

Virtual Section Meeting

May 13, 2020 7 p.m. ET

Vulnerability of Older Adults to COVID-19: Emerging Role for Biological Hallmarks of Aging and Geroscience-Guided Therapies

Dr. George A. Kuchel, MD, FRCP, AGSF, FGSA





Panel Discussion

- James Kirkpatrick, MD, FACC- Chair
- Karen Alexander, MD, FACC- *Immediate Past*Chair
- Mathew Maurer, MD, FACC
- Michael Rich, MD, FACC



Abstract Presentations and Discussion

Scott Hummel, MD
Jon Afilalo, MD, FACC



Congratulations!

Lina Brinker, MD

Hospitalist and Clinical Lecturer University of Michigan

Polypharmacy, Multimorbidity and Therapeutic Competition in HFpEF #13057





Congratulations!

Kamil Faridi, MD

Assistant Professor of Medicine Section of Cardiovascular Medicine Yale School of Medicine New Haven, CT

Assessing Frailty-Associated Characteristics and Outcomes in the Dual Antiplatelet Therapy (DAPT) Study Using Medicare Claims: Insights from the EXTEND-DAPT Study #16839





Polypharmacy, Multimorbidity and Therapeutic Competition in HFpEF

Lina M. Brinker MD, Mary E. Tinetti MD, Matthew C. Konerman MD, Cristen J. Willer PhD, Jennifer L. McNamara MS, Scott L. Hummel MD MS, and Parag Goyal MD MSc

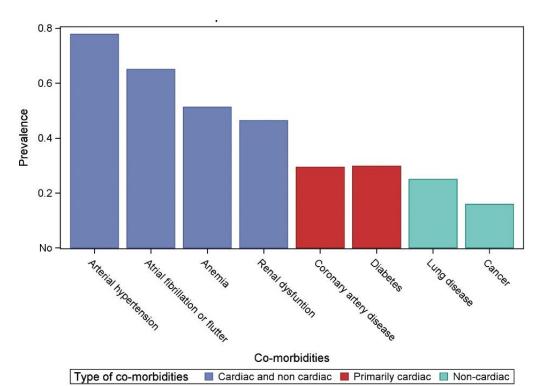


Disclosures - None

Background I

Heart failure with preserved ejection fraction (HFpEF): Challenging to manage

- Complex condition with high burden of comorbidities
- Primarily a geriatric disease (mean age >70 in most studies)
- Inconsistent diagnostic criteria
- Lack of interventions that improve outcomes
- Medication complexity further complicates these issues



Background II

Medication complexity in HFpEF

- Given the high prevalence of comorbidities, cardiac and noncardiac medications are common
- This leads to complex medication regimens, which predispose older adults to adverse outcomes
- Several factors contribute to medication complexity, including:
 - 1. Polypharmacy: taking multiple medications (we defined as \geq 10)
 - 2. Potentially inappropriate medications (PIMs): medications with risks that may outweigh benefits—especially in older adults
 - 3. Therapeutic competition: a type of drug-disease interaction
- Medication complexity—well-described in the geriatrics literature—has not been studied in HFpEF

Objective

To investigate potentially harmful medication patterns in older adults with HFpEF by characterizing polypharmacy, PIMs, and therapeutic competition

Methods I

- Study population: 231 adults seen in a specialized HFpEF clinic at the University of Michigan from 7/2016 - 9/2019
- Chart review to characterize the following:
 - Comorbidity burden
 - Polypharmacy
 - Use of PIMs, defined 3 ways
 - Therapeutic competition



Methods II: PIMs

1. AHA HF-exacerbating agents:

Medications considered by the American Heart Association to increase the risk of causing or worsening heart failure, per their 2016 Scientific Statement

Drugs That May Cause or Exacerbate Heart Failure

A Scientific Statement From the American Heart Association

Common examples include:

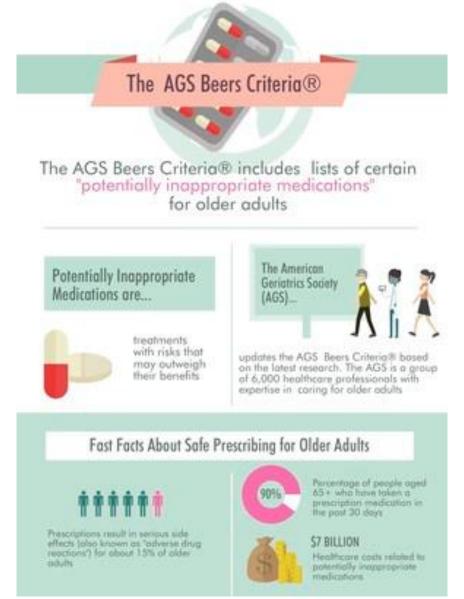
- Non-dihydropyridine CCBs
- Metformin
- Sulfonylureas
- Thiazolidinediones
- DPP-4 inhibitors
- NSAIDs
- Citalopram



Methods III: PIMs

2. Beers agents:

- Medications strongly recommended against using in older adults by the American Geriatric Society's Beers Criteria 2019 update
- Common examples include:
- NSAIDs
- Sulfonylureas
- Thiazolidinediones
- Tricyclic antidepressants
- First generation antihistamines (diphenhydramine, hydroxyzine)
- Nonselective alpha-1 blockers (terazosin, doxazosin, prazosin)



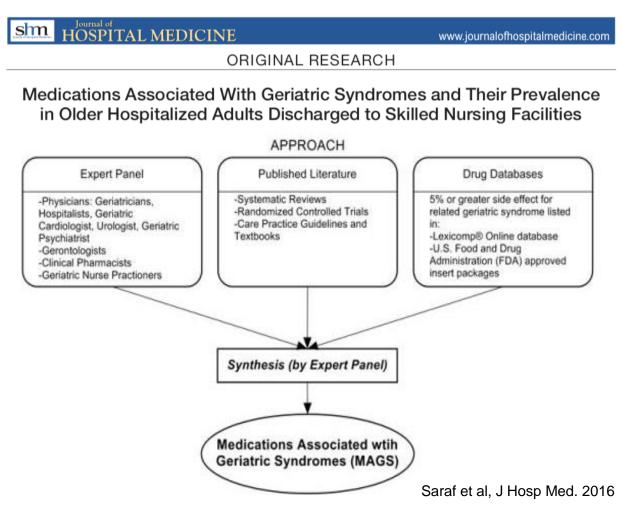
American Geriatrics Society

Methods IV: PIMs

3. Medications associated with geriatrics syndromes (MAGS)

Medications which may contribute to one or more of these common geriatric syndromes:

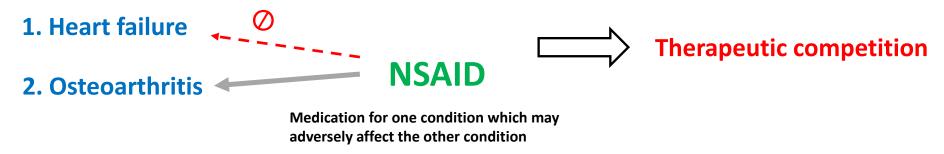
- Falls
- Urinary incontinence
- Delirium
- Reduced appetite or weight loss
- Depression
- Cognitive impairment



Methods V: Therapeutic competition

• Defined as a type of drug-disease interaction in which the treatment for one condition may adversely affect a coexisting condition

Competing conditions:



OPEN & ACCESS Freely available online



Potential Therapeutic Competition in Community-Living Older Adults in the U.S.: Use of Medications That May Adversely Affect a Coexisting Condition

Songprod Jonathan Lorgunpai¹, Marianthe Grammas², David S. H. Lee³, Gail McAvay², Peter Charpentier², Mary E. Tinetti^{2,4}*

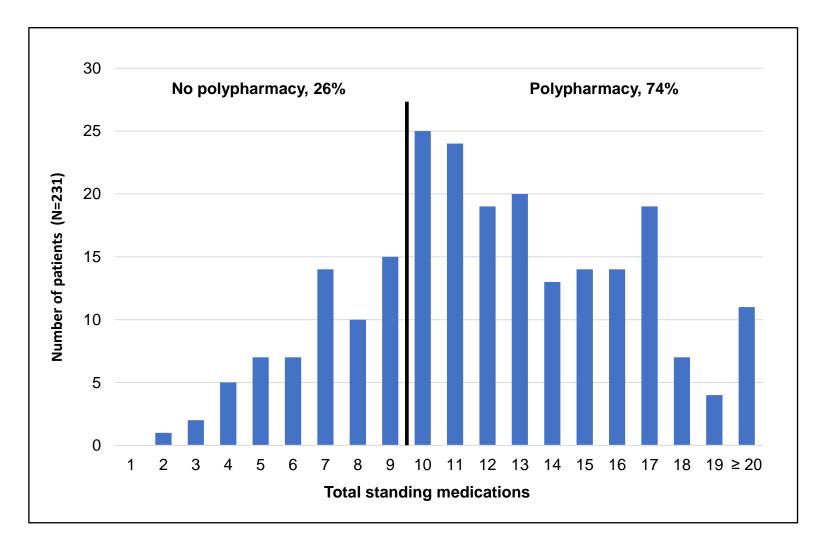
Results I: HFpEF patient characteristics

Demographics				
Age in years, mean ± SD	68 ± 12			
Age ≥ 65 years, n (%)	146 (63)			
Female, n (%)	147 (64)			
White, n (%)	209 (90)			
Count of comorbid conditions, median (IQR)	7 (5-8)			
Prevalence of comorbid conditions, n (%)				
Hypertension	191 (83)			
Hypertension Obesity	191 (83) 167 (72)			
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Obesity	167 (72)			
Obesity Obstructive sleep apnea	167 (72) 156 (68)			

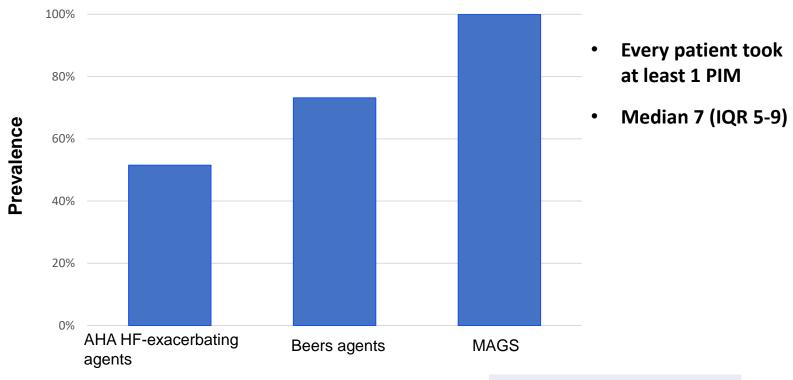
Prevalence, cont'd				
Atrial fibrillation/flutter	105 (45)			
Gastroesophageal reflux disease or peptic ulcer disease	102 (44)			
Chronic kidney disease	98 (42)			
Coronary artery disease	86 (37)			
Osteoarthritis	86 (37)			
Hypothyroidism	69 (30)			
Depression	62 (27)			
Chronic obstructive pulmonary disease	62 (27)			
Osteoporosis	27 (12)			
Benign prostatic hyperplasia	12 (5)			
Dementia or cognitive impairment	9 (4)			
Cirrhosis	7 (3)			

Results II: Polypharmacy

- 170 of 231 (74%) were taking 10 or more medications (polypharmacy)
- Mean 12 ± 4, Median 12 (IQR 9-15)



Results III: PIMs



AHA HF-exacerbating agents, n (%)			
Metformin	43 (19)		
Non-dihydropyridine CCB	26 (11)		
Citalopram or escitalopram	18 (8)		
Sulfonylurea	16 (7)		
NSAID	16 (7)		
Hydroxychloroquine	13 (6)		
Albuterol	9 (4)		
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Beers agents, n (%)				
Proton pump inhibitor	115 (50)			
Non-dihydropyridine CCB	26 (11)			
Sulfonylurea	16 (7)			
NSAID	16 (7)			
Amiodarone	13 (6)			
Sliding scale insulin	11 (5)			
Paroxetine	9 (4)			
_				

MAGS, n (%)	
Loop diuretic	186 (81)
Beta blocker	153 (66)
ACEI/ARB	112 (48)
Calcium channel blocker	62 (27)
Selective serotonin reuptake inhibitor	60 (26)
Insulin	54 (23)
Thiazide diuretic	53 (23)

Results IV: Therapeutic competition

- 81% had at least one pair of competing health conditions
- Among those with a pair of competing conditions, 49% were taking a medication which could worsen one condition
- Inhaled beta agonists were most common

Most common combinations of possible therapeutic competition:

Condition pairs, n (%)			Medication causing therapeutic competition, n (%)	
HTN	COPD	55 (24)	Beta agonist	31 (13)
HF	DM	112 (48)	Beta blocker with alpha-blocking activity	20 (9)
HTN	DM	106 (46)	Beta blocker with alpha-blocking activity	19 (8)
HF	COPD	62 (27)	Nonselective beta blocker, or Beta blocker with alpha-blocking activity	14 (6)
HTN	COPD	55 (24)	Nonselective beta blocker, <i>or</i> Beta blocker with alpha-blocking activity	13 (6)
CAD	COPD	28 (12)	Beta agonist	13 (6)
GERD	Osteoporosis	20 (9)	Proton pump inhibitor	14 (6)

Limitations

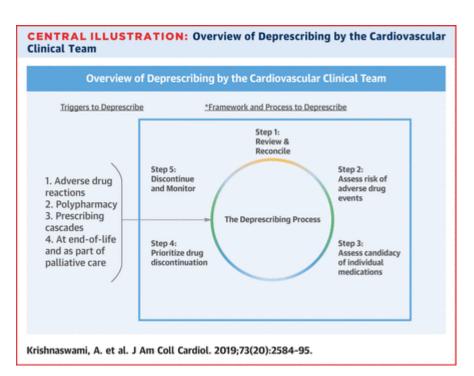
- Single-site observational study
- Medication and comorbidity data were collected from the electronic medical record and may be incomplete or inaccurate
- We included only scheduled medications

Conclusions

- Polypharmacy is highly prevalent, affecting around 75% of HFpEF patients
- Additional factors that contribute to medication complexity include:
 - Universal PIM use
 - Therapeutic competition, which is more prevalent in HFpEF than in the general geriatric population

Implications

- Need for patient-centered approaches to manage medication complexity in HFpEF, including:
 - Developing strategies to address medication burden in this primarily geriatric population
 - Considering how PIMs are defined in the setting of HFpEF, for example:
 - There is some data that Metformin may actually be beneficial in HFpEF
 - Some patients require loop diuretics to manage their symptoms
 - Developing approaches for addressing therapeutic competition: medications with benefits > risks should be continued, those with risks > benefits discontinued
- Formal processes to maximize medication reconciliation and deprescribing should be broadly incorporated into the care of HFpEF patients:
 - From specialized HFpEF programs to primary care



Thank you!

- Dr. Parag Goyal, Weill Cornell Medicine
- Dr. Scott Hummel, University of Michigan
- Jen McNamara, MS, UofM Cardiovascular Center clinical research manager
- linab@med.umich.edu
- @LinaBrinker

Discussion





Assessing Frailty-Associated Characteristics and Outcomes in the Dual Antiplatelet Therapy (DAPT) Study Using Medicare Claims:
Insights from the EXTEND-DAPT Study

Kamil Faridi, MD

Assistant Professor of Medicine

Section of Cardiovascular Medicine Yale School of Medicine New Haven, CT

ACC Geriatric Cardiology Section Virtual Meeting May 13th, 2020







Financial Disclosures

- I have no relevant disclosures
- Robert Yeh (PI) has investigator-initiated research grants from Abbott Vascular, Abiomed, AstraZeneca, BD Bard, Boston Scientific, Cook Medical, Edwards Life Sciences, Medtronic, and Philips.

Background

- Frailty is a state of decreased physiologic reserve with increased vulnerability to stressors, and is associated with increased mortality¹
- Frailty is not frequently assessed in observational studies or in randomized clinical trials of patients with coronary artery disease (CAD)
- Nearly 1 in 5 patients ≥65 years old with CAD and nearly 1 in 3 patients with acute coronary syndrome (ACS) have evidence of frailty^{2,3}

¹Bergman H et al. J Gerontol A Biol Sci Med. 2007.

²Singh M et al. Circ Card Qual Outcomes. 2011.

³Bebb et al. Eur Heart J Acute Card Care. 2018.

Background

- Many of the risks and benefits of therapies for CAD in frail patients remain uncertain
- Frail patients with CAD are less likely to receive therapies such as cardiac catheterization or medications for secondary prevention¹
- Older, frailer patients have increased short-term risk of bleeding and bleeding-related death²⁻⁴

¹Afilalo J et al. J Am Coll Cardiol. 2014.

²Alonso GL, et al. Int J Cardiol. 2016.

³Perera V, et al. Age Ageing. 2009.

⁴Dodson JA, et al. JACC Card Interv. 2018.

Background

- Dual antiplatelet therapy (DAPT) with ASA + P2Y12 inhibitor is recommended for 6 to 12 months following PCI, and continuation of DAPT for longer can be considered based on ischemic and bleeding risks¹
- The long-term impact of frailty on adverse clinical events for patients who undergo PCI and receive DAPT is unknown
- Use of administrative claims data in clinical trials may help us better assess risks and benefits associated with frailty

Study Objectives

- Assess whether frailty-associated characteristics derived from claims data are associated with adverse clinical events in patients after PCI
- Determine whether frailty-associated characteristics impact treatment effects of extended duration DAPT after PCI

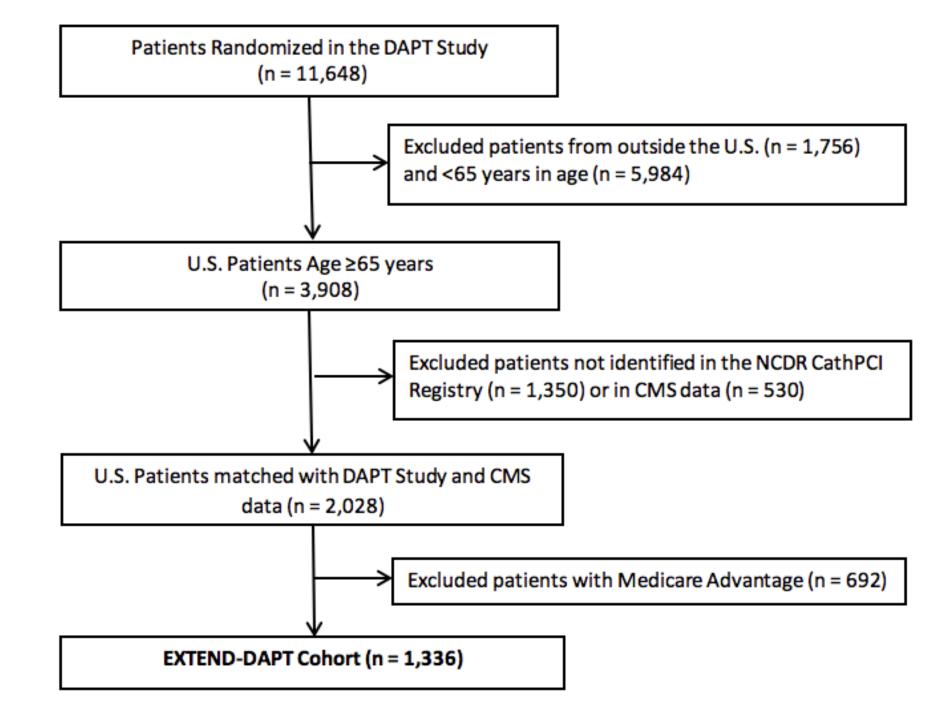
Methods

- DAPT Study¹
 - Randomized, placebo-controlled clinical trial comparing outcomes for patients who received 30 vs. 12 months of DAPT following PCI
 - Patients randomized at 1 year following PCI if no ischemic or bleeding events occurred following PCI
 - Extended duration DAPT reduced the risk of MACCE events but also increased risk of bleeding

Methods

 DAPT Study data was linked with Medicare claims data as part of the Extending Trial-Based Evaluations of Medical Therapies Using Novel Sources of Data (EXTEND) Study

 Deterministic algorithms were used to match patients ≥65 years old with the NCDR CathPCI Registry linked to Medicare claims



Methods

- Frailty-Associated Characteristics
 - Used inpatient ICD-9 diagnosis codes associated with frailty in prior studies^{1,2}
 - Patients analyzed based on number of frailty characteristics

¹Segal JB, et al. Med Care. 2017.

²Kim DH, et al. J Gerontol A Biol Sci Med Sci. 2018.

Frailty characteristics based on ICD-9 codes:

- impaired mobility or falls
- hospital beds and associated supplies
- wheelchairs and accessories
- walking aids and attachments other supplies
- transportation services including ambulance
- ill-defined and unknown causes of morbidity
- psychotic conditions
- hereditary/degenerative diseases of the central nervous system
- other psychoses
- neurotic and other nonpsychotic mental disorders
- open wound of lower limb
- contusion with intact skin surface
- chronic skin ulcer
- diseases of veins/lymphatics
- arthropathies and other musculoskeletal disorders
- other bacterial diseases
- pneumonia and influenza
- skin and soft tissue infections
- mycoses
- other diseases of the urinary system

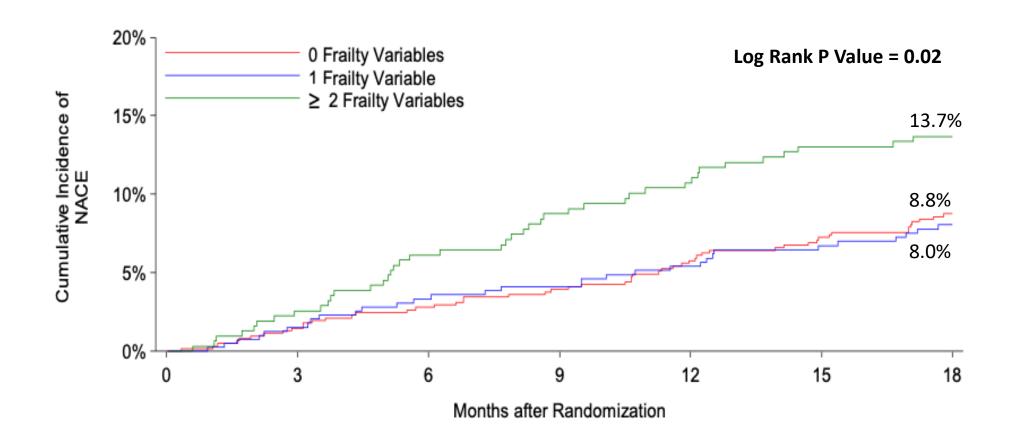
Methods

- Outcomes
 - Net adverse clinical events (NACE; composite of death, MI, stroke, and bleeding)
 - Death
 - MI
 - Stroke
 - Bleeding
- Rates of adverse outcomes and treatment effects of extended duration DAPT were determined based on presence of frailtyassociated characteristics

Results – Baseline Characteristics

Characteristic	0 Frailty Characteristics (n = 614)	1 Frailty Characteristic (n = 397)	≥2 Frailty Characteristics (n = 315)	P value
Age (mean, SD)	71.3, 5.2	72.1, 5.5	72.5, 5.8	0.003
Female	26.1	32.7	46.3	<0.001
Nonwhite Race	5.7	7.6	7.1	0.485
BMI (mean, SD)	29.5, 5.0	29.5, 5.3	29.9, 5.9	0.462
Diabetes Mellitus	32.8	33.6	34.0	0.928
Hypertension	84.6	84.4	87.5	0.422
Current/Recent Smoking	8.1	15.0	12.8	0.002
Prior Stroke/TIA	4.4	5.8	7.0	0.244
History of Major Bleeding	0.8	0.8	1.6	0.461
Prior PCI	33.2	39.3	33.3	0.103
Prior CABG	16.2	15.7	18.4	0.579
Prior Myocardial Infarction (MI)	21.3	22.3	18.5	0.483
Atrial Fibrillation	4.8	5.8	3.5	0.348
Index PCI - STEMI	3.7	4.5	3.8	0.807
Index PCI - NSTEMI	11.4	10.1	11.1	0.799
Index PCI – Unstable Angina	15.6	13.4	16.5	0.457
Index PCI – Stable Angina	44.1	45.6	45.7	0.859

Results – Cumulative Incidence of NACE Based on Frailty-Associated Characteristics



Results – Cumulative Incidence of Events Based on Frailty-Associated Characteristics

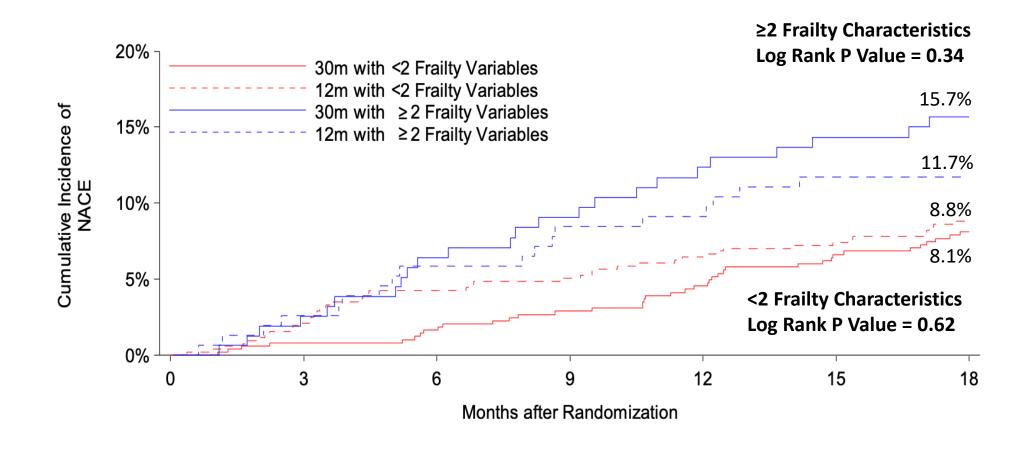
Outcome	0 Frailty Characteristics	1 Frailty Characteristic	≥2 Frailty Characteristics	Log Rank P Value
NACE	8.8	8.0	13.7	0.02
Death	2.3	1.8	3.9	0.19
MI	3.8	2.9	4.6	0.48
Stroke	0.5	2.4	2.0	0.03
Bleeding	4.0	3.4	6.0	0.21

Results – Adjusted* Risks of Adverse Outcomes by Number of Frailty-Associated Characteristics

Outcome	1 Frailty-Associated Characteristic HR [95% CI]	P value
NACE	0.92 [0.60-1.39]	0.68
Death	0.76 [0.32-1.79]	0.53
MI	0.81 [0.43-1.56]	0.53
Stroke	3.73 [1.18-11.81]	0.03
Bleeding	0.93 [0.50-1.71]	0.81
Outcome	≥2 Frailty-Associated Characteristics HR [95% CI]	P value
Outcome NACE		P value 0.03
	HR [95% CI]	
NACE	HR [95% CI] 1.57 [1.06-2.33]	0.03
NACE Death	HR [95% CI] 1.57 [1.06-2.33] 1.72 [0.82-3.60]	0.03 0.15

^{*}The following variables were included in adjusted analyses: age, sex, total number of lifestyle cardiovascular risk factors [diabetes, hypertension, smoking, and obesity], presence of an additional cardiovascular risk factor (any one of the following variables: prior PCI, prior CABG, prior stroke or TIA, prior MI, or MI at time of index PCI), and DAPT Study arm (30 or 12 months of DAPT)

Results – Cumulative Incidence of NACE Based on Frailty-Associated Characteristics and Duration of DAPT



Results – Cumulative Incidence of Outcomes for Patients with < 2 Frailty-Associated Characteristics Based on DAPT Duration

Outcome	30 Months <2 Frailty Ch.	12 Months <2 Frailty Ch.	Log Rank P Value
NACE	8.1	8.8	0.62
Death	2.5	1.8	0.44
MI	3.3	3.5	0.83
Stroke	1.1	1.4	0.61
Bleeding	4.4	3.2	0.34

Results – Cumulative Incidence of Outcomes for Patients with ≥ 2 Frailty-Associated Characteristics Based on DAPT Duration

Outcome	30 Months ≥2 Frailty Ch.	12 Months ≥2 Frailty Ch.	Log Rank P Value
NACE	15.7	11.7	0.34
Death	5.2	2.6	0.24
MI	4.0	5.3	0.59
Stroke	2.0	2.0	1.00
Bleeding	8.7	3.3	0.06

Results – Risk of Outcomes for 30 vs. 12 Months of DAPT by Number of Frailty-Associated Characteristics

Outcome	30 vs. 12 Mos. DAPT <2 Frailty Characteristics HR [95% CI]	30 vs. 12 Mos. DAPT ≥2 Frailty Characteristics HR [95% CI]	Interaction P value
NACE	0.90 [0.59-1.38]	1.35 [0.73-2.48]	0.28
Death	1.40 [0.59-3.32]	2.01 [0.61-6.68]	0.63
MI	0.93 [0.47-1.82]	0.75 [0.26-2.16]	0.74
Stroke	0.74 [0.24-2.35]	1.00 [0.20-4.94]	0.77
Bleeding	1.27 [0.69-2.33]	2.38 [0.92-6.20]	0.08

Limitations

- ICD codes have not been previously used to directly define frailty
- Results could differ with use of ICD-10 codes
- Analyses may be underpowered
- Inclusion and exclusion criteria may limit generalizability
- All treatment effects should be considered exploratory

Conclusions

- Frailty-associated characteristics in claims data were associated with higher NACE in a subgroup of older patients in the DAPT Study
- No differences in treatment effects for extended duration DAPT following PCI were seen for patients with increased frailty
- However, our findings suggest further study in this area is needed
- This study supports use of claims data to augment risk prediction and better assess heterogeneity of treatment effects in cardiovascular clinical trials

Acknowledgements

Co-investigators: Harun Kundi, Jordan B. Strom, Jeptha Curtis,
 Qi Gao, Yang Song, Hector Tamez, Laura Mauri, Changyu Shen

PI: Robert Yeh

 American College of Cardiology National Cardiovascular Data Registry (NCDR)

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Thank you!

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Discussion



FIT and Early Career Opportunities

Parag Goyal, MD, MSc, FACC



Opportunities for FITs and ECPs in Geriatric Cardiology

Parag Goyal MD, MSc, FACC
Chair, FIT/ECP Geriatric Cardiology Section WG
Assistant Professor of Medicine, Weill Cornell Medicine
pag9051@med.cornell.edu

My Early Career Path

- 2015: Joined ACC GCS
- 2016: Get Going RCT, ECCOA/JGC
- 2017: U13 Conference/Travel Award



- 2018: JACC Council Perspective
- 2018: Started HFpEF Program for the Aging
- 2019: Submitted Beeson application



ACC Geriatric Cardiology Section as a Resource



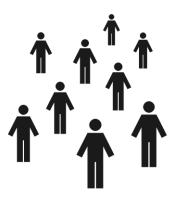
Exposure to Geriatric Cardiology content/expertise



Career development opportunities



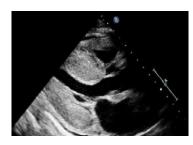
Acquisition of skills and knowledge



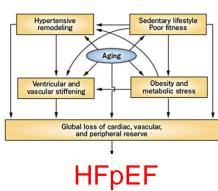
Network of sponsors and mentors

The Time is Now for Geriatric Cardiology





Cardiac Amyloidosis





Palliative Care

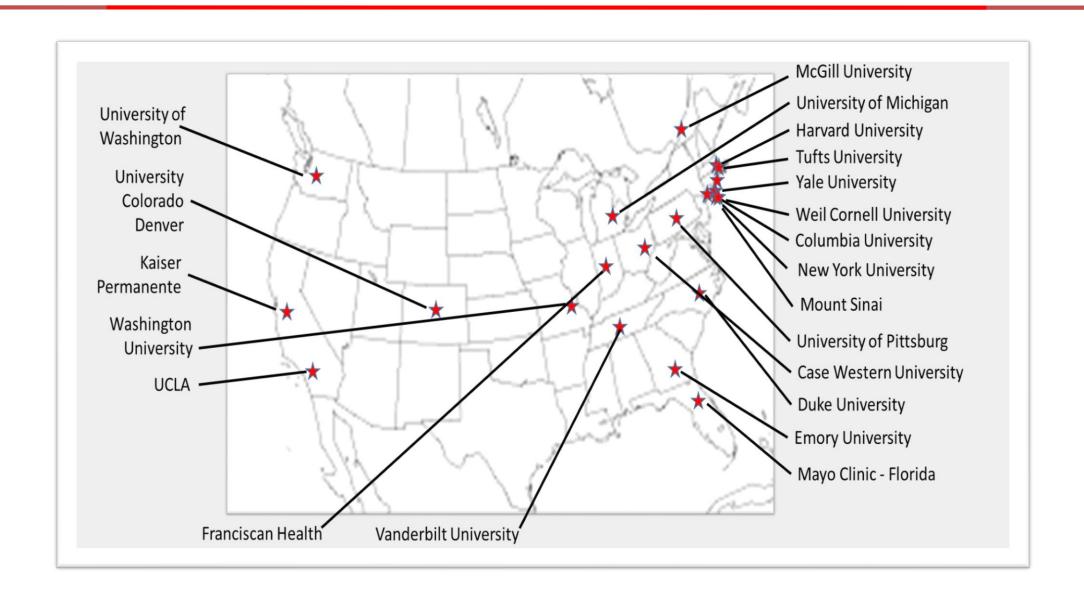


Polypharmacy/Deprescribing



Statins for Primary Prevention

A Growing Field/Network



How to get involved?

- ACC GCS Membership
 - Sign-up for membership
- ACC GCS FIT/ECP Working Group
- ACC Microvolunteering
 - Sing up via Membership Home Page
- ACC GCS FIT/ECP Working Group Leadership Panel
 - Call for applications



ACC GCS FIT/ECP Leadership Panel

Purpose:

- To engage and cultivate the careers of trainees and junior faculty with an interest in Geriatric Cardiology.
- Through this position, Representatives will have the opportunity to actively participate in ACC, work with thought leaders in Geriatric Cardiology, and advance the field to ultimately improve the care of older adults with cardiovascular disease

• Eligibility:

- Current FIT or ECP (within 5 years of training)
- Interest in joining the Research WG, Palliative Care WG, or ACC.org Editorial WG

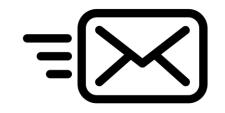
ACC GCS FIT/ECP Leadership Panel

Responsibilities:

- Participate in Working Group activities, which include regular conference calls and various ad-hoc activities intended to advance the field of Geriatric Cardiology.
- Serve on the ACC Geriatric Cardiology Section
 FIT/ECP Working Group Leadership Panel

(INSERT LINK FOR APPLICATION HERE)

Thank you!



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@ParagGoyalMD



Virtual Reception and Discussion

- Please use the chat box to the right of your screen
- We will attempt to unmute participants when possible
- Email <u>mdisch@acc.org</u> with any issues or follow up questions





James N. Kirkpatrick, MD, FACC
Chair, ACC Geriatric Cardiology Section Leadership Council

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Email: kirkpatj@cardiology.washington.edu

Social media: @Kirkpatj1

Clinical/research interests: Cardiac palliative care, cardioethics, end of life management of cardiac devices,

advance care planning for older adults.



Karen P. Alexander, MD, FACC
Immediate Past Chair, ACC Geriatric Cardiology Section Leadership Council

Institution: Duke University/Duke Clinical Research Institute

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Clinical/research interests: Frailty assessment, clinical registries, outcomes in older patients with CV disease considering geriatric conditions.





Aaron J. Bagnola, PharmD, BCPS, BCCP

Member, Geriatric Cardiology Section Leadership Council

Institution: Inova Fairfax Medical Campus

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Clinical/research interests: Cardiac Critical Care, Advanced Heart Failure, and

Pulmonary Hypertension



Susan Cheng, MD

Member, Geriatric Cardiology Section Leadership Council



Kelsey M. Flint, MD, FACC
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Scott L. Hummel, MD

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Richard A. Josephson, MS, MD, FACC, FAHA, FACP, FAACVPR

Member, Geriatric Cardiology Section Leadership Council

Institution: University Hospitals Health System & Case Western

Reserve University

Email: Richard.Josephson@UHhospitals.org

Clinical/research interests: Secondary prevention of CVD and bio-behavioral correlates

of CVD in older adults, Cardiac Rehab.





Maureen B. Julien, MSN, CPNP, AACC Member, Geriatric Cardiology Section Leadership Council



Ashok Krishnaswami, MBBS, FACC
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Radmila Lyubarova, MD, FACC
Member, Geriatric Cardiology Section Leadership Council





John P. Mulrow, MD, FACC
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Michael W. Rich, MD, FACC

Member, Geriatric Cardiology Section Leadership Council



Eiran Gorodeski, MD, MPH, FACC Member, Geriatric Cardiology Section Leadership Council

Social media: @EiranGorodeski



Join Us!

The Geriatric Cardiology Member Section is only able to accomplish its mission through support from members.

Visit ACC.org Member Sections page today or email mdisch@acc.org to join!

