ENROLLING SOON:

cvMOBIUS Registry

EPIC message: Research MHIF Patient Referral

CONDITION:

Recent ASCVD Event

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Amgen

DESCRIPTION:

The purpose of cvMOBIUS (Cardiovascular Multi-dimensional Observational Investigation of the Use of PCSK9 Inhibitors) is to evaluate the effectiveness of PCSK9 inhibitors to reduce cardiovascular events among subjects with a recent ASCVD event or revascularization procedure.

While large randomized trials have shown additional lipid-lowering through PCSK9i can further reduce risk of ASCVD events, realworld effectiveness of PCSK9i in subjects with ASCVD events has yet to be established.

CRITERIA LIST/ QUALIFICATIONS:

Inclusion:

- ≥ 40 v.o.
- Hospitalization for a clinical ASCVD event (acute MI, unstable angina, IS or CLI) within 18 months of enrollment and/or coronary, peripheral, or carotid revascularization including percutaneous or surgical revascularization in the past 18 months
- LDL ≥ 70 mg/dL with no immediate plans for statin change or newly started on PCSK9i after index hospitalization/procedure (no more than 6 months prior to enrollment)

Exclusion: ESRD, on a PCSK9i prior to qualifying event





Equity in Cardiovascular Health Outcomes-Facts, Figures, and MHIF Strategies to Close the Gap

> Courtney Jordan Baechler, MD, MS Mosi Bennett, MD, PhD Mario Goessl, MD, PhD





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Creating a World without Heart and , Vascular Disease...

FOR **ALL**





Context: Racial Disparities in Health

- African Americans have higher death rates than Whites for 12 of the 15 leading causes of death.
- Blacks and American Indians have higher agespecific death rates than Whites from birth through the retirement years.
- Hispanics have higher death rates than whites for diabetes, hypertension, liver cirrhosis & homicide
- Minorities get sick younger, have more severe illness and die sooner than Whites





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Historical Perspective

- Health disparities between blacks and whites since first settlers arrived
- Tuskegee Syphilis Trials
- 1990's University study on "genetic etiology of aggressive behavior"
- 2002 IOM *Unequal Treatment* disparities in health care delivery less likely to be given appropriate cardiac meds, CABG
- 2004—systemic review of angiography, angioplasty, CABG, and lytics-21/23 showed that African Americans were less likely to get CABG





What is Race?

"Pure races in the sense of genetically homogenous populations do not exist in the human species today, nor is there any evidence that they have ever existed in the past... Biological differences between human beings reflect both hereditary factors and the influence of natural and social environments. In most cases, these differences are due to the interaction of both."

American Association of Physical Anthropology, 1996





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Racial and Ethnic Disparities

Blacks have the highest rates of heart disease in the country

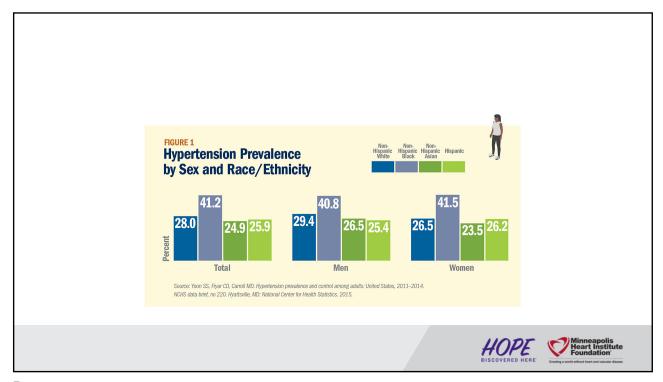
Blacks are 2-3 times more likely to die from heart disease

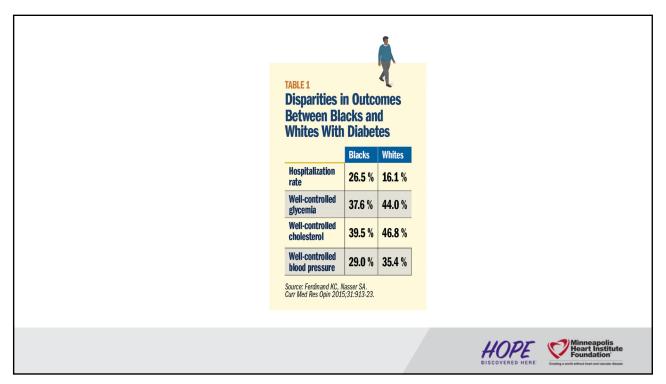
While the cardiovascular mortality rates have been decreasing, this is not seen for blacks

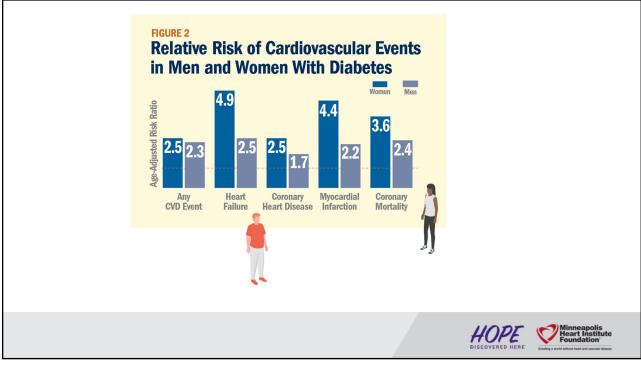
Blacks were 42% less likely to receive an ICD in the Medicare population after a heart attack











Black and Hispanic women have the highest rates of obesity in the country

Black and Hispanic women have the highest rates of diabetes and high blood pressure

Women and blacks are less likely to get a cardiac catheterization when they present with chest pain (then men and whites)





Atrial Fib Outcomes

- Blacks and Hispanics have a lower incidence of atrial fib than whites
- However, blacks are less likely to be aware they have the condition
- · Higher overall risk of stroke and stroke mortality in black patients
- · Greater risk of major bleeding with warfarin
- · No major difference noted with DOAC's







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Heart Transplant Wait Times

2014 Outcomes:

- 19.8 months average wait time for African-Americans
- 12 months for white patients
- 12.3 months for Hispanic patients

2016 Outcomes:

- 10.4 months for African-Americans
- 8 months for white patients
- 7.4 months for Hispanic patients







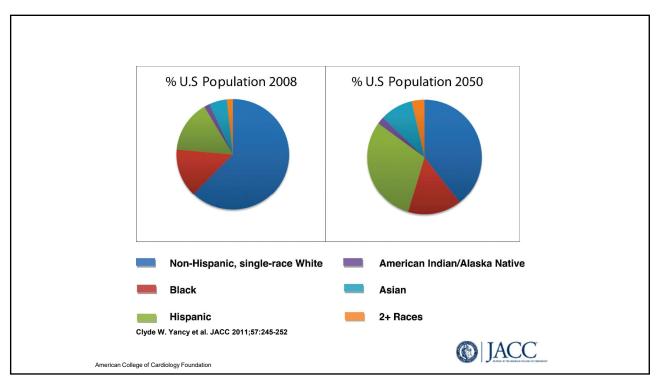
The Coalition to Reduce Racial and Ethnic Disparities in Cardiovascular Disease Outcomes (*credo*)

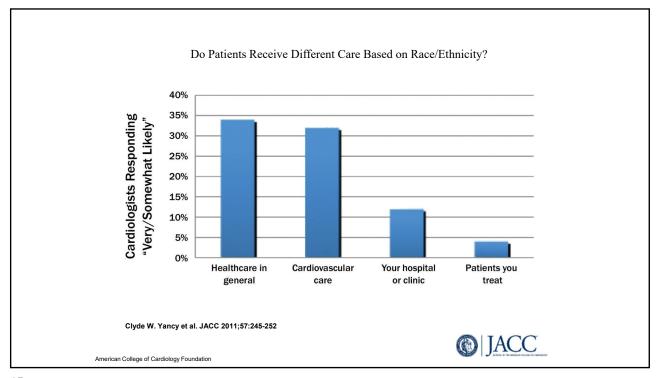
- · Launched in 2009
- Help the cardiology community meet the needs of an increasingly diverse patient population
- · Evidenced-based tools
- · Performance improvement data
- · Provider education including cultural competency training
- · Patient education approaches
- · Goal was equitable care and outcomes for all patients, regardless of race, ethnicity, sex, and age

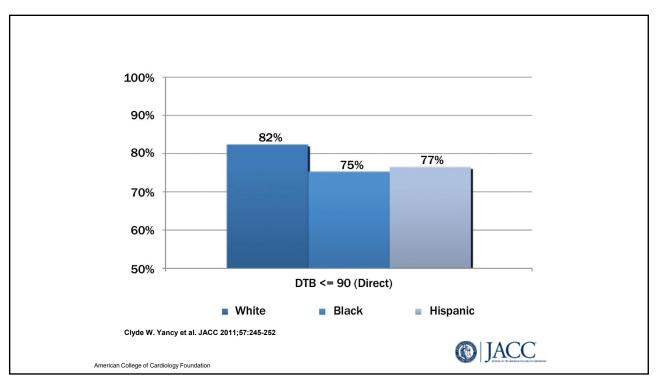




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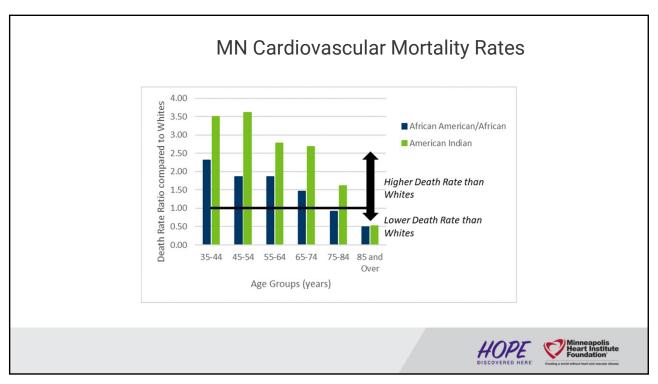


MN has had the best cardiovascular mortality rates since 1999....





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Cardiovascular Mortality Rates in MN

Age Group (years)	Crude Death Rate (per 100,000), Whites	Crude Death Rate (per 100,000), African Americans/ Africans	Rate Ratio of African Americans/Afric ans to Whites	Crude Death Rate (per 100,000), American Indians	Rate Ratio of American Indians to Whites
35-44	14.4	33.3	2.31	50.5	3.51
45-54	46.1	85.6	1.86	166.7	3.62
55-64	101.1	188.6	1.87	280.5	2.77
65-74	220.8	322.6	1.46	590.8	2.68
75-84	726.3	665.7	0.92	1175.3	1.62
85 and Over	3286.3	1637.5	0.50	1714.0	0.52





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Controlling High Blood Pressure by Race and Ethnicity, Minnesota Health Care Program Members, 2017

Race	Rate Comparison to State Rate		
American Indian/Alaskan Native	69%	No difference*	
Black/African American	57%	Lower	
Asian	72%	No difference*	
White	74%	Higher	
Ethnicity	Rate	Comparison to State Rate	
Hispanic	74%	No difference*	





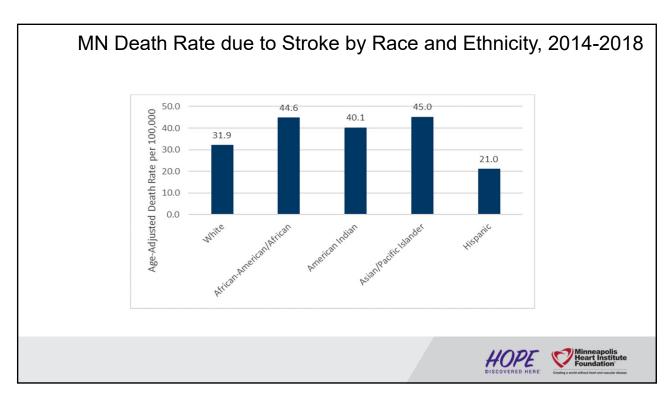
Optimal Vascular Care Goals among Minnesota Adults with Ischemic Vascular Disease by Race, 2017

Race/Ethnicity	% Meeting all four goals	
American Indian	45%	
Asian	68%	
African American/African	45%	
Multi-Racial	50%	
Native Hawaiian/Other Pacific Islander	55%	
White	63%	





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What is Health Equity?

"Health equity is a state where all persons, regardless of race, creed, income, sexual orientation, gender identification, age or gender have the opportunity to reach their full health potential without the limits of structural barriers."

 Minnesota Department of Health, "Advancing Health Equity in Minnesota: Report to the Legislature." 2014.





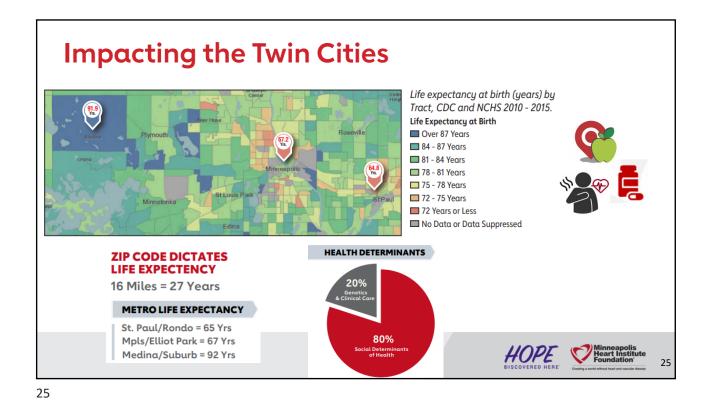
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A Tale of Two Cities

https://youtu.be/Eu7d0BMRt0o







MDH directed by Legislature in 2013 to prepare a report on Advancing Health Equity in Minnesota
Report to the Legislature

MDH directed by Legislature in 2013 to prepare a report on Advancing Health Equity in MN

1. To provide an overview of MN's health disparities and health inequities

2. To identify inequitable conditions that produce health disparities

3. To make recommendations to advance HE in MN

Statewide Health Assessment

- Shows a picture of health and well-being across the state, including:
 - Who is healthy and who is not?
 - What conditions shape health for all the different populations in Minnesota?
 - What do we have, and what do we need, to assure that all people in Minnesota can enjoy healthy lives and healthy communities





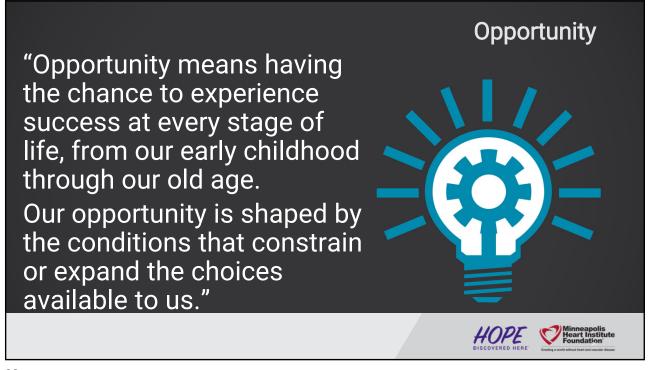
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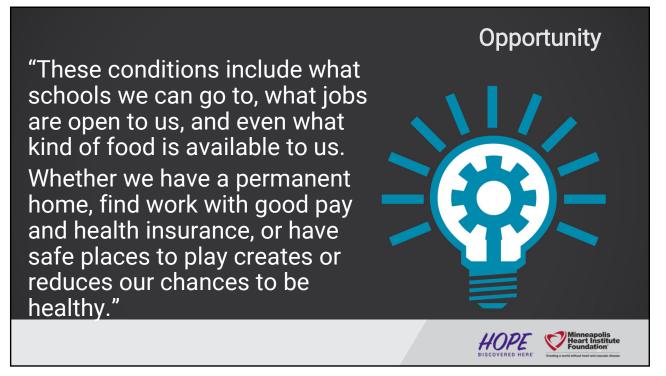
People: highlights

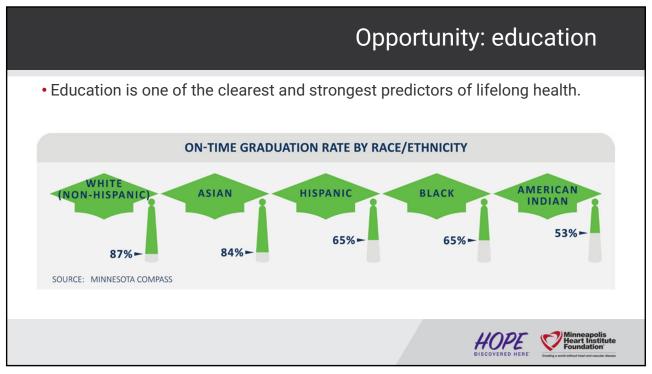
- About 14% of all children in Minnesota live in poverty.
- About 9% of Minnesotans 18-64 have a disability; almost 1 in 5 families with children have a child with special health needs.
- Racial and ethnic diversity is expected to increase to about 25% by 2035.
- The LGBTQ population in Minnesota faces many challenges and barriers to health.
- The population over 65 is growing rapidly.

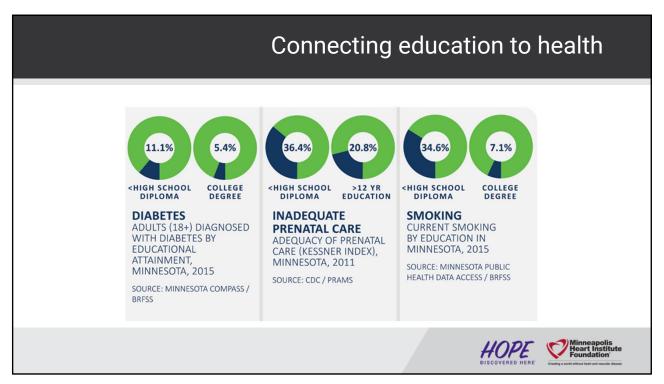


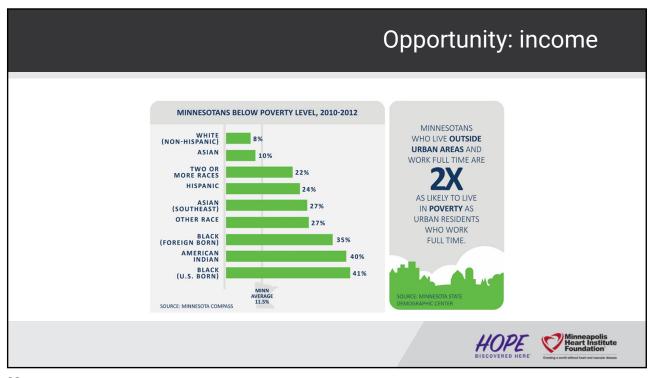


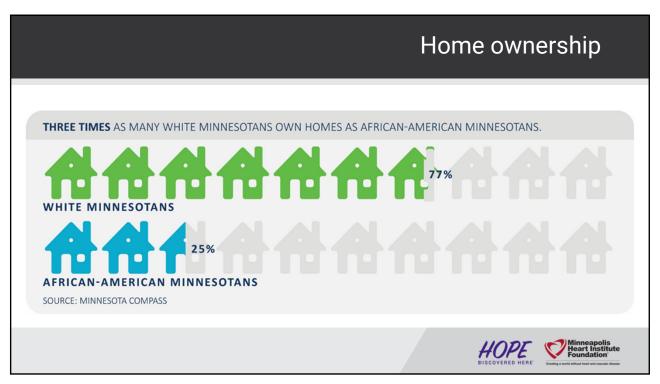












Connecting income, housing and health

Financial stress about housing, adults 18-64 only	Usually or always	Sometimes	Rarely or never
Ever had cancer (other than skin cancer)	8%	4%	4%
Ever had COPD1	10%	4%	2%
Ever had arthritis	29%	16%	14%
Ever had a depressive disorder	49%	24%	15%
Ever had diabetes	9%	5%	5%
Currently have asthma	14%	9%	6%
Currently smoke cigarettes	39%	23%	14%
Report binge drinking in past 30 days	25%	24%	22%
Are obese	35%	30%	24%

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Inequities in recreational opportunity

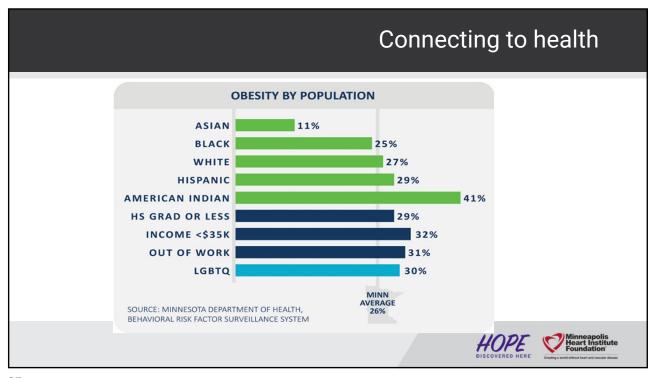


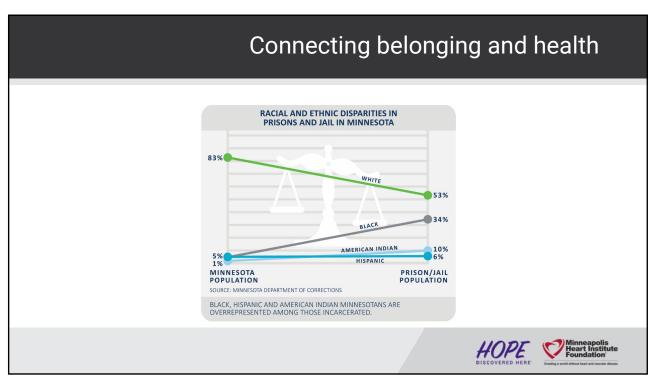
POPULATIONS OF COLOR ARE MORE LIKELY TO NOTE SAFETY CONCERNS ABOUT BEING IN REGIONAL PARKS.

SOURCE: METROPOLITAN COUNCIL









Consistency in opportunity inequities

 Populations of color and American Indians in Minnesota experience consistently lower opportunities in education, employment, income, housing, transportation, paid leave, health insurance, health care





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Minnesota Maternal Mortality Review Committee

Work on behalf of the Commissioner of Health to review all pregnancy- associated deaths of Minnesota residents.

<u>Purpose:</u> The Minnesota Maternal Mortality Review Committee (MMMRC) is tasked with addressing maternal mortality in Minnesota. The MMMRC works to identify factors contributing to maternal deaths and the health inequities impacting maternal health in the state. Leads the charge of disseminating recommendations it improve maternal outcomes for our Minnesota mothers.

<u>Vision</u>: The vision of the Minnesota Maternal Mortality Review Committee is to eliminate preventable maternal deaths, reduce maternal morbidities,, and improve population health and health equity for women of reproductive age in Minnesota.





Identifying Racism and Discrimination as Contributing Factors

- Women of color report more experiences of discrimination, food insecurity, and depression
- Women of color experience higher levels of chronic stress during pregnancy - results in compromised endocrine and immune function
- Burden remains higher across all income and education levels
- Results in greater rates of hypertensive disorder, preterm birth, low birth weight neonates and perinatal mortality among Black women

Grobman W479-99.Am J Perinatol. 2016 Dec;33(14):1426-1432.; Slaughter-Acey JC. Womens Health Issues. 2013 Nov-Dec;23(6):e381-7.; Borders AE. J Perinatol. 2015 Aug;35(8):580-4.; Mendez DD. Ethn Health. 2014;19(5):





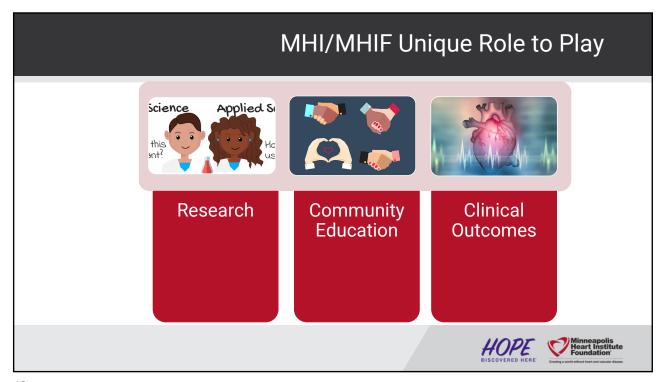
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NEW* Contributing Factors

- **DISCRIMINATION** Treating someone less or more favorably based on the group, class or category they belong to resulting from biases, prejudices, and stereotyping. It can manifest as differences in care, clinical communication and shared decision-making. (Smedley et al, 2003 and Dr. Rachel Hardeman)
- INTERPERSONAL RACISM Discriminatory interactions between individuals based on differential assumptions about the abilities, motives, and intentions of others and resulting in differential actions toward others based on their race. It can be conscious as well as unconscious, and it includes acts of commission and acts of omission. It manifests as lack of respect, suspicion, devaluation, scapegoating, and dehumanization. (Jones, CP, 2000 and Dr. Cornelia Graves)
- STRUCTURAL RACISM The systems of power based on historical injustices and contemporary social factors that systematically disadvantage people of color and advantage white people through inequities in housing, education, employment, earnings, benefits, credit, media, health care, criminal justice, etc. (Adapted from Bailey ZD. Lancet. 2017 and Dr. Carla Ortique)











The Insight2Health Fitness Challenge (I2H) inspires lifestyle changes in participants through fitness, yoga and nutrition and life coaching.

I2H introduces lifestyle changes that are sustainable. To date, we have hosted 14 I2H sessions with more than 300 participants who have lost more than 1,100 pounds and 430 inches.

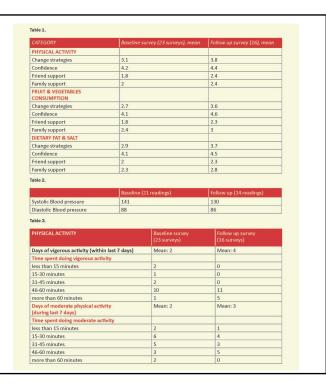
A few factors that have supported the program's success: 1) providing solid programming led by experienced personnel, 2) sharing participant success stories through Insight News and 3) receiving technical and financial support from organizations such as NorthPoint Health & Wellness.

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The interventions from the physical activity portion of the program appear to have had an overall positive effect. Participants became more physically active than they were at baseline and more motivated to continue physical activity as part of lifestyle changes.



"Based on the identified health concerns, our team would recommend emphasizing the importance of living a healthy lifestyle and control of risk factors for heart disease prevention especially in the African American community. Utilizing the medical community for any educational tools on these topics could be a way to better inform the participants of these diseases."



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Demonstrated Past Success

- Level 1 STEMI program
- HONU
- Creating strong relationships in the community
- · National reputation in research





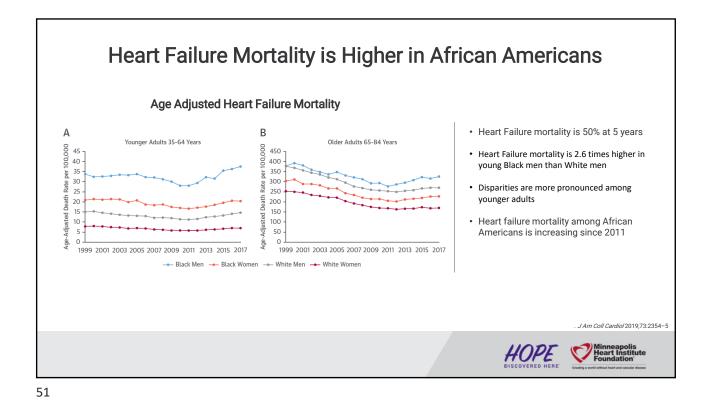
Dr. Mosi Bennett MD, PhD





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Heart Failure is More Prevalent in African-Americans Atherosclerosis Risk in Communities Study: **Annual incidence Heart Failure** 40.0 35.0 31.4 More than 6.5 million people with heart failure in 30.0 1,000 Person Years 26.2 25.0 Risk of heart failure increases with age for both 20.0 17.9 sexes and all races 15.0 11.2 Risk of heart failure is highest in African-Americans 10.0 Black men age 55-64: 3 times the risk of heart failure compared to white men 65-74 55-64 Age (years) ■White Males ■Black Males ■White Females ■Black Females Virani SS, Alonso A, Benjamin EJ, et al. Heart Disease and Stroke Statistics—2020 Update: a report from the American Heart Association. *Circulation* 2020;141:e139-e596.



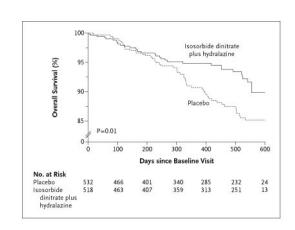
African Americans are Poorly Represented In Heart Failure Clinical Trials Overlay plot showing the trend in total patient enrollment vs % of • In a review of 25 randomized clinical trials for heart failure black patients enrolled over time • 19 for pharmacotherapies Total Study Population % Black • 6 for implantable cardioverter defibrillators 12000 - 100 · Among these studies 10000 78,816 patients 4,640 Black patients (5.9%) 8000 60 • Median Black participation per trial was 162 patients 6000 40 · Overall patient enrollment among the 25 trials increased while percentage of black patients decreased over time 2000 · Black patients are poorly represented among pivotal trials · Inclusion is necessary to ensure that study findings can be 1985 2000 2010 generalized to all patients with Heart Failure Year Total Study Population —

Black Individuals are Poorly Represented Among Trials for **Chronic Heart Failure** ELITE II (2000)39 67 (2.1%) patient enrollment (%) Val-HeFT (2001)⁴⁰ Study (year) [ref.] V-HeFT I (1986)²⁹ Veterans 18-75 y of age Ejection fraction <45% 180 (28.0%) NYHA functional class III-IV y of age with prior myocare Ejection fraction ≤30% NYHA functional class I-IV 8 y of age who were intolern Ejection fraction ≤40% NYHA functional class III-IV tients ≥18 y of age on an A Ejection fraction ≤40% MADIT-II (2002)⁴² NEIM ICD vs 1232 102 (8.3%) 73 (3.6%) CHARM-Added (2003)⁴⁴ 2548 127 (4.9%) NEIM Enalapril vs placebo 401 (9.48%) NEJM 217 (19.8%) NEIM 4800 897 (1.5.5%) Patients ≥18 y of age Ejection fraction ≤35% NYTHA functional class II-III Ejection fraction ≤40% NYTHA functional class III 55 y of age or ≥18 y of age Ejection fraction <40% NYTHA functional class III Patients ≥21 y of age Ejection fraction ≤30% NYTHA functional class III therapy ICD vs SCD-HeFT (2005) 46 NEJM 2521 425 (16.9%) BEAUTIFUL (2008)⁴⁸ ICD vs ICD-CRT MERIT-HE (1999)³⁸ The Lance 208 (5.2%)

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African-American Heart Failure Trial (A-HeFT)

- Subgroup analysis of earlier V-HeFT trial suggested combination therapy with isosorbide dinitrate plus hydralazine was beneficial in black patients
- A-HeFT: compare isosorbide dinitrate/hydralazine with placebo among black patients with advanced heart failure
- 1050 patients, EF 24% NYHA Class III, 18 months follow up
- Mortality was lower in the combination therapy group
- Survival differences at six months after randomization



Taylor AL, Ziesche S, Yancy C, et al. Combination of isosorbide dinitrate and hydralazine in blacks with heart failure. N Engl J Med 2004;351:2049-57.





Lessons Learned From the A-HeFT Trial

- Among black patients with advanced heart failure, treatment with isosorbide dinitrate plus hydralazine improves survival and reduces hospitalizations
- Inclusion of a group historically under represented in clinical trials ultimately led to the approval
 of a therapy with a specific survival benefit
- Current guidelines recommend the addition of nitrates and hydralazine for African American heart failure patients that are on optimal medical therapy

Taylor AL, Ziesche S, Yancy C, et al. Combination of isosorbide dinitrate and hydralazine in blacks with heart failure. N Engl J Med 2004;351:2049-57.





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Disparity in Access to Heart Failure Care

32018 BY THE AMERICAN COLLEGE OF CARDIOLOGY FOUNDATION

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VOL. 6, NO. 5, 2018

African Americans Are Less Likely to Receive Care by a Cardiologist During an Intensive Care Unit Admission for Heart Failure



Khadijah Breathett, MD, MS,* Wenhui G. Liu, PHD,* Larry A. Allen, MD, MHS,* Stacie L. Daugherty, MD, MSPH,* Irene V. Blair, PHD,* Jacqueline Jones, PHD, RN,* Gary K. Grunwald, PHD,* Marc Moss, MD,* Tyree H. Kiser, Phasub,** Blene Burnham, MD,* R. William Vandivier, MD,* Brendan J. Clark, MD, MS,* Eldrin F. Lewis, MD, MPH,* Sula Mazimba, MD, MPH,* Catherine Battaglia, PHD, RN,** P. Michael Ho, MD, PhD,** Pamela N. Peterson, MD, MSPH**

- Black Patients with Heart Failure are less likely to receive care by a cardiologist in the ICU
- ICU Care by a cardiologist is associated with better in-hospital survival

J Am Coll Cardiol HF 2018;6:413-20





Clinical Bias in Heart Failure

- A 2019 Study: 400 heart failure health care professionals
- · Randomized to a patient vignette about either an African-American or White man with identical profiles
- · Participants rated the appropriateness for heart transplant or LVAD
- The most important factors contributing to the decision to recommend heart transplant LVAD were social support and adherence
- · African-American patient with the same profile: less trustworthy, less social support, worse adherence
- · Heart transplant was recommended more often for the White patient





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Disparities in Heart Failure Care at MHI?

- Black patients at Minneapolis Heart Institute:
 - 11% of heart failure hospitalizations
 - · 14% of advanced heart failure hospital consults
 - 5% of clinic visits for heart failure
 - · 8% of advanced Heart Failure clinic visits
 - · Questions to consider:
 - Is there equity in access to cardiology and advanced heart failure care?
 - · Are Black patients represented in clinical trials at MHIF?
 - Is there clinical bias in consideration for Heart Transplants and VAD?



Alina Health & ABBOTT NORTHWESTERN





What Are The Factors That Drive Disparities in Heart Failure?

- · Social determinants of health: insurance, education, nutrition, housing, income transportation
- Healthcare provider discrimination and bias: reduced delivery of evidence based heart failure treatments to racial/ethnic minorities and women
- Disparities in participation in clinical trials: Racial minorities are underrepresented in heart failure research studies
- Lack of preventive care: **modifiable diseases** that increase the risk of developing heart failure, such as hypertension, diabetes, obesity, and atherosclerosis





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A Multi-Faceted Approach to Achieve Health Equity

- · Increase awareness. Recognize that disparities and bias exist
- **Promote favorable lifestyle changes** that are associated with reduced risk of developing heart failure, particularly among African-Americans
- · Address structural inequalities in education, income, and health insurance coverage
- Change policy in order to expand access to care and the distribution of health services
- Educate and train in bias reduction, anti-racism, and cultural competency
- Encourage informed involvement in clinical research





Dr. Mario Goessl, MD, PhD





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