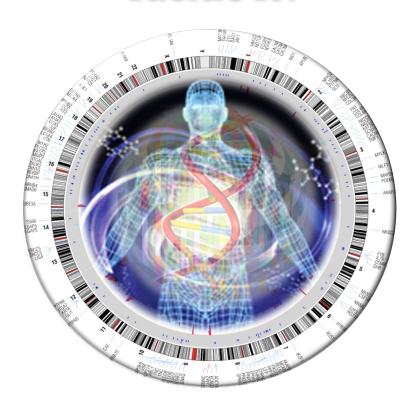
# Cancer Disparity from Different Angles: How to Tackle It?



John D. Carpten, Ph.D.

Director, City of Hope Comprehensive Cancer Center

Director, Beckman Research Institute

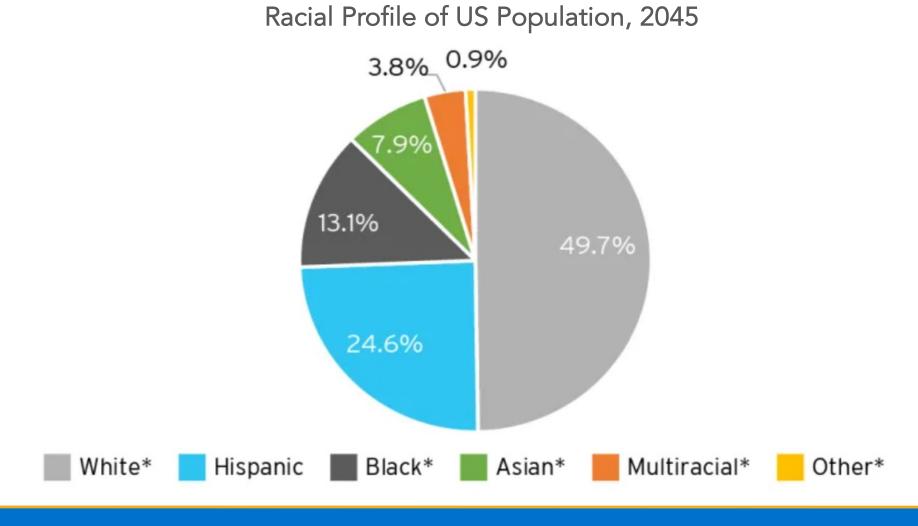


# Cancer Disparities and Cancer Health Equity

This presentation will focus mostly on cancer disparities among underrepresented racial/ethnic minorities and their related global diaspora, but we do not want to ignore the fact that there are other important diseases and also other medically underserved communities/populations.



# Cancer Disparities and Cancer Health Equity - Significance





# Cancer Statistics and Cancer Disparities

Table 5. Comparison of Cancer Incidence Rates between Non-Hispanic (NH) Blacks and Whites, US, 2008-2012

Female Male NH NH NH NH Black White Absolute Black White Absolute Rate Rate Rate\* Rate\* Difference<sup>†</sup> Ratio<sup>‡</sup> Rate\* Rate\* Difference<sup>†</sup> Ratio<sup>‡</sup> Cancer Cancer Kaposi sarcoma 3.96 Kaposi sarcoma 1.7 0.5 1.2 3.57 0.2 < 0.1 0.1 Myeloma 14.8 7.8 Myeloma 4.3 6.8 2.58 7.0 2.11 11.1 Stomach Stomach 15.1 7.8 7.3 1.93 8.0 3.5 4.5 2.30 Liver & intrahepatic bile duct Liver & intrahepatic bile duct 16.5 9.3 7.2 1.77 4.8 3.2 1.6 1.52 85.7 7.1 2.9 1.41 Prostate 208.7 123.0 1.70 Uterine cervix 10.0 9.3 6.3 3.0 1.48 Pancreas 14.4 10.6 3.8 1.36 Larynx Breast 20 1.4 0.6 1.45 Esophagus 2.5 1.8 0.7 1.34 Colon & rectum 60.3 47.4 12.9 1.27 Colon & rectum 44.1 36.2 7.9 1.22 1.7 17.2 3.2 1.23 Kidney & renal pelvis 13.0 11.3 1.15 Pancreas 14.0 Lung & bronchus 128.1 -3.80.97 93.4 79.3 14.1 1.18 Breast 124.3 Kidney & renal pelvis 24.2 2.4 1.11 Uterine corpus 23.0 25.5 -2.5 0.90 21.8 Hodgkin lymphoma -0.2Hodgkin lymphoma 2.7 0.88 3.2 3.4 0.95 2.4 -0.3Esophagus 8.0 8.8 -0.80.90 Lung & bronchus 51.4 58.7 -7.30.87 Leukemia 13.2 -4.5Leukemia 10.7 0.80 17.7 0.75 8.6 -2.1Oral cavity & pharynx Oral cavity & pharynx -1.5 0.78 15.3 18.1 -2.80.84 5.2 6.7 Non-Hodgkin lymphoma 0.77 -6.912.4 -2.8 17.2 24.1 0.71 Ovary 9.6 Brain & other nervous system Non-Hodgkin lymphoma 0.72 4.9 8.8 -3.90.56 12.0 16.6 -4.6Urinary bladder 19.8 -20.40.49 Urinary bladder 6.7 9.9 -3.20.68 40.2 21.9 0.59 Thyroid 3.7 7.7 -4.00.48 Thyroid 12.9 -9.0 Brain & other nervous system Testis 1.4 6.8 -5.43.6 6.3 -2.70.58 0.21 Melanoma of the skin 0.05 31.3 -30.20.04 Melanoma of the skin 1.0 20.6 -19.61.1 592.3 528.9 All sites 63.4 1.12 All sites 408.1 436.2 -28.1 0.94

www.cancer.org

City of Hope Compre

Note: Sites listed in descending order by rate ratio. \*Rates are per 100,000 and age adjusted to the 2000 US standard population. †Absolute difference is the rate in blacks minus the rate in whites. ‡Rate ratio is the unrounded rate in blacks divided by the unrounded rate in whites.

Source: North American Association of Central Cancer Registries. 183



# Cancer Statistics and Cancer Disparities

Table 6. Comparison of Cancer Death Rates between Non-Hispanic (NH) Blacks and Whites, US, 2008-2012

Male Female NH NH NH NH Black White Absolute Black White Absolute Rate Rate Ratio<sup>‡</sup> Difference† Rate\* Difference† Ratio<sup>‡</sup> Cancer Rate\* Rate\* Cancer Rate\* 5.8 2.58 2.7 Stomach 9.4 3.6 Stomach 4.5 1.8 2.48 47.2 2.38 Myeloma 3.0 2.22 Prostate 19.9 27.3 5.4 2.4 Uterine cervix Larynx 3.7 1.8 1.9 2.02 4.1 2.0 2.1 2.00 Myeloma 7.8 Uterine corpus 4.0 3.8 1.95 4.1 3.7 1.92 Liver & intrahepatic bile duct Liver & intrahepatic bile duct 12.8 5.2 4.4 3.1 1.3 1.43 7.6 1.69 Colon & rectum 27.6 1.42 18.2 9.4 1.52 Breast 31.0 21.9 9.1 Oral cavity & pharynx 5.2 12.9 3.8 1.4 1.36 Colon & rectum 18.2 5.3 1.41 Lung & bronchus 74.9 12.7 9.5 62.2 1.20 12.6 3.1 1.32 Pancreas 15.4 12.7 2.7 1.21 Esophagus 2.0 1.6 0.4 1.28 Pancreas Kidney & renal pelvis 5.7 5.9 -0.2Urinary bladder 2.3 1.12 0.97 0.3 Hodgkin lymphoma 0.94 Kidney & renal pelvis 0.4 0.5 -0.12.6 0.0 1.02 Lung & bronchus Esophagus 0.89 7.1 8.0 -0.936.7 41.1 -4.4 0.89 Leukemia Leukemia 0.89 8.1 -1.8 0.82 4.8 5.4 -0.6Non-Hodgkin lymphoma 5.9 -2.4 Hodgkin lymphoma 0.0 0.89 0.71 0.3 0.3 Urinary bladder 0.83 5.4 -3.00.65 Ovary 6.8 -1.48.4 Brain & other nervous system Non-Hodgkin lymphoma 3.2 -2.80.53 3.6 -1.40.71 6.0 Melanoma of the skin Brain & other nervous system 0.5 5.0 -4.5 0.09 2.2 3.9 -1.7 0.55 Melanoma of the skin 0.4 2.1 -1.70.18 All sites 267.7 210.6 57.1 1.27 All sites 170.4 149.2 21.2 1.14

www.cancer.org

Source: National Center for Health Statistics, Centers for Disease Control and Prevention as provided by the SEER program in the SEER\*Stat database.186

blacks minus the rate in whites. ‡Rate ratio is the unrounded rate in blacks divided by the unrounded rate in whites.

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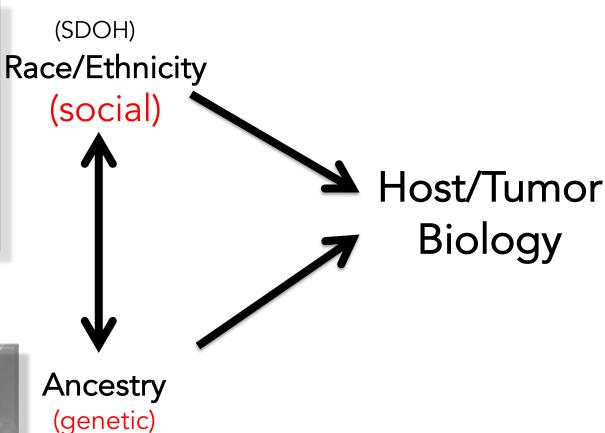
# Race and Ancestry Represent Different but Related Factors

Social/Societal

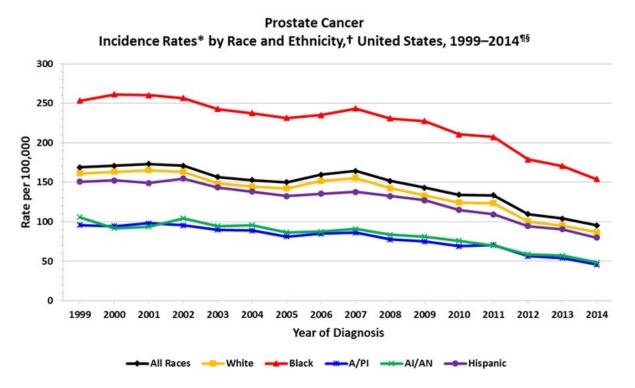
**VS** 

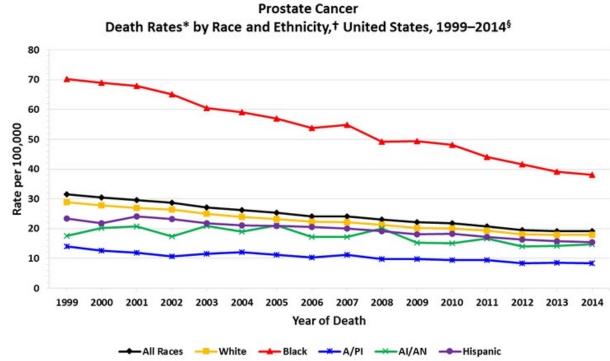
Biological/Genetic





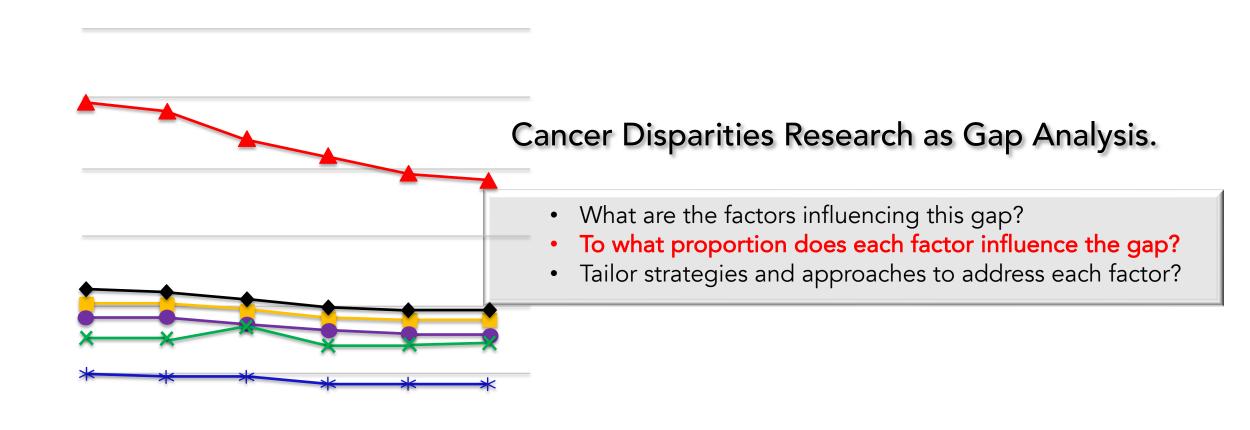
# Disparities as Gaps



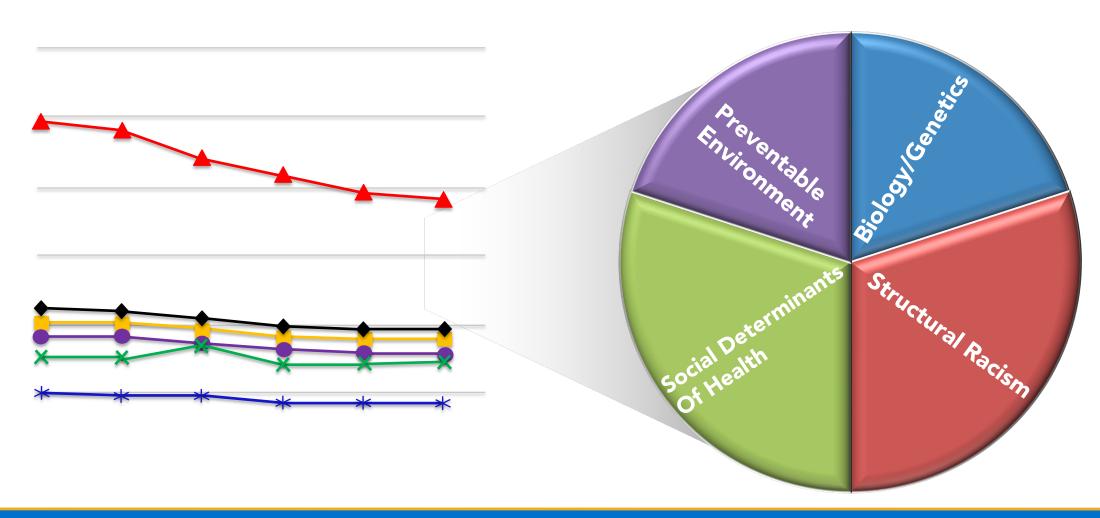


https://www.cdc.gov/

# Disparities as Gaps – Reducing the Gaps

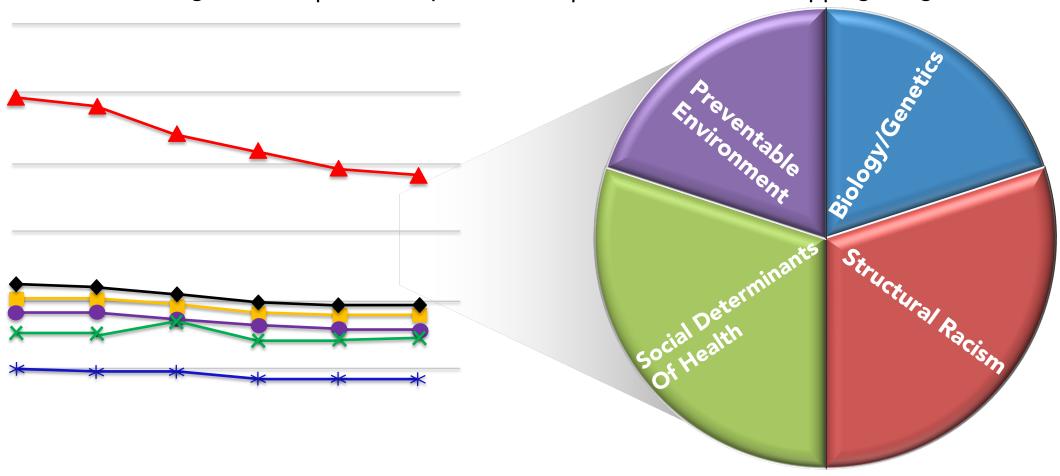


# Disparities as Gaps – Reducing the Gaps



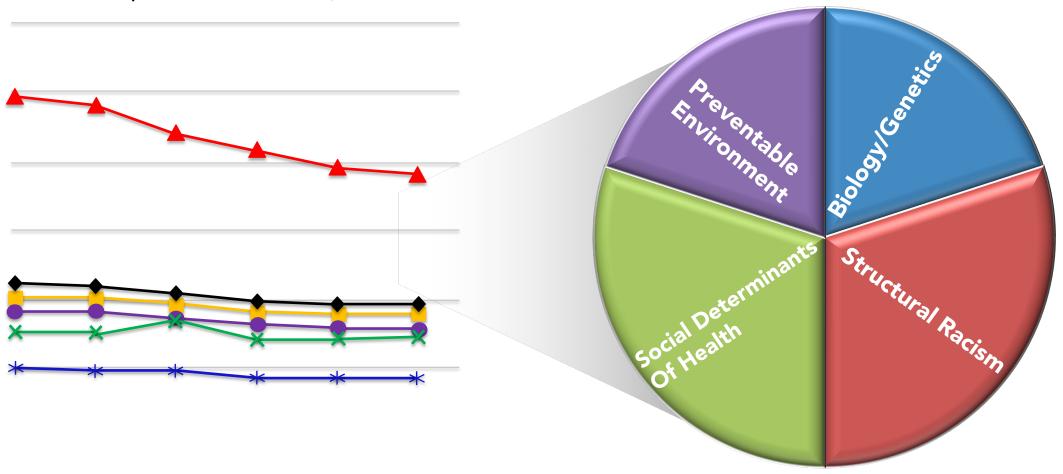
# Disparities as Gaps – Reducing the Gaps

Addressing each component requires some specific but also overlapping mitigators



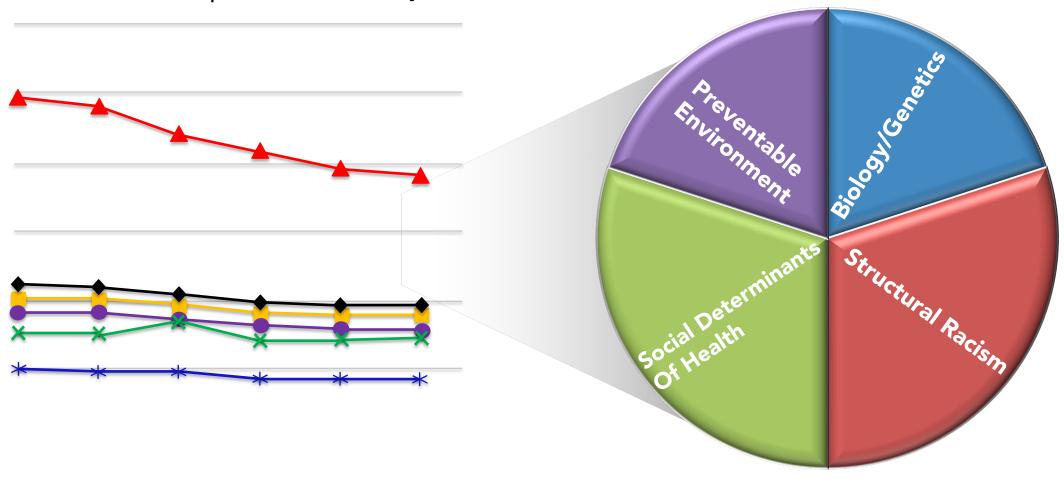
# Disparities as Gaps – Reducing the Gaps – Context Dependent

Context Dependent – Differ by Indication (PrCa vs Cervical; or BrCa vs CRC; LuCa vs Endometrial)



# Disparities as Gaps – Reducing the Gaps – Context Dependent

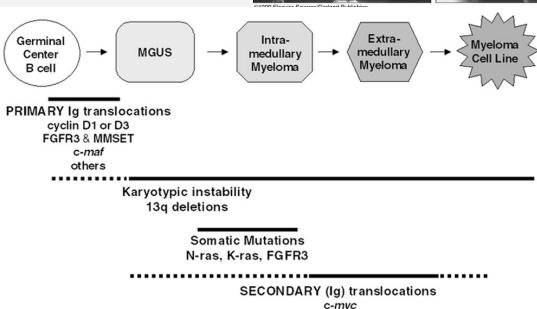
Context Dependent – Differ by Clinical Issue (Incidence versus Outcomes)



# Multiple Myeloma Pathogenesis

Cancer of the plasma cells in the bone marrow leading to bone destruction and bone marrow failure.





- ~31,000 new Myeloma cases and an estimated
   ~13,000 deaths attributed.
- Historically treated with Dexamethasone and bone marrow transplant.
- More recent development of myeloma "block buster "drugs.
  - iMids Lenalidomide
  - Proteosome inhibitors Bortezomib,
     Carfilzomab
- 5 year survival rate has increased from ~30% to >50% in last 10-15yrs!!

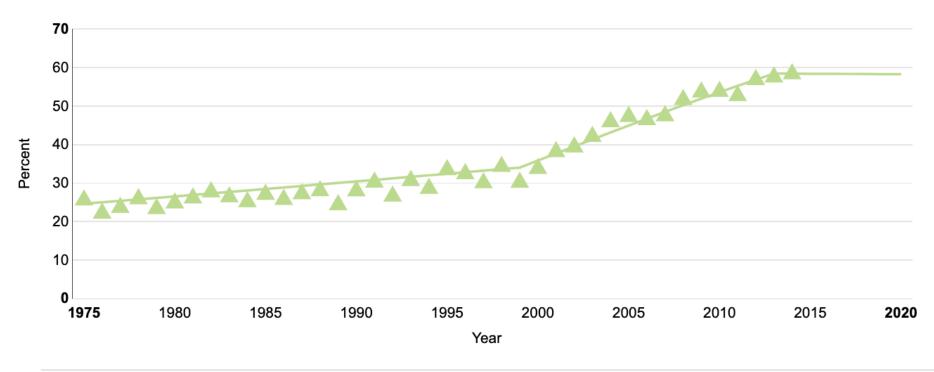


Dalton W et al, Amer Soc Hematology 2001

# Multiple Myeloma Outcomes



## 5-Year Relative Survival



SEER 8 5-Year Relative Survival Percent from 1975–2014, All Races, Both Sexes.

Modeled trend lines were calculated from the underlying rates using the <u>Joinpoint Survival Model</u> Software.

www.cancer.org

Table 5. Comparison of Cancer Incidence Rates between Non-Hispanic (NH) Blacks and Whites, US, 2008-2012

Male					Female					
Cancer	NH Black Rate*	NH White Rate*	Absolute Difference <sup>†</sup>	Rate Ratio‡	Cancer	NH Black Rate*	NH White Rate*	Absolute Difference <sup>†</sup>	Rate Ratio‡	
Kaposi sarcoma	1.7	0.5	1.2	3.57	Kaposi sarcoma	0.2	<0.1	0.1	3.96	
Myeloma ⊁	14.8	7.0	7.8	2.11	Myeloma ⊁	11.1	4.3	6.8	2.58	
Stomach	15.1	7.8	7.3	1.93	Stomach	8.0	3.5	4.5	2.30	
Liver & intrahepatic bile duct	16.5	9.3	7.2	1.77	Liver & intrahepatic bile duct	4.8	3.2	1.6	1.52	
Prostate	208.7	123.0	85.7	1.70	Uterine cervix	10.0	7.1	2.9	1.41	
Larynx	9.3	6.3	3.0	1.48	Pancreas	14.4	10.6	3.8	1.36	
Breast	2.0	1.4	0.6	1.45	Esophagus	2.5	1.8	0.7	1.34	
Colon & rectum	60.3	47.4	12.9	1.27	Colon & rectum	44.1	36.2	7.9	1.22	
Pancreas	17.2	14.0	3.2	1.23	Kidney & renal pelvis	13.0	11.3	1.7	1.15	
Lung & bronchus	93.4	79.3	14.1	1.18	Breast	124.3	128.1	-3.8	0.97	
Kidney & renal pelvis	24.2	21.8	2.4	1.11	Uterine corpus	23.0	25.5	-2.5	0.90	
Hodgkin lymphoma	3.2	3.4	-0.2	0.95	Hodgkin lymphoma	2.4	2.7	-0.3	0.88	
Esophagus	8.0	8.8	-0.8	0.90	Lung & bronchus	51.4	58.7	-7.3	0.87	
Leukemia	13.2	17.7	-4.5	0.75	Leukemia	8.6	10.7	-2.1	0.80	
Oral cavity & pharynx	15.3	18.1	-2.8	0.84	Oral cavity & pharynx	5.2	6.7	-1.5	0.78	
Non-Hodgkin lymphoma	17.2	24.1	-6.9	0.71	Ovary	9.6	12.4	-2.8	0.77	
Brain & other nervous system	4.9	8.8	-3.9	0.56	Non-Hodgkin lymphoma	12.0	16.6	-4.6	0.72	
Urinary bladder	19.8	40.2	-20.4	0.49	Urinary bladder	6.7	9.9	-3.2	0.68	
Thyroid	3.7	7.7	-4.0	0.48	Thyroid	12.9	21.9	-9.0	0.59	
Testis	1.4	6.8	-5.4	0.21	Brain & other nervous system	3.6	6.3	-2.7	0.58	
Melanoma of the skin	1.1	31.3	-30.2	0.04	Melanoma of the skin	1.0	20.6	-19.6	0.05	
All sites	592.3	528.9	63.4	1.12	All sites	408.1	436.2	-28.1	0.94	

Note: Sites listed in descending order by rate ratio. \*Rates are per 100,000 and age adjusted to the 2000 US standard population. †Absolute difference is the rate in blacks minus the rate in whites. ‡Rate ratio is the unrounded rate in blacks divided by the unrounded rate in whites.

Source: North American Association of Central Cancer Registries. 183

@2016, American Cancer Society, Inc., Surveillance Research

Table 6. Comparison of Cancer Death Rates between Non-Hispanic (NH) Blacks and Whites, US, 2008-2012

Male

Female

Cancer	NH Black Rate*	NH White Rate*	Absolute Difference†	Rate Ratio‡	Cancer	NH Black Rate*	NH White Rate*	Absolute Difference†	Rate Ratio‡
Stomach	9.4	3.6	5.8	2.58	Stomach	4.5	1.8	2.7	2.48
Prostate	47.2	19.9	27.3	2.38	Myeloma ★	5.4	2.4	3.0	2.22
Larynx	3.7	1.8	1.9	2.02	Uterine cervix	4.1	2.0	2.1	2.00
Myeloma ⊁	7.8	4.0	3.8	1.95	Uterine corpus	7.8	4.1	3.7	1.92
Liver & intrahepatic bile duct	12.8	7.6	5.2	1.69	Liver & intrahepatic bile duct	4.4	3.1	1.3	1.43
Colon & rectum	27.6	18.2	9.4	1.52	Breast	31.0	21.9	9.1	1.42
Oral cavity & pharynx	5.2	3.8	1.4	1.36	Colon & rectum	18.2	12.9	5.3	1.41
Lung & bronchus	74.9	62.2	12.7	1.20	Pancreas	12.6	9.5	3.1	1.32
Pancreas	15.4	12.7	2.7	1.21	Esophagus	2.0	1.6	0.4	1.28
Kidney & renal pelvis	5.7	5.9	-0.2	0.97	Urinary bladder	2.6	2.3	0.3	1.12
Hodgkin lymphoma	0.4	0.5	-0.1	0.94	Kidney & renal pelvis	2.6	2.6	0.0	1.02
Esophagus	7.1	8.0	-0.9	0.89	Lung & bronchus	36.7	41.1	-4.4	0.89
Leukemia	8.1	9.9	-1.8	0.82	Leukemia	4.8	5.4	-0.6	0.89
Non-Hodgkin lymphoma	5.9	8.3	-2.4	0.71	Hodgkin lymphoma	0.3	0.3	0.0	0.89
Urinary bladder	5.4	8.4	-3.0	0.65	Ovary	6.8	8.2	-1.4	0.83
Brain & other nervous system	3.2	6.0	-2.8	0.53	Non-Hodgkin lymphoma	3.6	5.0	-1.4	0.71
Melanoma of the skin	0.5	5.0	-4.5	0.09	Brain & other nervous system	2.2	3.9	-1.7	0.55
					Melanoma of the skin	0.4	2.1	-1.7	0.18
All sites	267.7	210.6	57.1	1.27	All sites	170.4	149.2	21.2	1.14

Note: Sites listed in descending order by rate ratio. \*Rates are per 100,000 and age adjusted to the 2000 US standard population. †Absolute difference is the rate in blacks minus the rate in whites. ‡Rate ratio is the unrounded rate in blacks divided by the unrounded rate in whites.

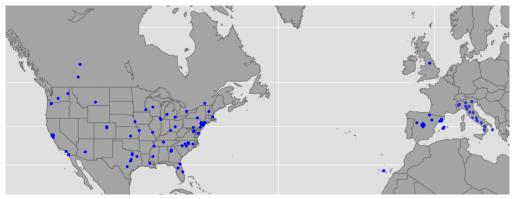
Source: National Center for Health Statistics, Centers for Disease Control and Prevention as provided by the SEER program in the SEER\*Stat database.186

@2016, American Cancer Society, Inc., Surveillance Research

www.cancer.org

#### **CoMMpass Enrollment**

1,000+ patients enrolled from >90 sites worldwide, with over 850 samples molecularly profiled at baseline and >100 sequentially.











Angela Baker

Zarko Manojlovic Jonathan Keats

RESEARCH ARTICLE

PLOS GENETICS

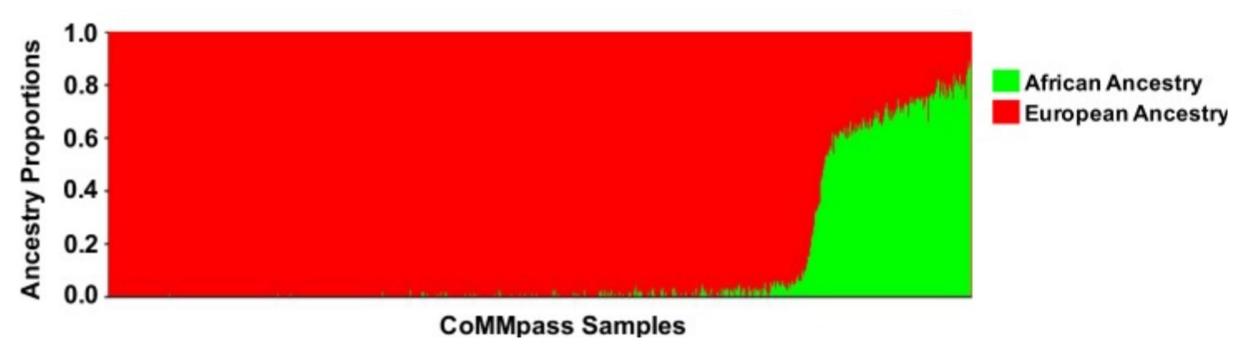
Comprehensive molecular profiling of 718 Multiple Myelomas reveals significant differences in mutation frequencies between African and European descent cases

Zarko Manojlovic<sup>1,2</sup>\*, Austin Christofferson<sup>2</sup>, Winnie S. Liang<sup>2</sup>, Jessica Aldrich<sup>2</sup>, Megan Washington<sup>2</sup>, Shukmei Wong<sup>2</sup>, Daniel Rohrer<sup>3</sup>, Scott Jeweli<sup>3</sup>, Rick A. Kittles<sup>4</sup>, Mary Derome<sup>5</sup>, Daniel Auclair<sup>5</sup>, David Wesley Craig<sup>1</sup>, Jonathan Keats<sup>2</sup>, John D. Carpten<sup>1,2</sup>

1 Department of Translational Genomics, Keck School of Medicine, University of Southern California, Los Angeles, CA, United States of America, 2 Translational Genomics Research Institute, Phoenix, AZ, United States of America, 3 Van Andel Research Institute, Grand Rapids, MI, United States of America, 4 Department of Surgery, Division of Population Genetics, University of Arizona, Tuscon, AZ, United States of America, 5 Multiple Myeloma Research Foundation, Norwalk, CT, United States of America

# Genomic Studies of Myeloma Cancer Disparities

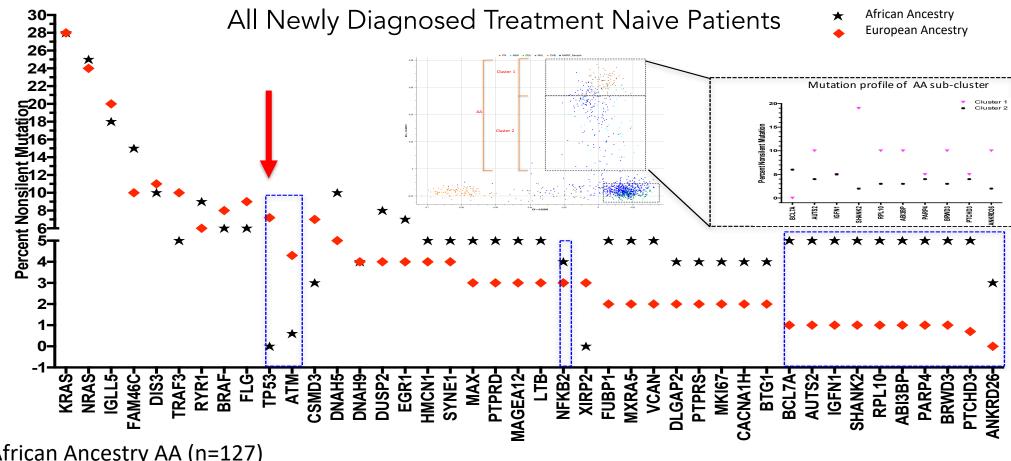
## All Newly Diagnosed Treatment Naive Patients



African Ancestry AA (n=127) European Ancestry EA (n=591)

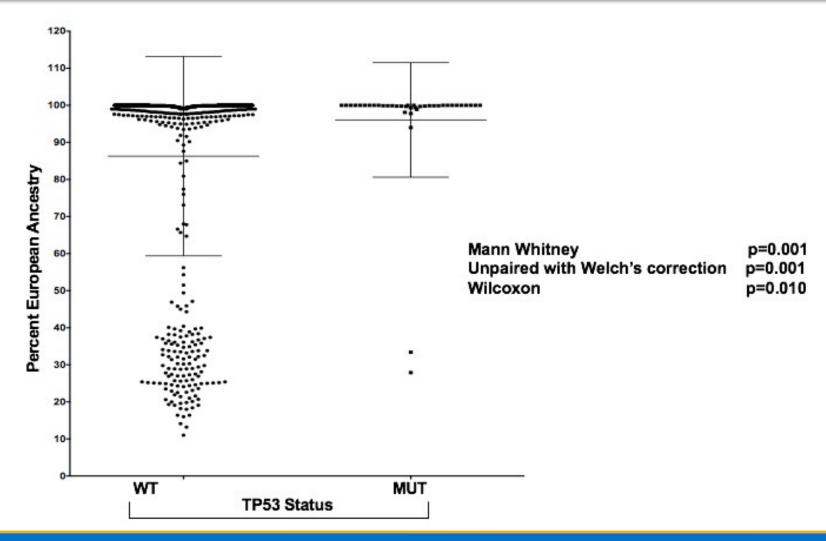
Manojlovic et al., PLoS Genet. 2017; 13(11):e1007087.

# Genomic Studies of Myeloma Cancer Disparities



African Ancestry AA (n=127) European Ancestry EA (n=591)

Manojlovic et al., PLoS Genet. 2017; 13(11):e1007087.



- These data provide clear evidence of differences in tumor profiles among AA and EA MM patients in a sufficiently powered tumor cohort.
- TP53 loss is associated with poor outcome, yet is enriched in tumors from EA cases.
- These data suggest that AA MM patients may have tumors with molecular features associated with more favorable outcomes.
- Perhaps in MM, equal treatment could lead to similar or better outcomes in AA patients.

#### STIMULUS REPORT

blood advances

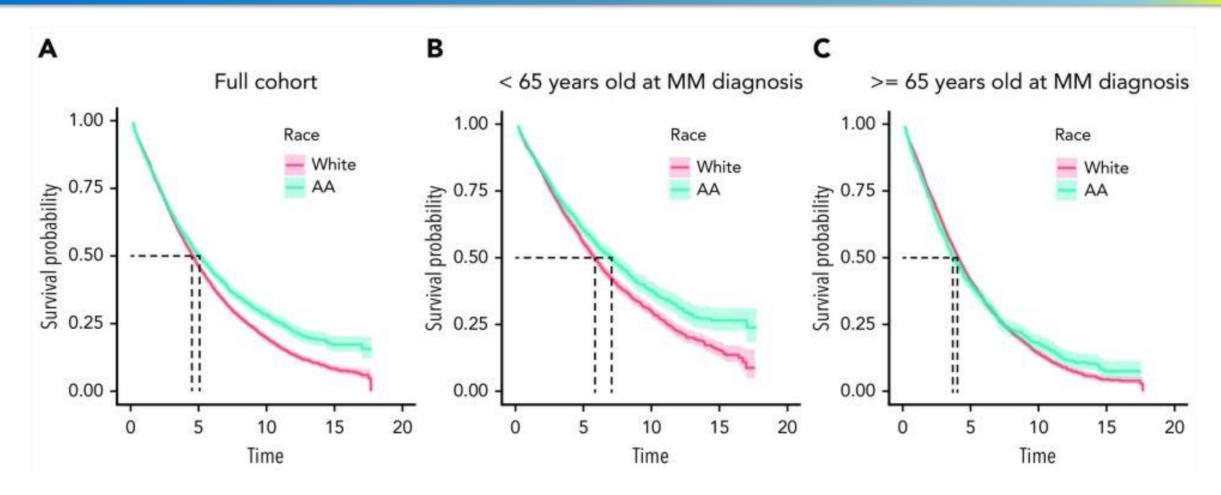
Survival of ethnic and racial minority patients with multiple myeloma treated with newer medications

E. Dianne Pulte, Lei Nie, Nicole Gormley, Kirsten B. Goldberg, Amy McKee, Ann Farrell, and Richard Pazdur

Center for Drug Evaluation and Research, US Food and Drug Administration, Silver Spring, MD

Pulte et al., Blood Advances. 2018 Jan 23;2(2):116-119.

- Results show a low participation of minorities in MM clinical trials of newer agents.
- Despite this limitation, mortality was slightly lower in participants of other racial and ethnic groups.
- Observed that Imid- and PI-class drugs may be working better in minorities.
- Examination of survival in the current SEER data shows that overall 5-year survival from 2007 to 2013 increased to 52.3% for African Americans and 50.6% for whites, suggesting that the earlier observed disparities were likely related to a temporary phenomenon (ie, differences in treatment utilization, access).



Fillmore et al., *Blood*. 2019 Jun 13;133(24):2615-2618.

Disparities in MM Outcomes Likely Due to Access to Quality Care

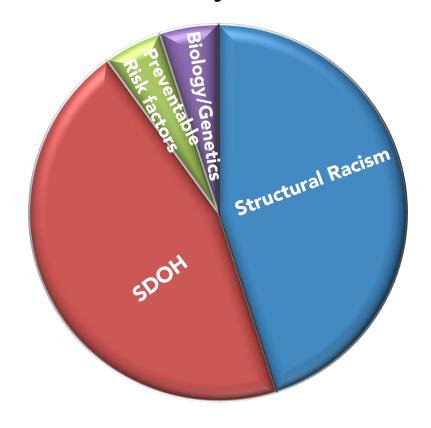


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Note: Sites listed in descending order by rate ratio. \*Rates are per 100,000 and age adjusted to the 2000 US standard population. †Absolute difference is the rate in blacks minus the rate in whites. ‡Rate ratio is the unrounded rate in blacks divided by the unrounded rate in whites.

Source: North American Association of Central Cancer Registries. \*\*ISIS\*\*

Melanoma of the skin

All sites

0.04

1.12

@2016, American Cancer Society, Inc., Surveillance Research

20.6

436.2

-19.6

-28.1

0.05

0.94

1.0

408.1

#### Disparities in MM Incidence Rates Remain

- Rate of MGUS and conversion rate to MM??
- Immunological Factors??
- Germline Genetic Risk??
- Environmental Exposure??
- Some combination of the above??

-30.2

63.4

Melanoma of the skin

All sites

1.1

592.3

31.3

528.9

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Larynx	9.3	6.3	3.0	1.48	Pancreas	14.4	10.6	3.8	1.36
Breast	2.0	1.4	0.6	1.45	Esophagus	2.5	1.8	0.7	1.34
Colon & rectum	60.3	47.4	12.9	1.27	Colon & rectum	44.1	36.2	7.9	1.22
Pancreas	17.2	14.0	3.2	1.23	Kidney & renal pelvis	13.0	11.3	1.7	1.15
Lung & bronchus	93.4	79.3	14.1	1.18	Breast	124.3	128.1	-3.8	0.97
Kidney & renal pelvis	24.2	21.8	2.4	1.11	Uterine corpus	23.0	25.5	-2.5	0.90
Hodgkin lymphoma	3.2	3.4	-0.2	0.95	Hodgkin lymphoma	2.4	2.7	-0.3	0.88
Esophagus	8.0	8.8	-0.8	0.90	Lung & bronchus	51.4	58.7	-7.3	0.87
Leukemia	13.2	17.7	-4.5	0.75	Leukemia	8.6	10.7	-2.1	0.80
Oral cavity & pharynx	15.3	18.1	-2.8	0.84	Oral cavity & pharynx	5.2	6.7	-1.5	0.78
Non-Hodgkin lymphoma	17.2	24.1	-6.9	0.71	Ovary	9.6	12.4	-2.8	0.77
Brain & other nervous system	4.9	8.8	-3.9	0.56	Non-Hodgkin lymphoma	12.0	16.6	-4.6	0.72
Urinary bladder	19.8	40.2	-20.4	0.49	Urinary bladder	6.7	9.9	-3.2	0.68
Thyroid	3.7	7.7	-4.0	0.48	Thyroid	12.9	21.9	-9.0	0.59
Testis	1.4	6.8	-5.4	0.21	Brain & other nervous system	3.6	6.3	-2.7	0.58
Melanoma of the skin	1.1	31.3	-30.2	0.04	Melanoma of the skin	1.0	20.6	-19.6	0.05
All sites	592.3	528.9	63.4	1.12	All sites	408.1	436.2	-28.1	0.94

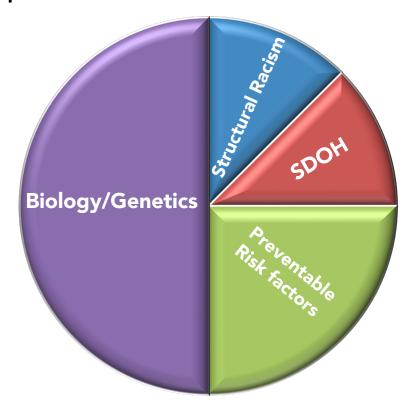
Note: Sites listed in descending order by rate ratio. \*Rates are per 100,000 and age adjusted to the 2000 US standard population. †Absolute difference is the rate in blacks minus the rate in whites. ‡Rate ratio is the unrounded rate in blacks divided by the unrounded rate in whites.

Source: North American Association of Central Cancer Registries.

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@2016, American Cancer Society, Inc., Surveillance Research

## Disparities in MM Incidence Rates

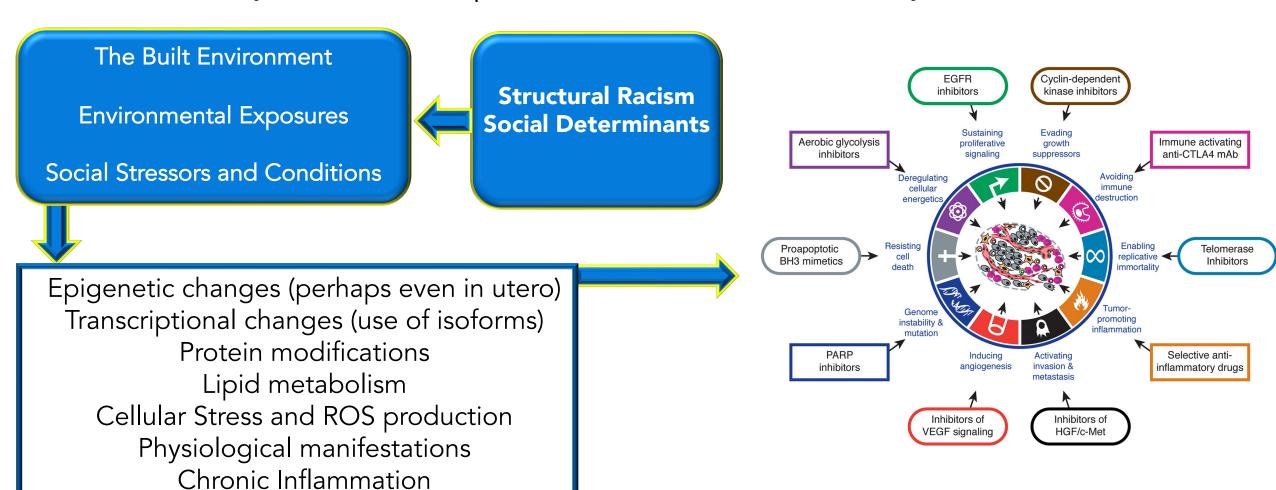


Perhaps a different proportion of factors influencing risk?



# Integration of Environment/Social/Biological Sciences

Cancer exacts a heavy burden on underrepresented communities and the medically underserved.

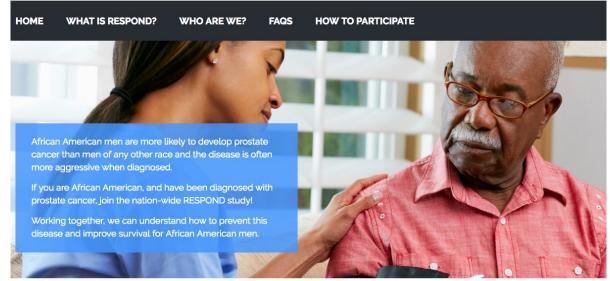


Hallmarks of Cancer: The Next Generation, Hanahan and Weinberg (2011), Cell, 144:5, 646-674

# Integration of Environment/Social/Biological Sciences



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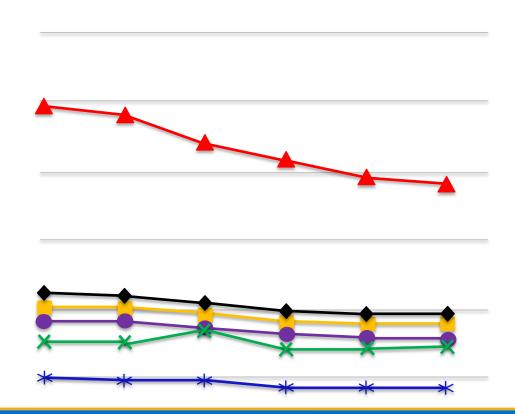


A large, national, population-based cohort study, RESPOND, (Research on Prostate Cancer in Men of African Ancestry: Defining the Roles of Genetics, Immunity and Access to Care) of 10,000 AA men with incident PCa identified through nine SEER and NPCR U.S. cancer registries from states that include 38% of all AA PCa cases in the U.S.





# Must Continue Analyzing the Gaps





# To Close the Gaps to Achieve Health Equity

