



XXXII CONGRESSO NAZIONALE AIRO
XXXIII CONGRESSO NAZIONALE AIRB
XII CONGRESSO NAZIONALE AIRO GIOVANI

AIRO2022

Radioterapia di precisione per un'oncologia innovativa e sostenibile

BOLOGNA, 25-27 NOVEMBRE
PALAZZO DEI CONGRESSI

 Associazione Italiana
Radioterapia e Oncologia clinica

 Società Italiana di Radiobiologia

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Impact of geriatric assessment on QoL of elderly onco-hematologic patients (ONCO-AGING) candidate to complex treatments: interim results

E. Ferrara¹, C. Pisani¹, F. Biello², G. Burrafato², R. Bruna³, A. Patriarca³, A. Mahmoud^{2,4}, V. Martini^{2,4}, C. Branni^{2,4},
M. Cappelli⁵, E. Catania⁵, D. Ferrante⁴, G. Gaidano^{3,4}, A. Gennari^{2,4}, M. Krengli^{1,4}

Carla Pisani, MD PhD

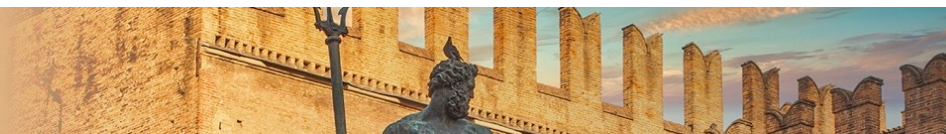
¹ Department of Radiation Oncology, University Hospital Maggiore della Carità (Novara, Italy)

² Department of Oncology, University Hospital Maggiore della Carità (Novara, Italy)

³ Department of Haematology, University Hospital Maggiore della Carità (Novara, Italy)

⁴ Department of Translational Medicine, Università del Piemonte Orientale (Novara, Italy)

⁵ Department of Palliative Care, University Hospital Maggiore della Carità (Novara, Italy)

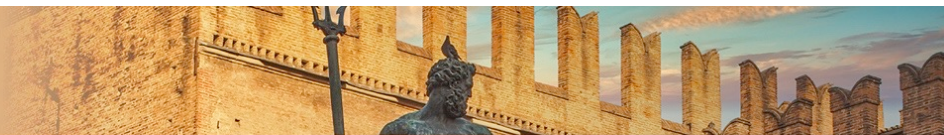


DICHIARAZIONE

Relatore: CARLA PISANI

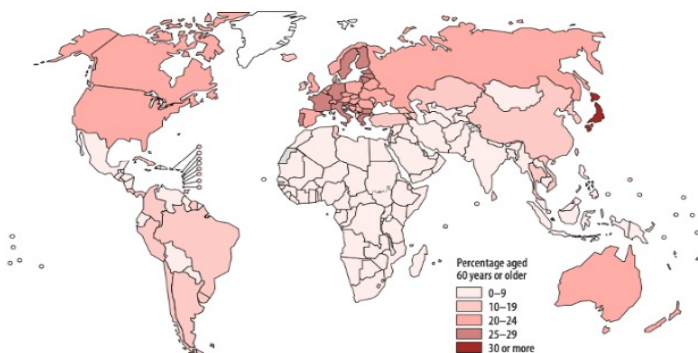
Come da nuova regolamentazione della Commissione Nazionale per la Formazione Continua del Ministero della Salute, è richiesta la trasparenza delle fonti di finanziamento e dei rapporti con soggetti portatori di interessi commerciali in campo sanitario.

- Posizione di dipendente in aziende con interessi commerciali in campo sanitario: **NIENTE DA DICHIARARE**
- Consulenza ad aziende con interessi commerciali in campo sanitario: **NIENTE DA DICHIARARE**
- Fondi per la ricerca da aziende con interessi commerciali in campo sanitario: **NIENTE DA DICHIARARE**
- Partecipazione ad Advisory Board: **NIENTE DA DICHIARARE**
- Titolarità di brevetti in compartecipazione ad aziende con interessi commerciali in campo sanitario: **NIENTE DA DICHIARARE**
- Partecipazioni azionarie in aziende con interessi commerciali in campo sanitario: **NIENTE DA DICHIARARE**
- Altro

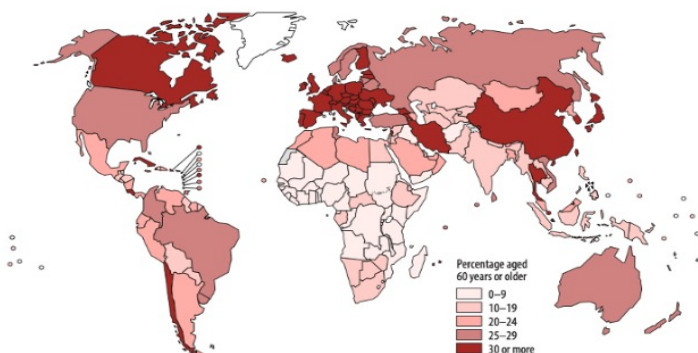


Oncological treatment in elderly patients

Proportion of population aged 60 years or older, by country, 2015



Proportion of population aged 60 years or older, by country, 2050 projections



The need of oncological treatment in elderly patients is rising together with the increase in life expectancy

→ **How could we quantify the frailty?**

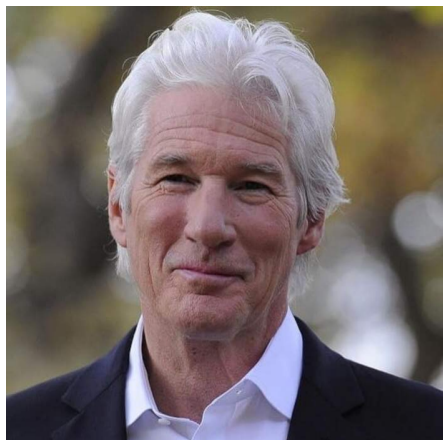
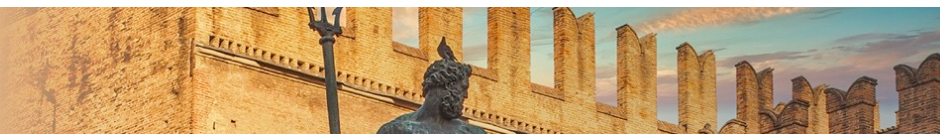
Older patients are under-represented in clinical trials

→ **less evidence-based information exists to guide the treatment of these patients**

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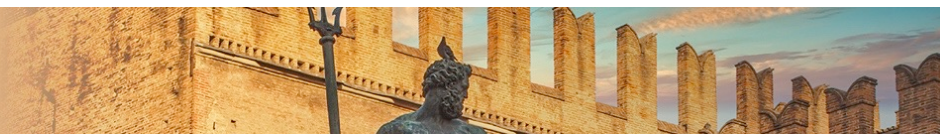
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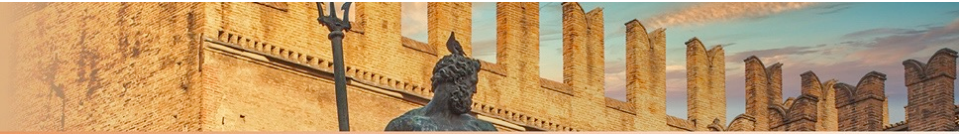
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Assessing older cancer patients

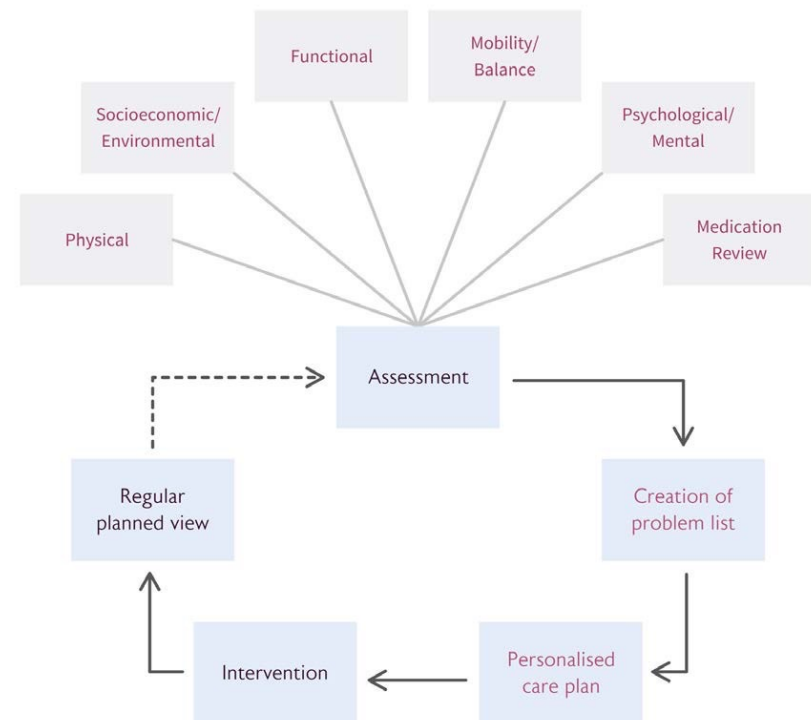
- Focus on **disease** characters (history, histology, staging).
- Focus on **patient's** characters and functional status (**NOT ONLY PS!**)
- Geriatric 8 (G8) can be used to screen and detect patients who could benefit from a geriatric assessment.

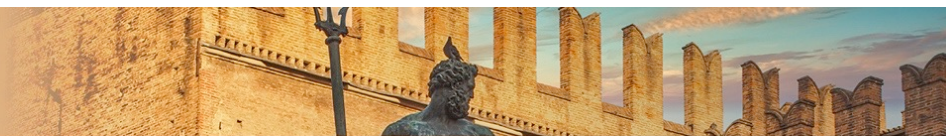
	Items	Possible answers (score)
A	Has food intake declined over the past 3 months due to loss of appetite, digestive problems, chewing or swallowing difficulties?	0 : severe decrease in food intake
		1 : moderate decrease in food intake
		2 : no decrease in food intake
B	Weight loss during the last 3 months	0 : weight loss > 3 kg
		1 : does not know
		2 : weight loss between 1 and 3 kgs
C	Mobility	3 : no weight loss
		0 : bed or chair bound
		1 : able to get out of bed/chair but does not go out
E	Neuropsychological problems	2 : goes out
		0 : severe dementia or depression
		1 : mild dementia or depression
F	Body Mass Index (BMI (weight in kg) / (height in m ²))	2 : no psychological problems
		0 : BMI < 19
		1 : BMI = 19 to BMI < 21
		2 : BMI = 21 to BMI < 23
H	Takes more than 3 medications per day	3 : BMI = 23 and > 23
		0 : yes
		1 : no
P	In comparison with other people of the same age, how does the patient consider his/her health status?	0 : not as good
		0.5 : does not know
		1 : as good
		2 : better
	Age	0 : >85
		1 : 80-85
		2 : <80
TOTAL SCORE		0 – 17



Comprehensive Geriatric Assessment (CGA)

- 9 questionnaires used to assess overall health status in geriatric patients.
- *CGA can be used to stratify patients screened with G8:*
 - **Unfit (prefrail)**
 - **Frail**
- Proposed to identify patients who require personalised interventions to improve outcomes.



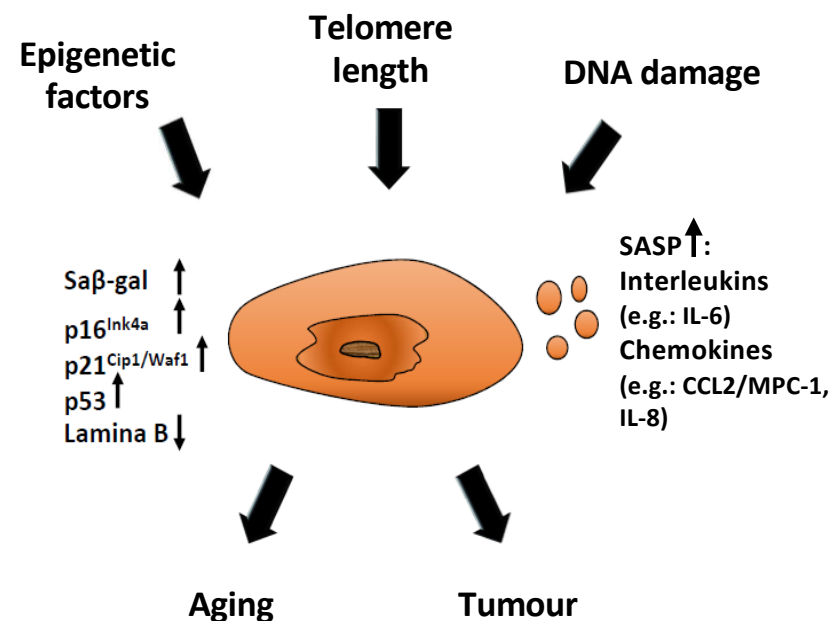


Aging and cellular senescence

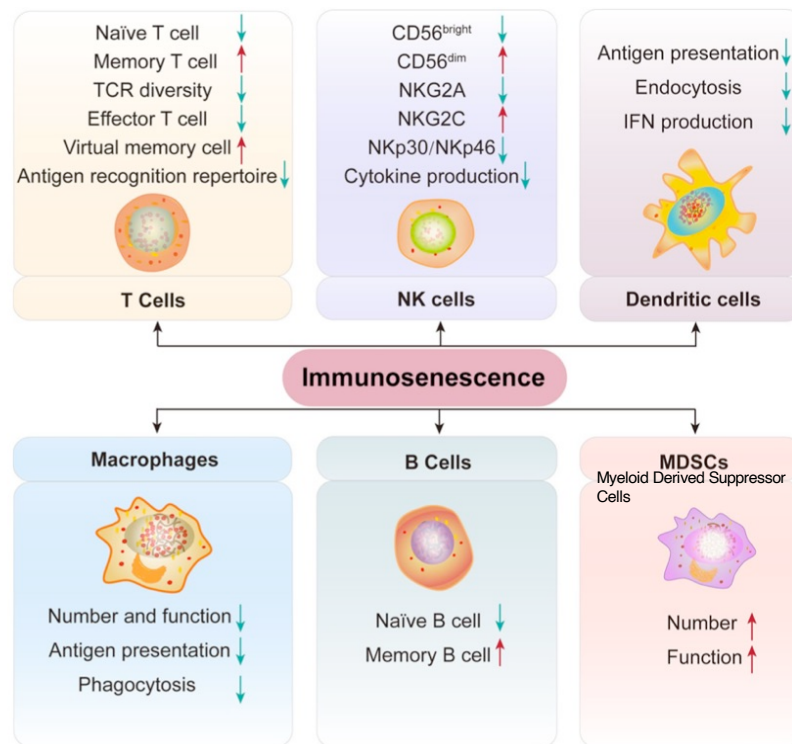
Induction of a senescent phenotype



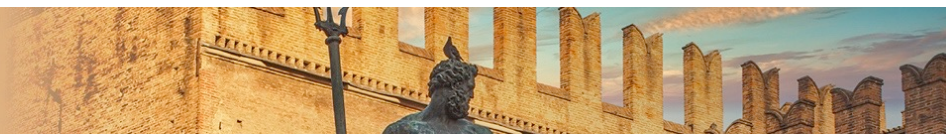
Promotion of aging-related diseases
(e.g.: chronic inflammation, cancer).



Immunosenescence

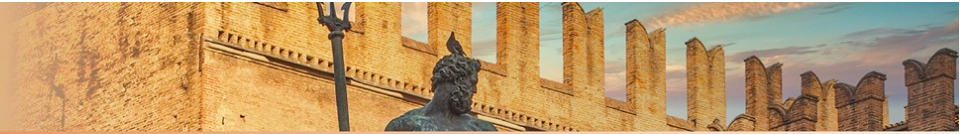


- Senescence in immune cells is associated with a decrease in adaptive immune functions, decrease infection resistance, and increased autoimmune risk
- the inability to induce a strong immune reaction against them results in higher susceptibility to cancer in people who have a less efficient immune system such as old people and immune-suppressive people
- The low number and low efficient immune system in older people are partially due to a high number of senescent



Aims of the study

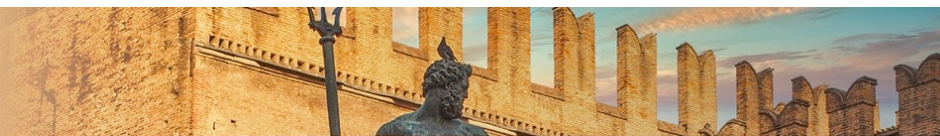
- Test the impact of the CGA on the QoL of elderly onco-haematological patients eligible for multimodal/targeted therapy, who resulted fragile at the G8 screening.
- Investigate the impact of CD3+ T-cells senescence and MDSCs in the peripheral blood on cancer progression and overall survival (OS).



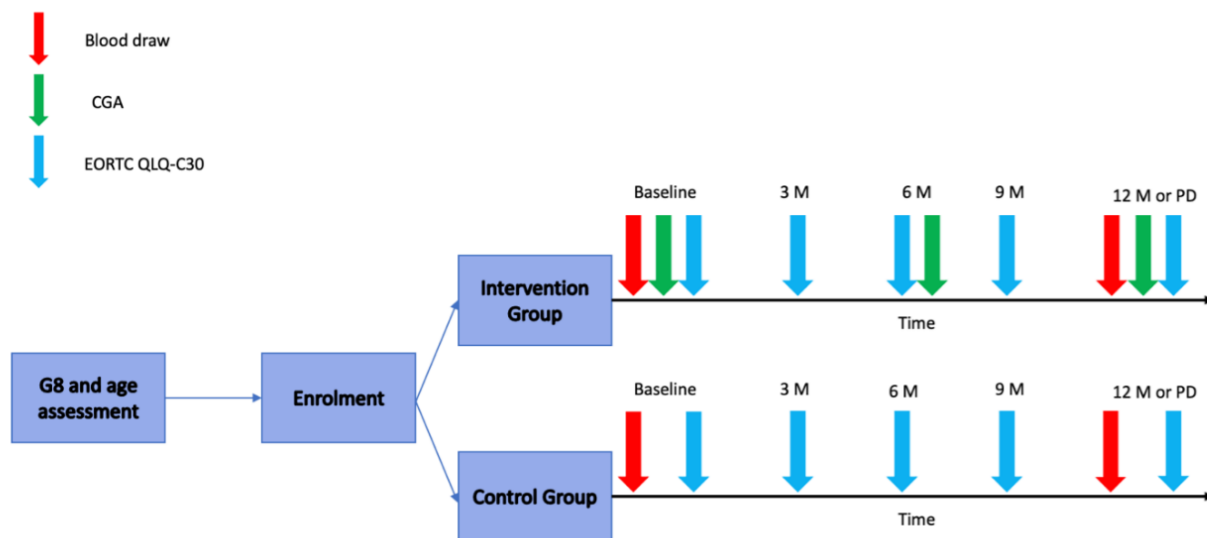
Materials and methods (1)

Inclusion criteria:

- Patients aged ≥ 65 years old diagnosed with solid/haematological neoplasia.
- Eligible to 1° line therapy with CT or biological/target drugs or concurrent radio-chemotherapy.
- Screening G8 score $\leq 14/17$.



Materials and methods (2)

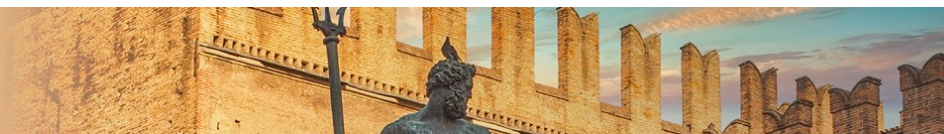


Intervention group:

- Senescent CD3+ T cells/MDSCs evaluation at baseline and after 12 months/PD.
- QLQ-C30 QoL questionnaire every 3 months.
- Geriatric evaluation + CGA at baseline, at 6 months and 12 months/PD.

Control group:

- Senescent CD3+ T cells/MDSCs evaluation at baseline and after 12 months/PD.
- QLQ-C30 QoL questionnaire every 3 months.



Gender (%)

Male	61.9
Female	38.1

Age

Median	75 (range: 65-91)
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Number of comorbidities (%)

< 4	53.5
≥ 4	46.5

Number of concomitant drugs (%)

< 4	29.1
≥ 4	70.9

Screening G8 score

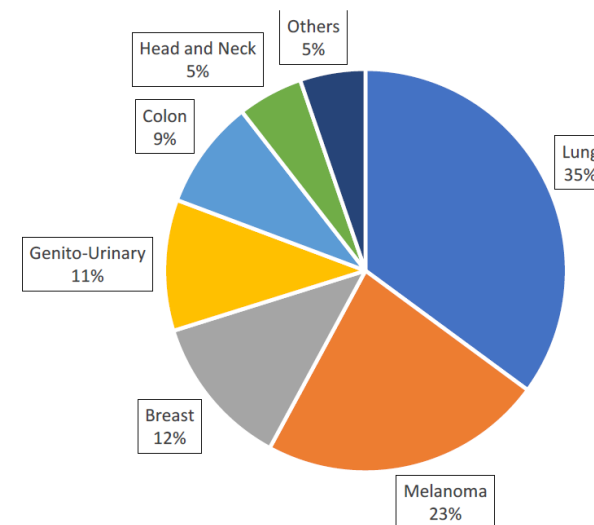
Median	12 (range: 3-14)
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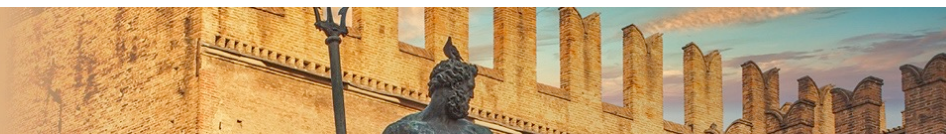
Treatment modality (%)

Targeted/biological therapy	76.1
Concurrent radio-chemotherapy	23.9

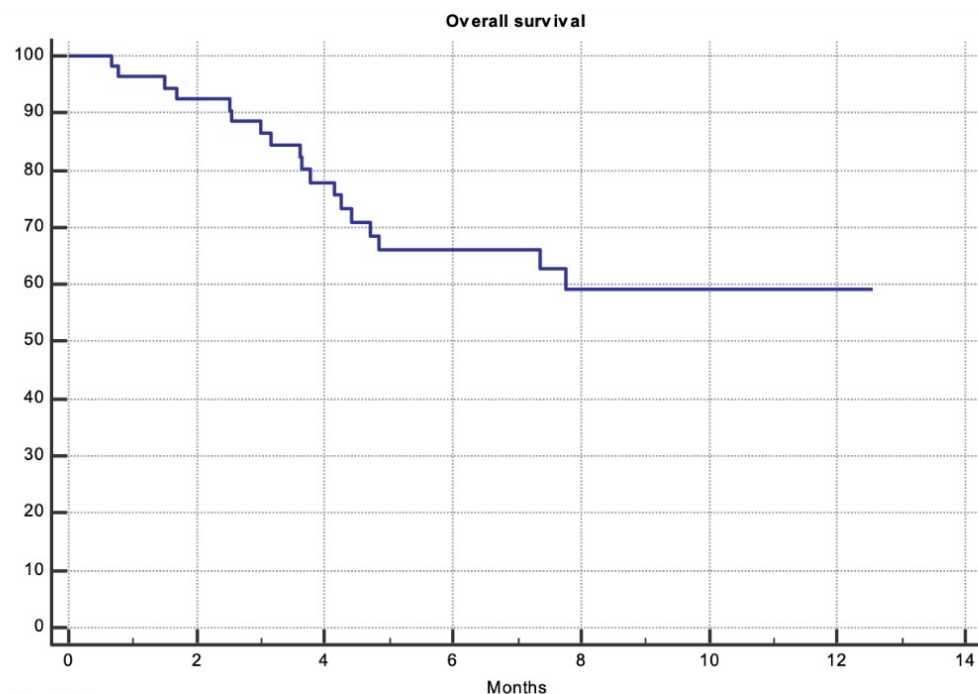
Pz characteristics

- ✓ 155 patients enrolled from 12/2019 to 02/2022
- ✓ Interim results for 57 patients (1:1 randomisation).



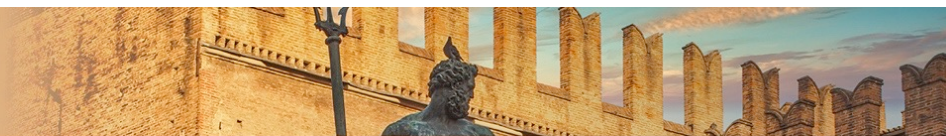


Overall survival (OS)

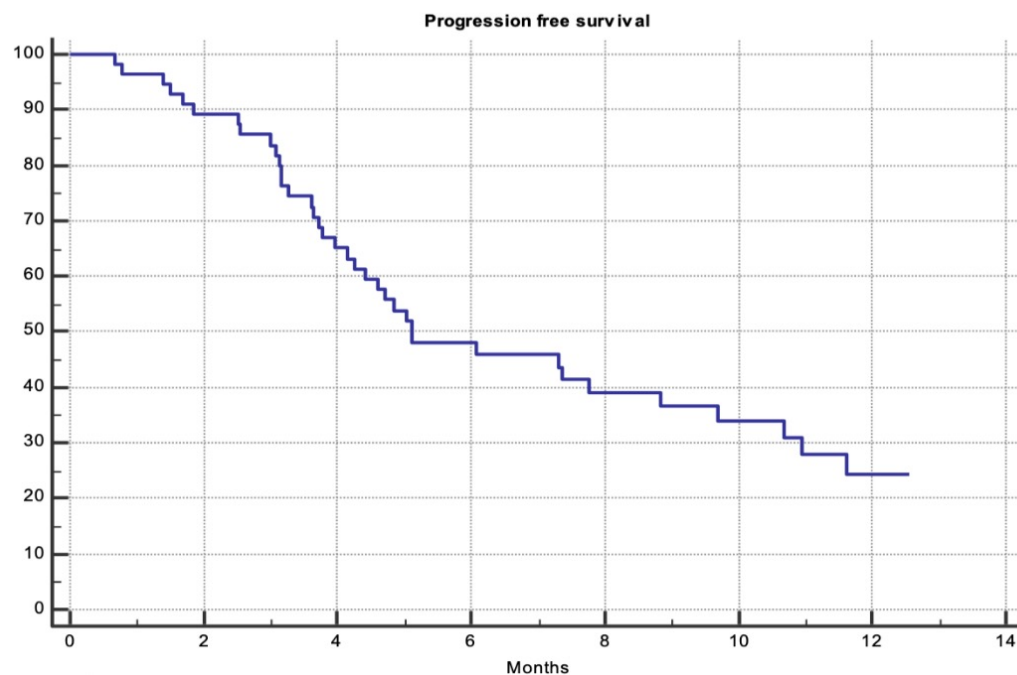


- 1° quartile: 4.1 months.
- Median OS: not reached.

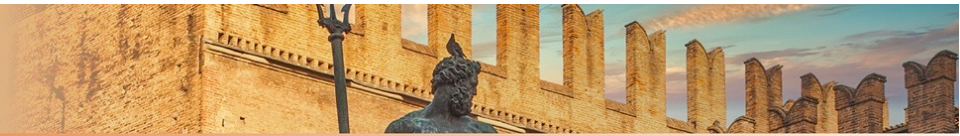
The median age of the cohort must be taken in mind (75, range 65-91) since as per se is a risk factor for death.



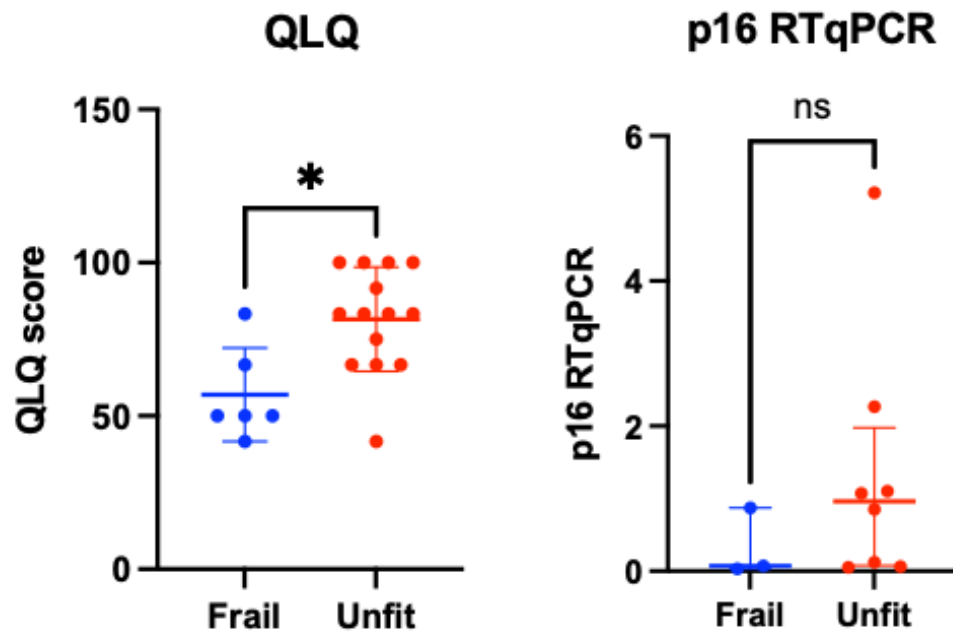
Progression-free survival (PFS)



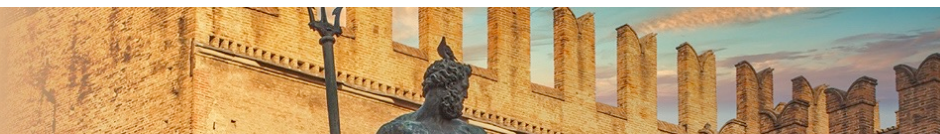
- 1° quartile: 3.1 months.
- Median PFS: 5 months.
- 3° quartile: 11.62 months



Frail vs unfit: differences in QoL and p16 expression



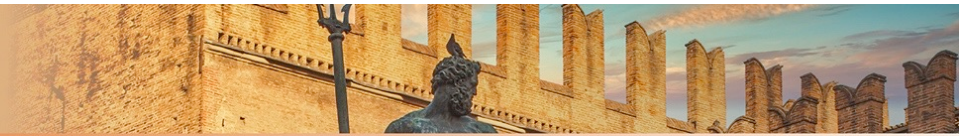
- Unfit patients present a significantly better QoL compared to frail ones (median QLQ score: 83.33 vs 48, $p < 0.05$).
- Difference in expression of p16 in CD3+ T cells between frail and unfit patients (although not significant at the moment).



Impact of CGA and p16 on OS and PFS

- *Although not statistically significant at the moment, both CGA and senescent CD3+ T-cells may identify patients at higher risk of death.*
- *No univocal correlation between the questionnaires/p16 expression and the risk of disease progression.*

G8		QLQ		RFI		p16 (RT)		p16(dd)		
ρ	p-value	ρ	p-value	ρ	p-value	ρ	p-value	ρ	p-value	
G8			0.19	0.157	-0.41	0.061	0.10	0.597	0.014	0.951
	QLQ				-0.63	0.002	0.17	0.357	0.093	0.668
			RFI				-0.74	0.009	-0.500	0.391
					P16					
						P16				



Conclusions

- CGA and T-cell senescence may predict patients' outcome in terms of OS.
- Frail patients at CGA have worse QLQ score → importance of QoL assessment in clinical setting.
- Correlation between CGA and p16 expression → potential biochemical biomarker for frailty.
- Preliminary results that still have to be confirmed and completed.
- Build evidence on the management of elderly and fragile cancer patients in order to personalise the therapeutic approach.



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Thanks for your attention



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