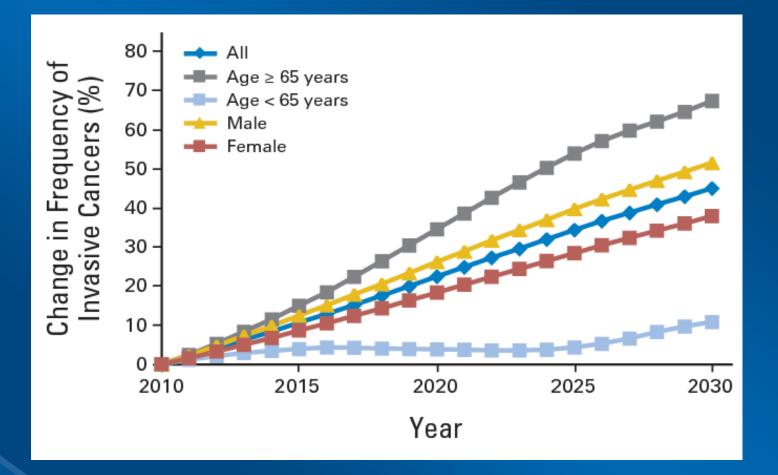
Updates in Geriatric Oncology

Allison Magnuson, DO, MS Associate Professor of Medicine University of Rochester Medical Center



Increasing Number of Older Patients with Cancer



Pal S.K., Hurria A. JCO (2010)



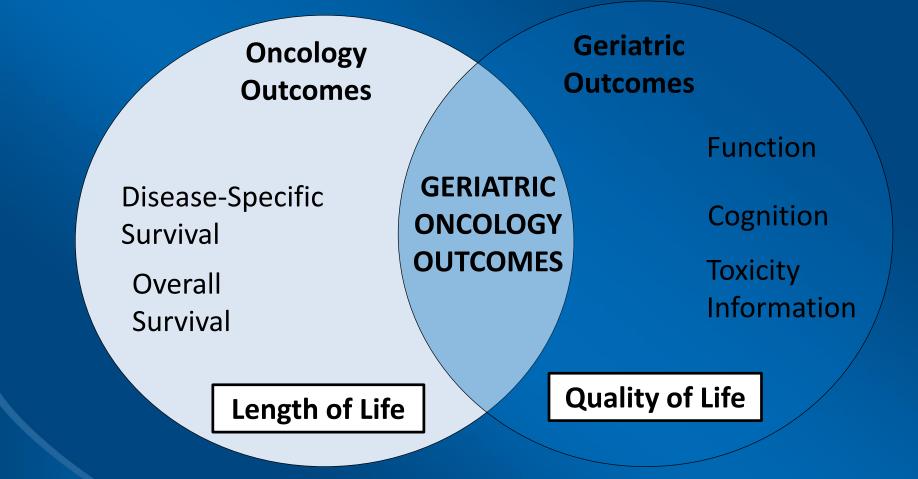
All Older Patients with Cancer are Not the Same







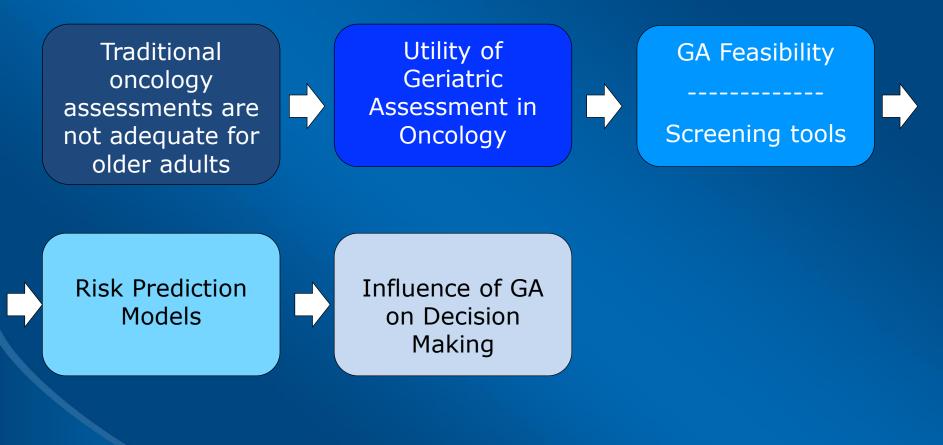
Geriatric Oncology



Williams GR, Mackenzie A, Magnuson A, et al. Journal of geriatric oncology 2016;7:249-57.



The Evolution of Geriatric Oncology





JOURNAL OF CLINICAL ONCOLOGY

ASCO SPECIAL ARTICLE

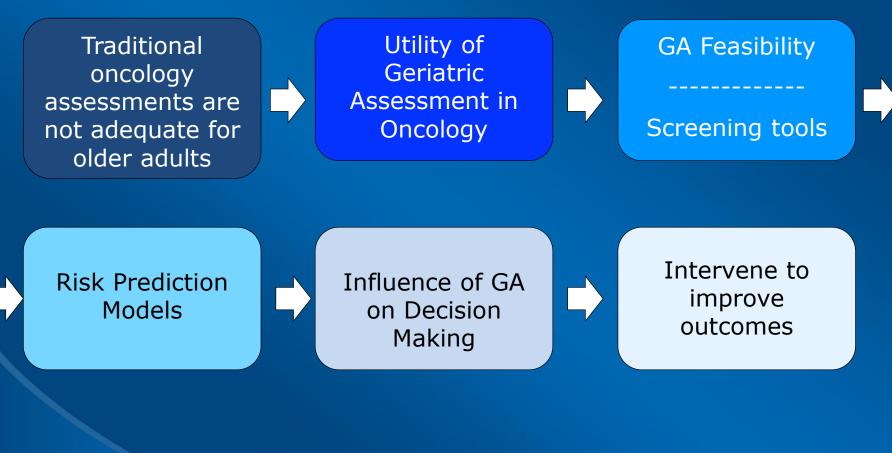
Practical Assessment and Management of Vulnerabilities in Older Patients Receiving Chemotherapy: ASCO Guideline for Geriatric Oncology

Supriya G. Mohile, William Dale, Mark R. Somerfield, Mara A. Schonberg, Cynthia M. Boyd, Peggy S. Burhenn, Beverly Canin, Harvey Jay Cohen, Holly M. Holmes, Judith O. Hopkins, Michelle C. Janelsins, Alok A. Khorana, Heidi D. Klepin, Stuart M. Lichtman, Karen M. Mustian, William P. Tew, and Arti Hurria

Mohile, et al, JCO, 2018



The Evolution of Geriatric Oncology





GA to Improve Communication



- To improve <u>communication</u> about age-related concerns of older patients with advanced cancer and their caregivers
 - Direct communication about age-related concerns in clinical encounters
 - Patient satisfaction with communication about age-related concerns
- Providing a summary of geriatric assessment results with recommendations for GA-guided interventions
 - Has potential to improve communication about age-related concerns of older patients with cancer and their caregivers

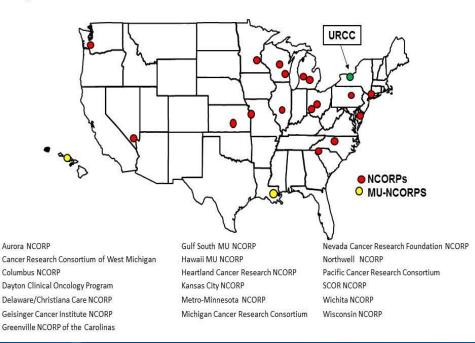
PI: Mohile; Funding: PCORI Mohile, et al. JAMA Onc, 2019



University of Rochester NCORP Research Base

University of Rochester Cancer Center NCORP Research Base

Map of Affiliates - 2016



The 2017 URCC NCORP Research Base Annual Meeting with 20 NCORP Affiliates



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Accrual: 305 oncologists, 541 patients, 414 caregivers





Patient and Caregiver Eligibility Criteria

Patients

- Age ≥70 years
- Diagnosis of advanced solid tumor or lymphoma
- Have ≥1 GA Impairment (other than polypharmacy)
- Will see their oncologist for next ≥3 months and willing to participate in study visits
- Have decision-making capacity, or, if not, oncologist has obtained consent from health-care proxy
- Able to read and understand English



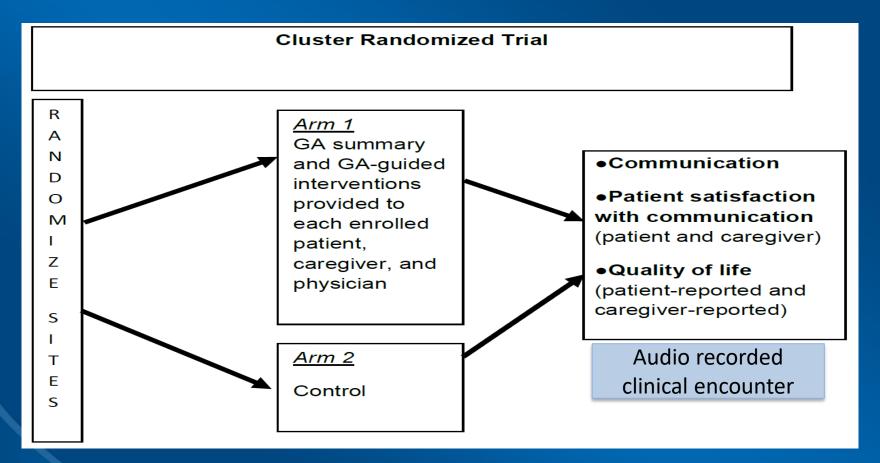


Caregivers

- One caregiver was chosen by the patient to enroll using the question:
 - "Is there a family member, partner, friend, or caregiver (age 21 or older) with whom you discuss or who can be helpful in health-related matters?"
- Caregivers not required for patients to participate
- Able to provide informed consent
- Able to read and understand English

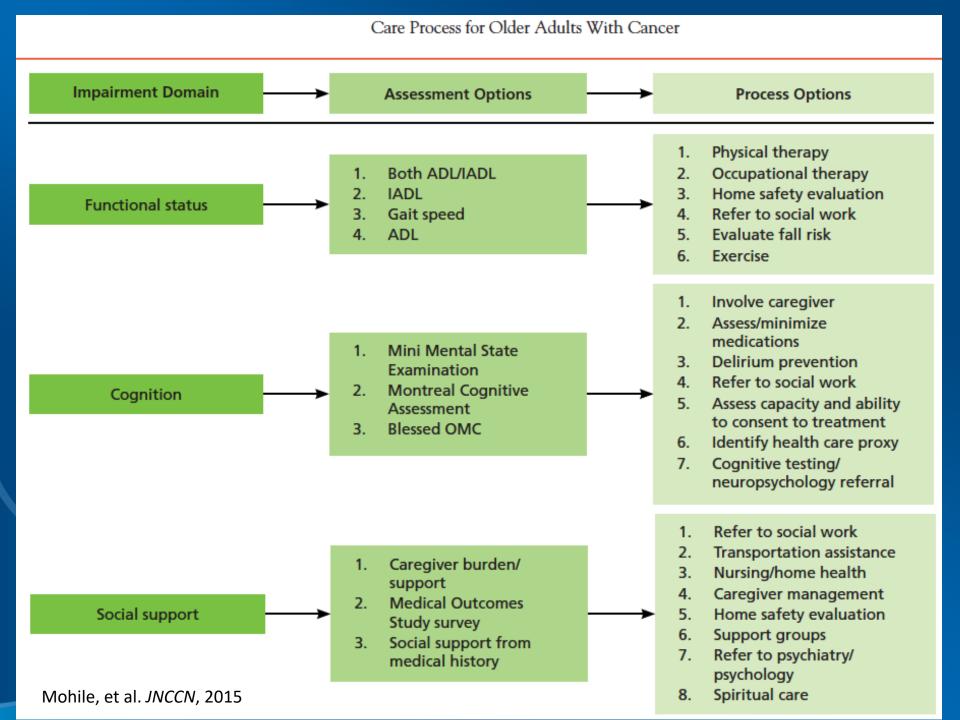


Study Design and Eligibility









GA-guided Communication Recommendations

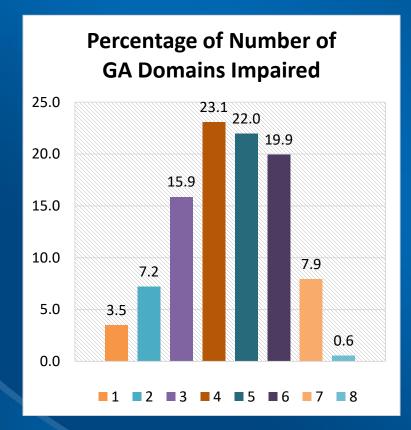
D:	d you or your staff complete any of the following with the	Comp	oleted	Not Con	Not Completed	
	d you or your staff complete any of the following with the tient during the clinic consultation (study visit)?	Yes	No	Not Appropriate	Patient Declined	
1.	Discuss patient's concerns about cognition.					
2.	Elicit input and perspectives from caregiver(s) about the patient's cognition.					
3.	Assess decision-making capacity.					
	 Elicit health care proxy information and input if the patient lacks decision-making capacity. 					
4.	Carefully weigh risks and benefits given limited data and potential for further cognitive impairment and functional impairment.					

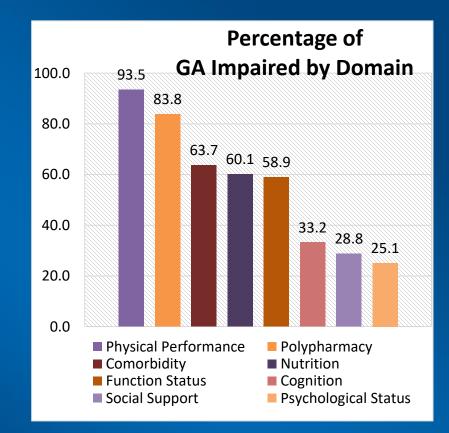




GA domains for all patients (N=541)

Mean age: 76.6 (Range 70-96)









Improving the Care of Older Adults with Cancer is Important: Oncologists Want Guidance

n=305 community oncologists	Agree (%)	Disagree (%)	Neutral (%)
I believe that the medical care of older adults with cancer needs to be improved	89%	3%	8%
I would appreciate additional training in topics related to the care of older adults with cancer	79%	4%	17%
I routinely ask my patients if they have a history of recent falls	70%	14%	16%
I frequently order home safety evaluations for my older patients	41%	35%	25%
I frequently enlist the help of a social worker	31%	37%	32%
I use standardized geriatric assessment tools to help me make decisions about my patients	23%	49%	29%

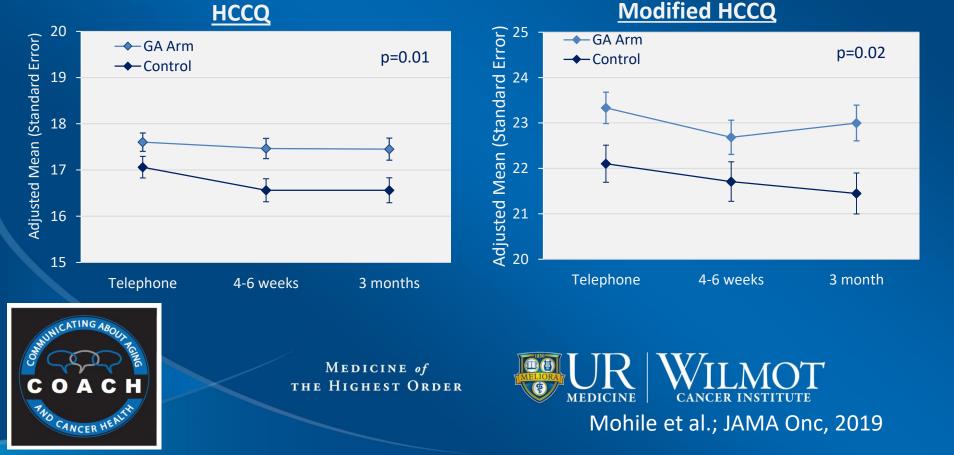
GA Improves Patient and Caregiver Satisfaction with Communication

-Health Care Climate Questionnaire

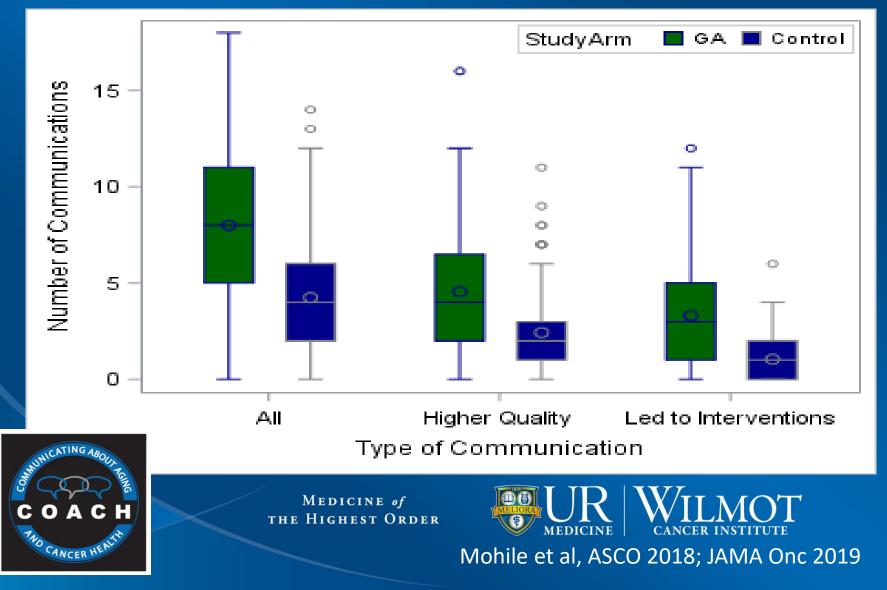
(5 questions, scale: 0-20)

-Health Care Climate Questionnaire modified for age-related concerns (modified)

(7 questions, scale: 0-28)



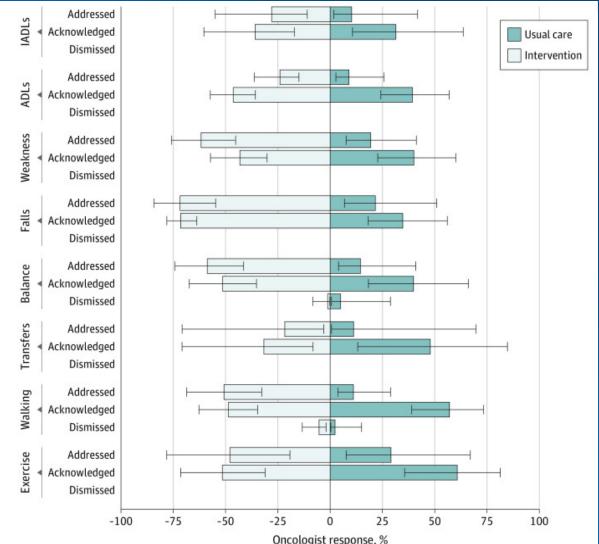
GA Improves Communication about Age-Related Concerns



Geriatric Assessment Improves Conversations about Geriatric Domains

Function and Physical Performance Conversations

Battaglia-Jensen et al., JNCCN 2022



Communication About Cognition

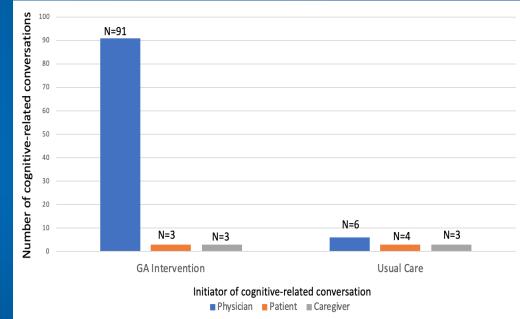
Of the 175 patients with abnormal cognitive screen:

- Mean age: 78 years
- Male: 54.3%
- >HS Education: 61.4%
- Condition:
 - 101=Intervention arm
 - 74=Usual Care

Conversations about cognition were more likely in the GA intervention arm

- 63.4% vs 12.1%; p<0.0001
- Majority oncologist initiated (overall 88.2%) MEDICINE of

THE HIGHEST ORDER



Patients randomized to the intervention were significantly more likely to have the cognitive conversation initiated by the oncologist (OR 20.22; 95% CI 3.66-111.78).





Magnuson, ASCO 2021

Quality of Cognitive Conversations

Oncologist discussing cognitive concerns with older patients with cancer and their caregivers. All patients in examples had an abnormal cognitive screening evaluation. P=Patient; C=Care Partner; D=Doctor

Example #1: Limited exploration of caregiver and patient cognitive concerns and how cognitive problems affect function and independence.

C: But her – her memory has not been real good before she had cancer to start with.
P: And now it's worse.
C: I'd noticed that, you know, couple years. And it's not getting any better.
D: Aha.

Example #2: Incorporating cognition as a potential treatment risk but limited integration of cognitive-related goals in the context of cancer treatment decision-making.

D: When you get sick, when you go through a lot of stress, particularly chemotherapy, this could get unmasked. Memory problems. Even dementia. This may be temporary because of the stresses you are going through but it becomes manifest whenever you are going through treatment...

P: That's not permanent, is it?

D. Yes, so that weakness that you have? You may actually lose track of time. Lose sleep. And then you may actually feel like you are confused sometimes but it should bounce back. It may get better.

Example #3: Caregiver uncertainty about expressing cognitive concerns.

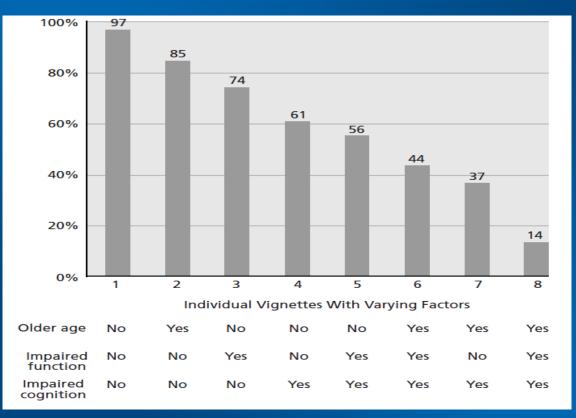
D (to caregiver): Okay, have you noticed anything? Gone bonkers on you? C: She's been – no. D: Okay.

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How does the Geriatric Assessment Influence Oncologist Decision-Making?

Percentage of Oncologists who Recommend Chemotherapy for an Older Vignette Patient with Advanced Pancreatic Cancer



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Mohile, Magnuson et al. JNCCN; 2018

Treatment Decisions

Patient	Comparison	OR (95%CI)	p-value	
Characteristic				

23% of oncologists surveyed reported using elements of geriatric assessment in clinical practice

		(1.56 to 5.60)	
Cognition	Not impaired vs impaired	8.56 (4.20 to 17.44)	<.0001*



GA to Reduce Treatment Toxicity



• Primary Aim:

 To evaluate if providing a GA summary with recommendations for management to oncologists reduces grade 3-5 toxicity (CTCAE) in patients aged 70+ starting a new regimen with chemotherapy and/or other agents which cause toxicity for advanced cancer

Secondary Aims:

- Survival at 6 months
- <u>Treatment decisions</u>
- Functional and Physical Performance

CTEP, NCI CTCAE v3.0 General Descriptions of Grade

0	No adverse event or within normal limits
1	Mild Adverse Event (minor; no specific medical intervention; asymptomatic laboratory findings only, radiographic findings only; marginal clinical relevance)
2	Moderate Adverse Event (minimal intervention; local intervention; noninvasive intervention [packing, cautery])
3	Severe and undesirable Adverse Event (significant symptoms requiring hospitalization or invasive intervention; transfusion; elective interventional radiological procedure; therapeutic endoscopy or operation)
4	Life-threatening or disabling Adverse Event (complicated by acute, life- threatening metabolic or cardiovascular complications such as circulatory failure, hemorrhage, sepsis. Life-threatening physiologic consequences; need for intensive care or emergent invasive procedure; emergent interventional radiological procedure, therapeutic endoscopy or operation)
5	Fatal adverse event

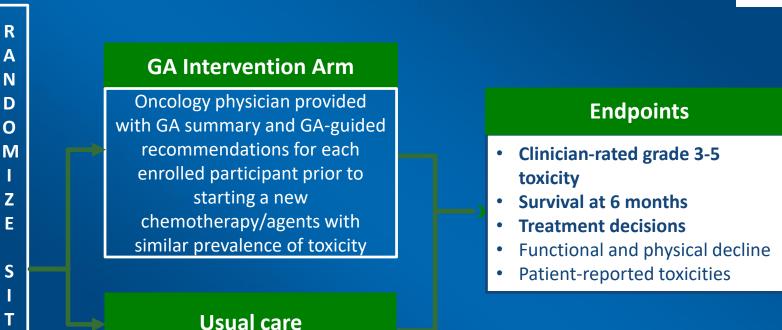
5 Fatal adverse event

PI: Mohile MEDICINE of THE HIGHEST ORDER Funding: NCI Mohile et al. ASCO, 2020; Lancet 2021



Study Schema Geriatric Assessment for Patients 70+





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Patient Eligibility and Characteristics

- Age ≥70 years
- Diagnosis of advanced solid tumor or lymphoma
- Have ≥1 GA Domain Impairment (other than polypharmacy)
- Starting a new chemotherapy regimen or another regimen with high risk of toxicity
- Have decision-making capacity, or, if not, oncologist has obtained consent from health-care proxy

	GA Intervention	Usual Care	
	N or Mean (% or SD)	N or Mean (% or SD)	P value
Age	77.2 (5.7)	77.2 (5.2)	0.98
Female	145 (41.5%)	166 (45.0%)	0.35
Race/Ethnicity			<0.01
Non-Hispanic White	281(80.5%)	347 (94.0%)	
African American	40 (11.5%)	12 (3.3%)	
Cancer Type			<0.01
Gastrointestinal	133 (38.1%)	114 (30.9%)	
Genitourinary	56 (16.0%)	53.0 (14.4%)	
Lung	63 (18.1%)	116 (31.4%)	
Stage IV	304 (87.1%)	324 (87.8%)	0.11
Cancer Treatments			0.53
Chemotherapy	305(87.4%)	328 (88.1%)	
Non-chemotherapy	44(12.6)	41(11.1%)	
Prior chemotherapy	104 (30.8%)	81 (22.7%)	0.02



Medicine of the Highest Order

Mohile et al. ASCO, 2020

DomainsToolsDescriptionsDefinitions of impairmentPrevalence of the most common GA-guided management recommendations chosen by oncologists in the intervention armPhysical performanceam Go"Assess mobility over 3 meters; longer time indicates worse performance>13.5-Conduct frequent toxicity checks (86-0%) - Provide information on exercise and exercise prescription (83.4%)(n=314/349 impaired in intervention arm)Short Physical PerformanceAssess balance, gait speed, and strength; higher score indicates better performance (range 0-12 points)> 9 points-Conduct frequent toxicity checks (86-0%) - Provide information on exercise and exercise prescription (83.4%)Falls HistoryAssess the number of fallsAny history of falls in the prior 6 monthsAny history of falls in the prior 6 monthsConduct frequent toxicity checks (86-0%) - Provide information on exercise and exercise prescription (83.4%)OARS Physical HealthAssess any limitation in activities (e.g. climbing several fights of stairs, walking more than a mile) as a result of his/her health (options: a lot, a litte, not at all)If the patient answered any question as "a lot"If the patient answered any question as "a lot"OARS (12.1%)Assess any limitation in activities (e.g. climbing several fights of stairs, walking more than a mile) as a result of his/her health (options: a lot, a litte, not at all)If the patient answered any question as "a lot"If the patient answered any question as "a lot" </th <th></th> <th></th> <th></th> <th></th> <th></th>					
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GA Intervention Management Recommendations Example: Physical Performance

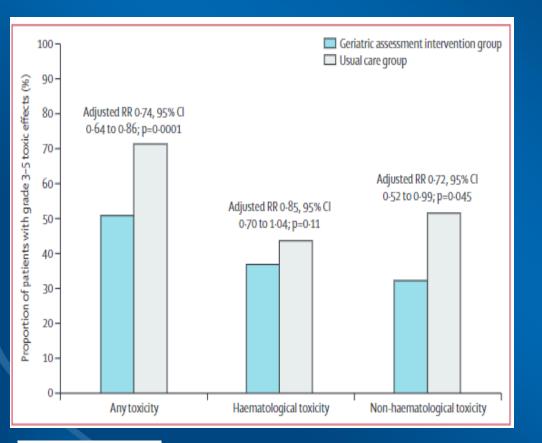
Decembra detione		nmended	Implemented?	Implemented by	
Recommendations	Yes	No	Enter Reason Code	Physician	Staff
1. Referrals:					
A. Physical Therapy (outpatient or home-based depending on eligibility for home care): request gait/assistive device evaluation, strength and balance training.					
B. Occupational therapy (if eligible for home care, OT referral to do safety evaluation).					
C. Aide services (SW may be able to assist).					
D. Personal Emergency Response information (PERS) especially if alone at any time while receiving treatment (SW may be able to assist).					



MEDICINE of THE HIGHEST ORDER



Any Grade 3-5 CTCAE Toxicity in 3 Months



- Any Grade 3-5 Toxicity
 Adjusted Risk Ratio: 0.74
 95% CI: (0.63-0.87), P < 0.01
 Clustering effect: P = 0.15
- Any Grade 3-5 Hematologic Toxicity Adjusted Risk Ratio: 0.85
 95%CI: (0.69-1.05), P = 0.13
 Clustering effect: P = 0.30
- Any Grade 3-5 Non-hematologic Toxic Adjusted Risk Ratio: 0.73
 95% CI: (0.53-0.996), P = 0.047
 Clustering effect: P < 0.01

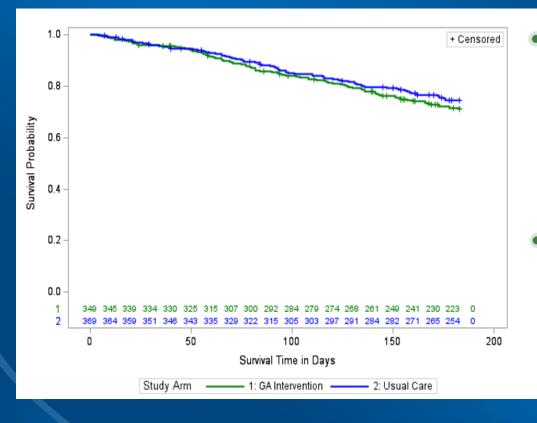


MEDICINE *of* the Highest Order



Mohile et al., Lancet 2021

Survival



Survival at 6 month based on Kaplan Meier Estimates GA Intervention: 71.3% (66.2%- 75.9%) Usual Care: 74.3% (69.5%-78.6%) P = 0.33

Adjusted Hazard Ratio: 0.87
 95% CI: (0.65-1.15), P = 0.33
 Clustering effect: P = 0.04



MEDICINE *of* the Highest Order



Mohile et al. ASCO, 2020; Lancet 2021

Mechanism

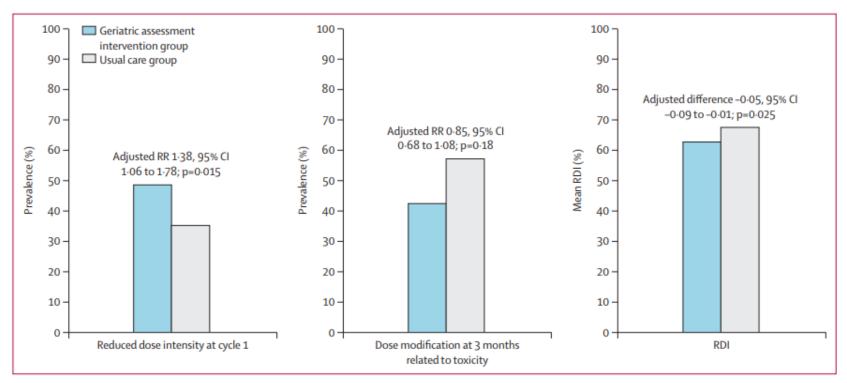


Figure 3: Treatment intensity by study group

(A) Prevalence of reduced treatment intensity at cycle 1. (B) Prevalence of dose modifications over 3 months. (C) RDI over 3 months. RDI=relative dose intensity. RR=risk ratio.

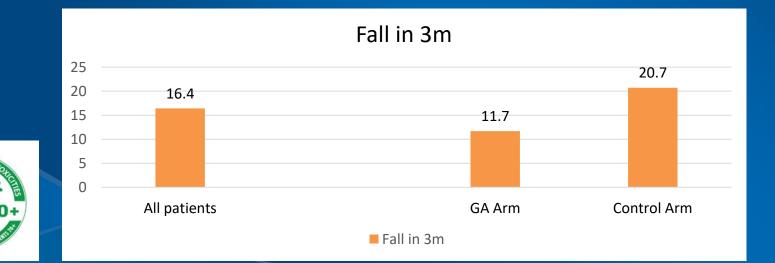






GA Intervention Improves Falls over 3 months

All Events: 103/627=16.4% GA Arm: 35/298=11.7% Control Arm: 68/329=20.7%	Risk Ratio	95% CI	P-value	P-value for cluster effect. Do results differ by site?
Unadjusted	0.58	(0.40-0.84)	0.004	N/A
Clustered Standard error	0.58	(0.45 - 0.75)	P<0.001	N/A
Bootstrap Standard error	0.58	(0.40 - 0.84)	0.004	N/A



Alternative Models of Care



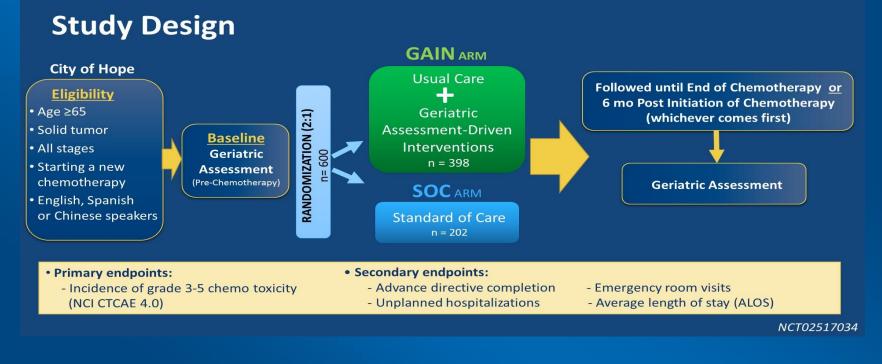


Infuse geriatrics principles indirectly

Direct involvement of geriatrics expertise in oncology care model



GAIN Trial



- •Prospective randomized trial
- •Single US cancer center
- •Enrolled patients aged 65+ with solid malignancy starting new chemo
- •Randomized patients to GAIN vs SOC

Li, et al. JAMA Oncology, 2021



Figure 2. Geriatric Assessment-Driven Intervention (GAIN) Used in This Study

Domain	Deficit		Interventions	
Functional status	 Limitations in activities of daily living or instrumental activities of daily livin History of falls Timed Up and Go >13 s Lack of energy 		ll risk y evaluation thening	GAIN GA and
Comorbidities	 Presence of comorbid conditions Hearing/visual impairments 	primary car • Referrals as		Interventions
Psychological status	 Feeling sad or depressed Anxiety Feeling nervous/worried 	 Social work Psychiatry i Psychology Chaplaincy Support pro 	referral referral referral	
Social activity Interference of physical or emproblems on social activity 		 Evaluation Social work Occupation 		
		Social support	 Lack of social support identified Patient lives alone 	 Counseling Social work referral Home safety evaluation Support programs Community resources
		Nutrition	 Weight loss ≥5% Body mass index ≤21 or ≥30 Problems with eating or feeding 	 Diet recommendations Supplements Oral care Physical/occupational therapy for food intake problems
		Cognition	 Abnormal cognitive screening Confusion Memory loss/impairment 	 Assess decision-making capacity Involve caregivers Review of medications Delirium prevention Cognitive testing
Li, et al. JAN Oncology, 2		Polypharmacy	 ≥5 Prescribed medications ≥1 Over-the-counter medication ≥1 Herb/vitamin supplement 	 Recommendations regarding drug interactions, potentially inappropriate medications, duplicati medications

GAIN GA and rventions

regarding drug interactions, opriate medications, duplicative

Results: Primary Aim

Incidence of Grade 3-5 Chemotherapy-Related Toxicity p=0.02 70.0% 60.0% GAIN 50.0% 60.4% 40.0% p=0.008 SOC p=0.003 30.0% 50.5% p=0.61 26.2% 20.0% 19.3% 21.1% 18.1% 14.9% 10.0% 11.3% 0.0% **Overall Toxicity Heme Toxicity** Non-Heme **Both Heme and** Only **Toxicity Only** Non-Heme Toxicity The GAIN arm had a statistically significant reduction of 9.9% (95% CI: 1.6-18.2%, **p=0.02**) in chemo-related toxicity compared to the SOC arm

Li, et al. JAMA Oncology, 2021



Geriatric Assessment-Driven Interventions \downarrow Toxicity Risk

GAIN STUDY (n = 605) – JAMA Oncology

Key finding:

- GA-driven INterventions (GAIN) reduced grade 3+ chemo-related toxicity (↓10%)
- Improved advance directive completion in older adults with cancer (^{24%}) compared to standard of care (SOC).
- Survival no different at 12 months.

GAP70+ STUDY (n = 718) – Lancet

Key finding:

- A GA summary with management recommendations provided to the oncologist prior to the start chemo reduces grade 3+ toxicity (↓20%) over 3 months.
- Survival no different at 6 months.
- Falls and polypharmacy \downarrow .

Li, et al. JAMA Oncology, 2021 MEDICINE of THE HIGHEST ORDER



GA systematic review and meta-analysis

17 RCTs (6 including toxicity outcome)

Risk of treatment toxicity:

	CGA	1	Usual	are		Risk Ratio	Risk Ratio
Study or Subgroup	Study or Subgroup Events Total		Events	Total	Weight	M-H, Random, 95% CI	M-H, Random, 95% Cl
Li 2021	203	402	123	203	33.2%	0.83 [0.72 to 0.97]	-
Lund 2021	20	71	28	71	4.5%	0.71 [0.45 to 1.14]	
Magnuson 2018	21	37	21	34	6.5%	0.92 [0.62 to 1.35]	
Mohile 2021	177	349	263	369	42.5%	0.71 [0.63 to 0.80]	
Nadaraja 2020	9	48	17	46	2.1%	0.51 [0.25 to 1.02]	
Puts 2022	54	153	65	162	11.3%	0.88 [0.66 to 1.17]	
Total (95% CI)		1060		885	100.0%	0.78 [0.70 to 0.86]	•
Total events	484		517				
Heterogeneity: Tau² =	0.00; Ch	i ^z = 5.83	2, df = 5 (/	P = .32)	;/ ² = 14%	, <u>For</u>	
Test for overall effect:	Z=4.92	(P < .00	001)			0.01	0.1 1 10 100

No differences observed in mortality, hospitalization, early treatment discontinuation, initial or subsequent dose reduction

MEDICINE of the Highest Order

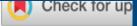


Anwar, et al. JNCI, 2023

Study	Design	Population	Intervention Delivery	Management Strategy	Outcomes
Mohile et al. -community oncology practices affiliated with University of Rochester NCORP Research Base	Cluster randomization by oncology practice COACH: n=542; GAP n=700	age 70+ with advanced solid tumor malignancies	GA summary results and recommendation given to oncology team	Established protocol based on Delphi consensus panel and guidelines	COACH: Communication, Satisfaction; GAP70+: Chemo toxicity (Gr3+), Survival, Function POSITIVE
Li et al. -City of Hope	2:1 Patient randomization n=600	age 65+ with any stage solid tumor malignancies starting a new chemo regimen (any line)	Study NP in collaboration with the primary oncologist and clinic nurse to follow up	Established protocol based on multidisciplinary team input and triggers based on GA results	GAIN: Endpoints: Chemo toxicity (Gr3+); Rate of hospitalization; Change in functional status; Change in psychosocial status POSITIVE
Soo et al. -mult-center study in Australia	1:1 Patient randomization N=150	age 70+ with cancer planned for chemotherapy, targeted therapy, and immunotherapy	Geriatrician-led longitudinal co- management	Clinical expertise	QoL; function, mood, nutrition, health utility, treatment delivery, healthcare utilization and survival POSITIVE
Puts et al. -multi-center study of centers in Canada	Patient randomization n=350	age 70+ with most solid tumor malignancies starting first/second line chemotherapy	Geriatric oncology with nurse follow up	Established protocol based on Delphi consensus and guidelines	QoL; Cost-effectiveness; Function; Chemo tox; Satisfaction; Cancer tx changes; Survival NEGATIVE

ASCO Guideline update

ASCO Special Articles



Practical Assessment and Management of Vulnerabilities in Older Patients Receiving Systemic Cancer Therapy: ASCO Guideline Update

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Dale, et al. JCO, 2023

MEDICINE of THE HIGHEST ORDER





PROs, Symptom science



caregiver



GA implementation in oncology



Polypharmacy, Data Science



GA-guided cancer treatment modifications



Cancer-related cognitive decline, dementia and cancer interface

Digital health interventions, exercise interventions, decision-making



ED-based cancer care interventions



GA in survivorship (among everything else)



Aging biomarkers,

health equity,

frailty interventions

Where are

we going?



Communication, advanced care planning



Mental health and social support interventions



Social support, treatment tolerability

University of Rochester Geriatric Oncology Research Program



https://www.urmc.rochester.edu/labs/geriatric-oncology.aspx







MEDICINE of THE HIGHEST ORDER