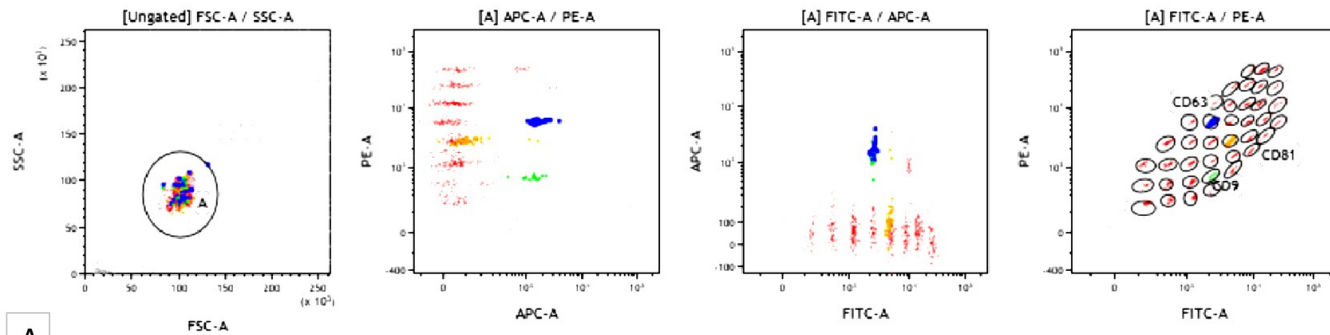
cooperator role in the
development of lung
diseases such as IPF

AIM AND METHODS

- To compare 37 exosomal surface markers through flow cytometry in BAL from patients affected by IPF, sarcoidosis and hypersensitivity pneumonitis (HP), enrolled at Siena Referral Centre for rare lung diseases.
- To corroborate these findings, a validation cohort was enrolled from two referral centers for ILDs (Barcelona and Foggia).

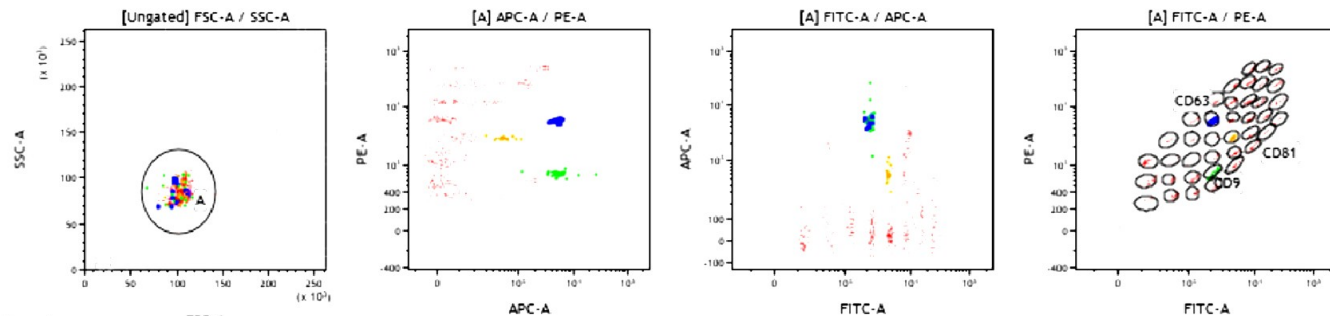
Parameters	Study Cohort			Validation Cohort		
	IPF (n = 17)	HP (n = 24)	Sarcoidosis (n = 42)	IPF (n = 44)	HP (n = 11)	Sarcoidosis (n = 10)
Age (years)	64.7 ± 23.8	68.23 ± 11.9	52.2 ± 21.2	62.6 ± 19.8	67.11 ± 9.2	50.2 ± 20.5
	13/4	14/10	16/36	31/13	6/5	3/7
Smoking habit (never/former)	6/11	6/18	15/27	15/28	4/7	4/6

37 ALVEOLAR EVS SURFACE MARKERS THROUGH FLOWCYTOMETRY



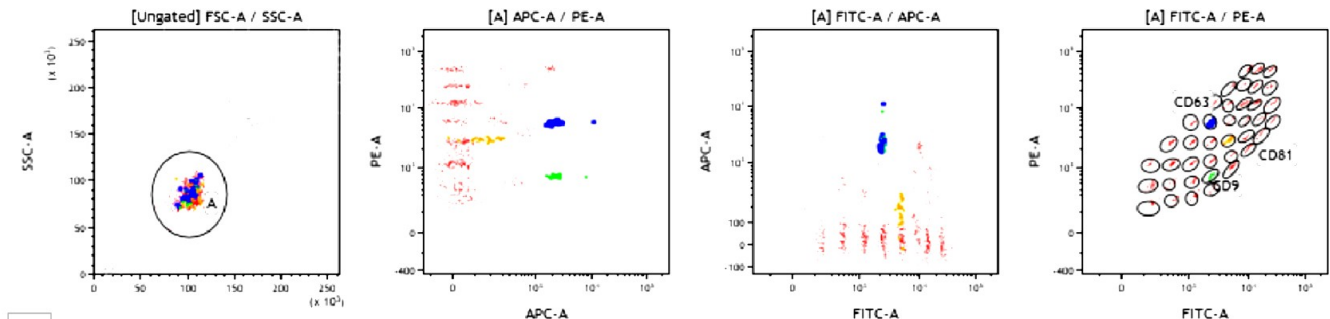
SIENA

Study
cohort



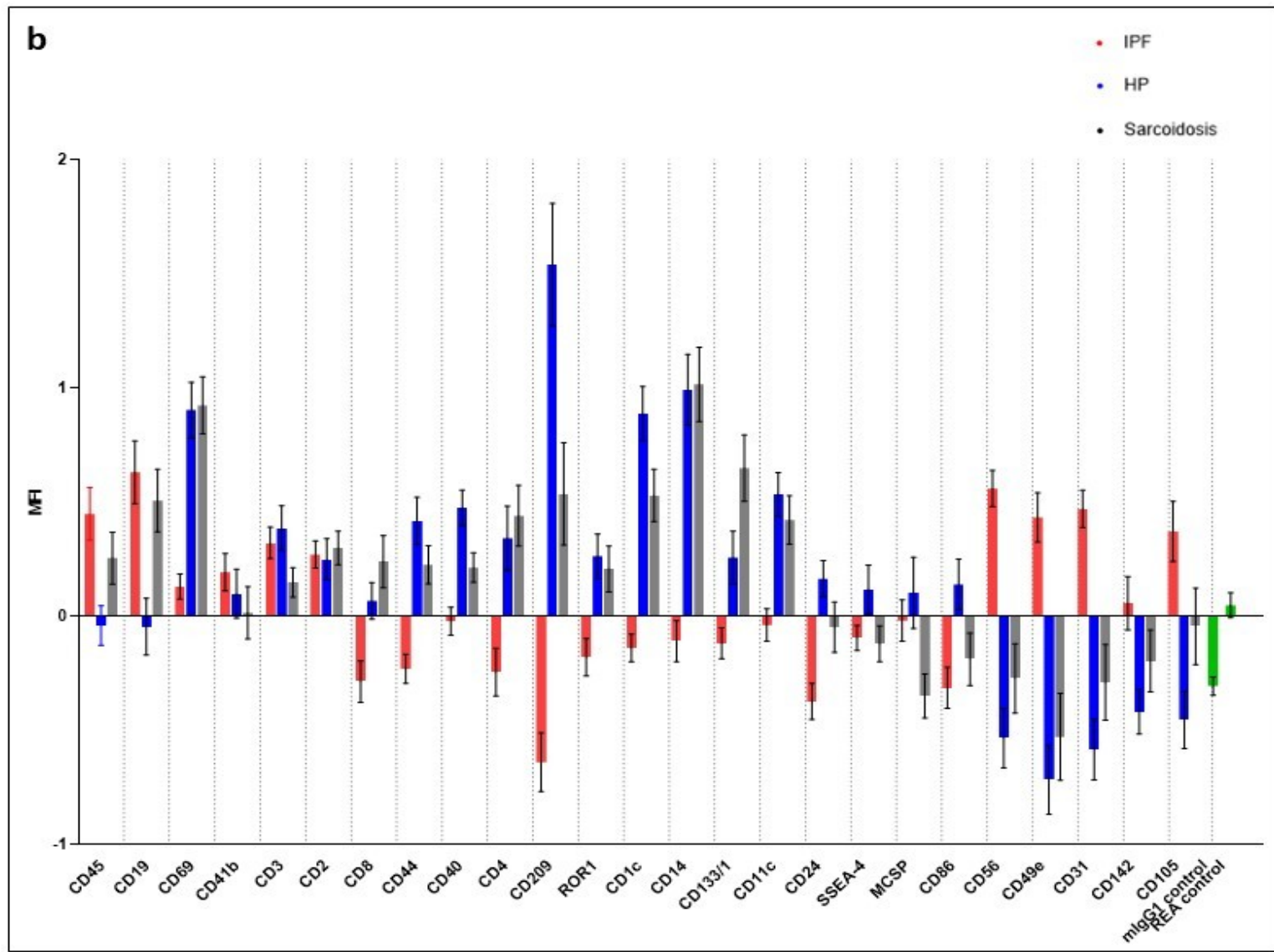
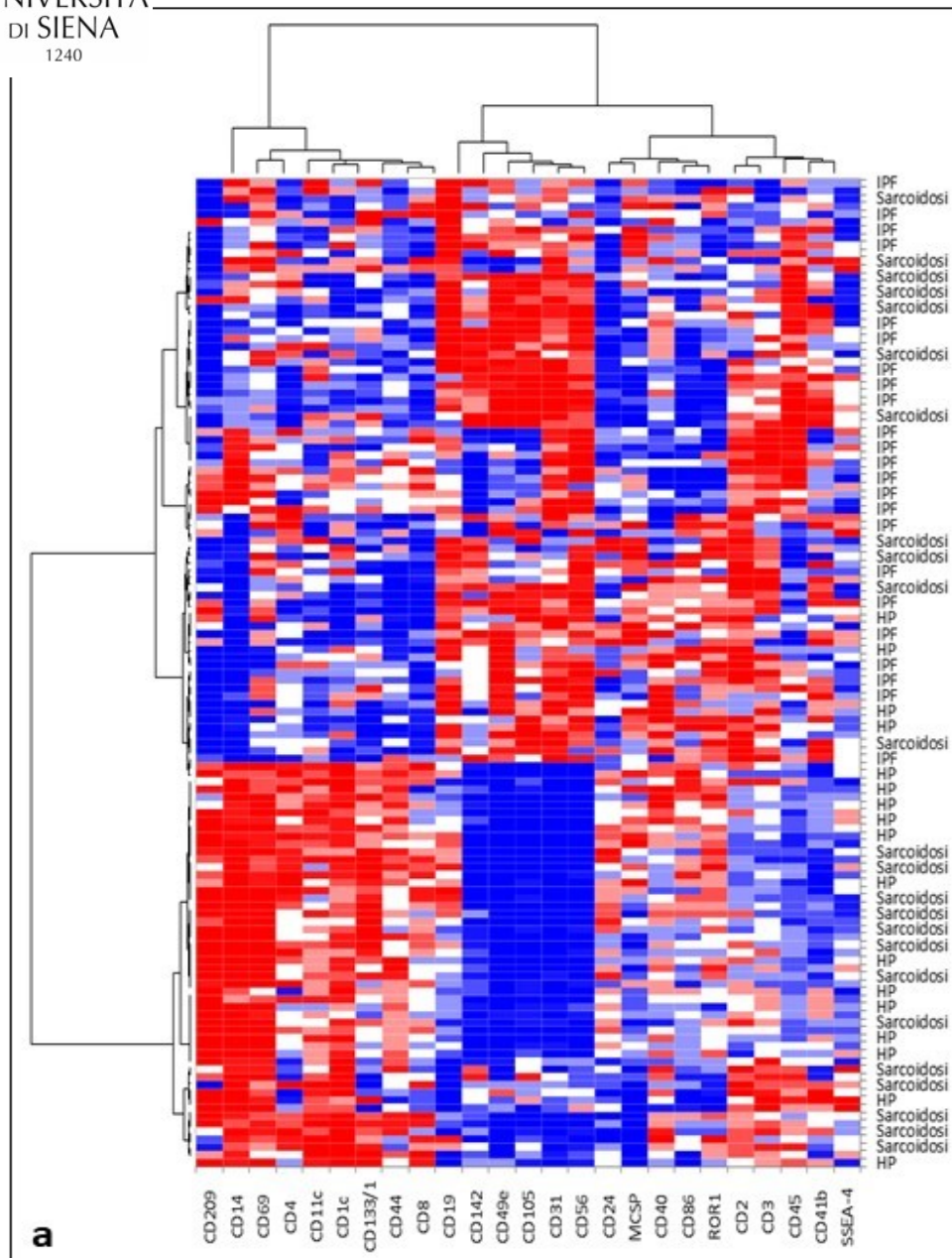
FOGGIA

Validation
cohort



BARCELLONA

d'Alessandro, M. et al. Int. J. Mol. Sci. **2023**,
24, 4071.

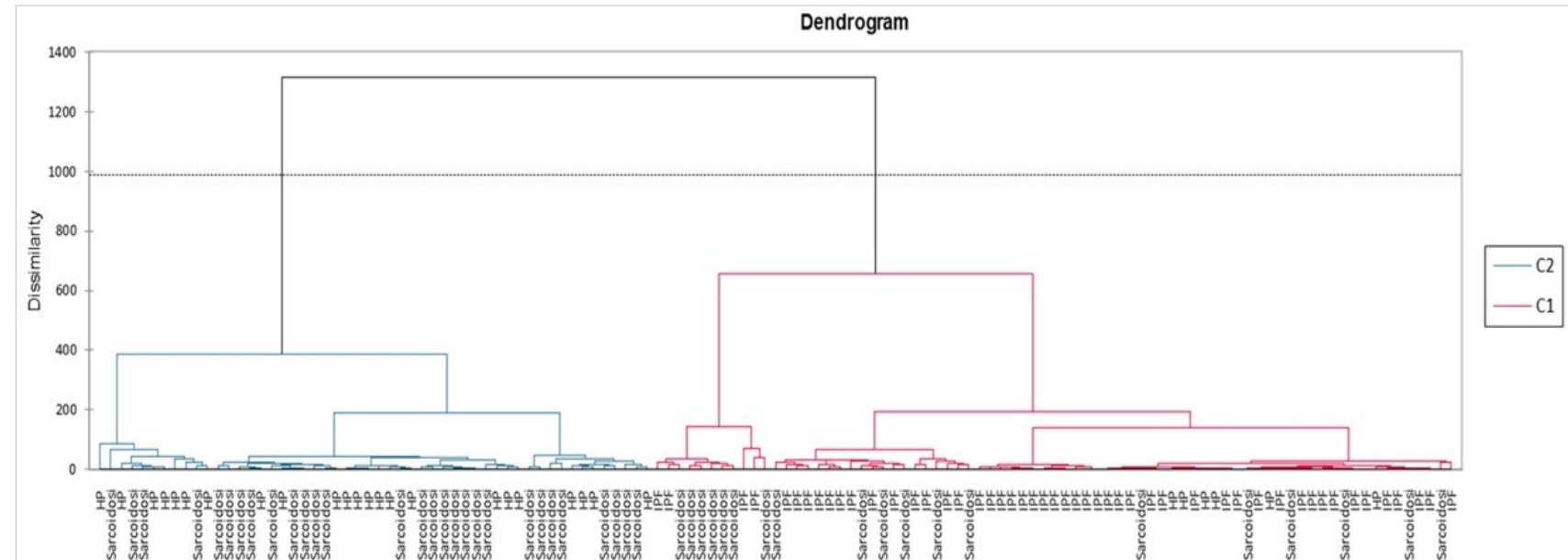


AGGLOMERATIVE HIERARCHICAL CLUSTERING (AHC) ANALYSIS

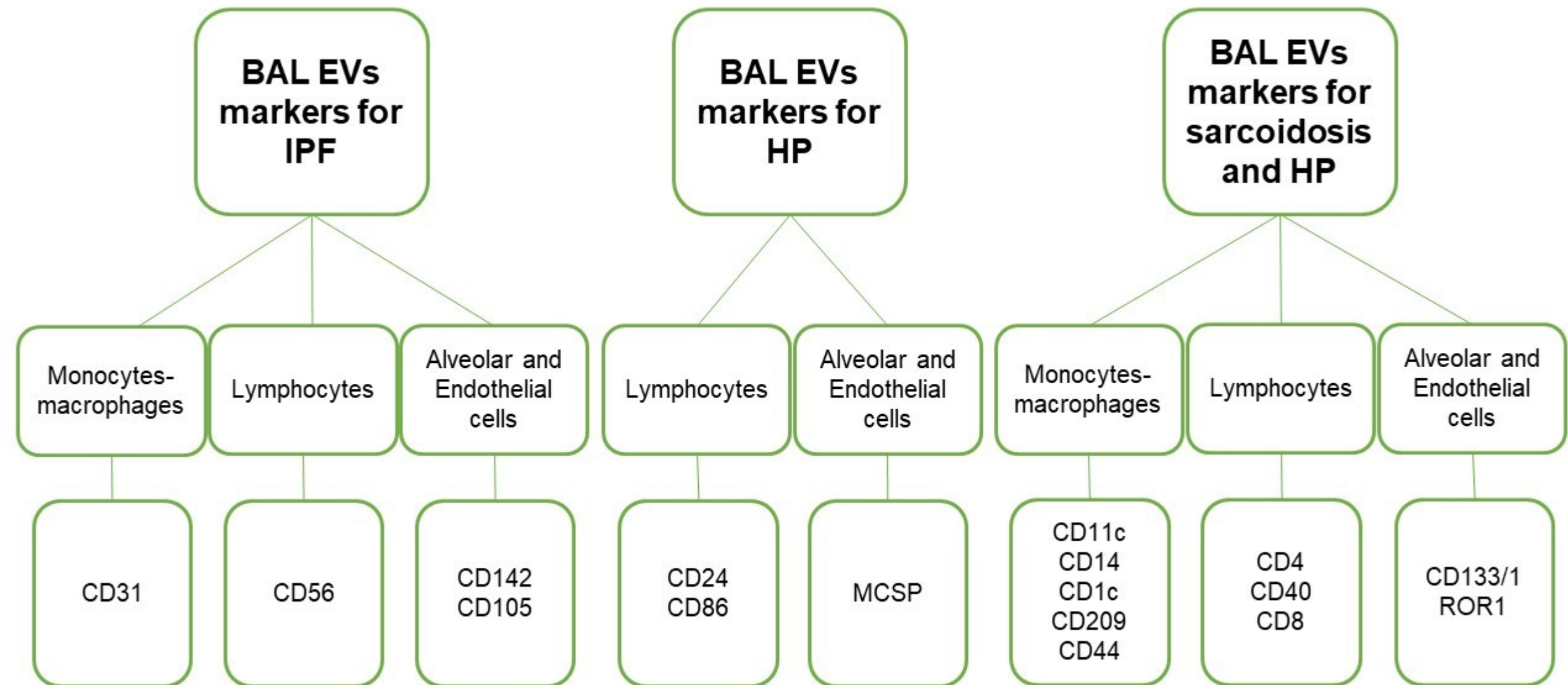
Dendrogram plot clustered
IPF, HP, sarcoidosis based on
dissimilarities between
values of 37 EV surface
markers

Evolution of indices:

Number of clusters	2	3	4	5
Silhouette index	0.276	0.275	0.296	0.231
Hartigan index (H)	28.861	19.192	9.649	9.906
H(k-1) - H(k)	18.815	9.669	9.543	-0.257
Calinski & Harabasz index	47.676	43.582	39.714	34.292



EV MARKERS DIVIDED ACCORDING TO THE DISEASES



TAKE-HOME MESSAGE

- validity of the flow cytometric method to phenotype and characterize EV surface markers
- viability of the alveolar compartment to identify lung-specific markers for IPF and HP
- fibrotic HP: post-inflammatory cell-regulated ILD
- IPF: related to tissue remodeling and repair



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Thank you for the attention



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SANTA CREU I
SANT PAU

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UNIVERSITAT AUTÒNOMA DE BARCELONA



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& Harefield

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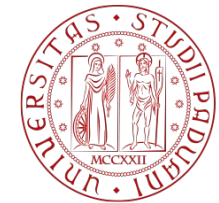


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