Assessment of Prevention Research Measuring Leading Risk Factors and Causes of Mortality and Disability Supported by the US National Institutes of Health

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Council of Councils January 24, 2020



National Institutes of Health Office of Disease Prevention

### Strategic Priority I

- Systematically monitor NIH investments in prevention research and assess the progress and results of that research.
  - ODP defines prevention research to include primary and secondary prevention in humans, together with relevant methods development.
  - ODP's definition includes research designed to promote health; to prevent onset of disease, disorders, conditions, or injuries; and to detect, and prevent the progression of, asymptomatic disease.
  - Prevention research includes studies for:
    - Identification and assessment of risk and protective factors,
    - Screening and identification of individuals and groups at risk,
    - Development and evaluation of interventions to reduce risk,
    - Translation, implementation, and dissemination of effective, preventive interventions into practice, and
    - Development of methods to support prevention research.



### Selection of Activity Codes

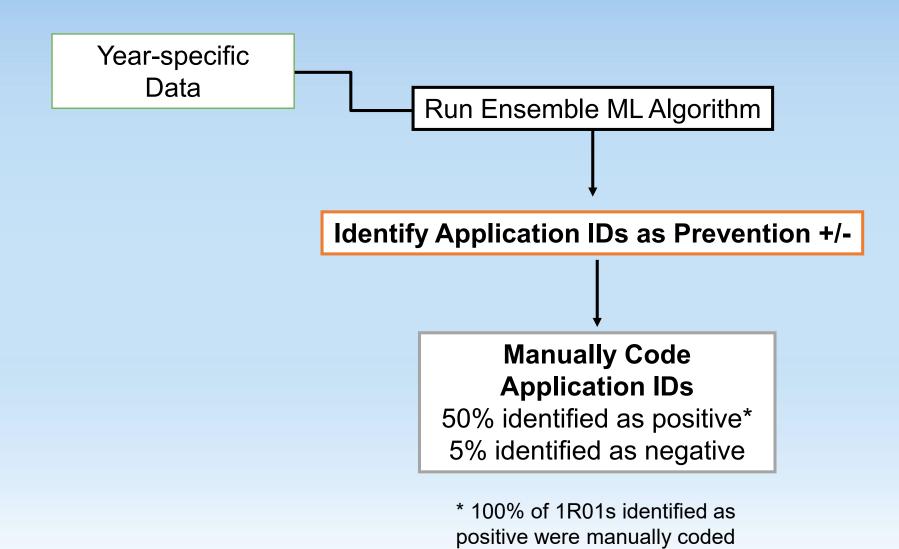
- ODP worked with staff from many ICs to identify activity codes likely to support NIH prevention research that met ODP's definition.
  - Basic and preclinical research were excluded.
  - Awards for community services, facilities, infrastructure, loan repayment, meetings, planning, and training were excluded.
  - Intramural research was excluded to focus on extramural research.
  - Contracts proved too difficult to code using our methods.
  - Methodological research was included only if it yielded products that were applicable to prevention research without additional development.
  - We included all remaining R, P, and U activity codes with at least 500 awards across FY12-17 or at least \$500M awarded across FY12-17.
  - Several of these activity codes involved awards with multiple subprojects; as a result, we sampled Application IDs (Appl IDs) instead of awards.

### Portfolio Coverage by These 12 Activity Codes

			Research	ODP's	% Research
		R, P, U	R, P, U	Selected	R, P, U
	All Activity	Activity	Activity	Activity	Activity
	Codes	Codes	Codes	Codes	Codes
Total Awards	111,626	68,757	63,381	58,104	91.7%
Total Costs	\$57.5 B	\$32.6 B	\$30.6 B	\$25.7 B	84.1%

All figures based on Type I, 2, and 9 awards from FY12-17, excluding parent awards for projects with sub-awards to avoid double counting.

### Sampling of Application IDs



### Coding Based on a Prevention Research Taxonomy

- A classification system to characterize projects or subprojects on:
  - Study Focus
    - Rationale
    - Exposures
    - Outcomes
  - Population focus
  - Study design/purpose
  - Prevention research category
- 128 topics, 29-page protocol
  - Applied to title, abstract, public health significance
- Input from the PRCC

				Rater	
				Date:	
Prevention Taxonomy Form CHECK ALL THAT APPLY IN EACH COLUMN (TOPICS ARE NOT MUTUALLY EXCLUSIVE) See accompanying protocol for definitions and examples					
ApplIDPILast N			Project I	ile:	
Study focus	Rationale	Exposure	Outcome	Population focus	
1. Alcohol				1. Incarcerated/institutionalized	
2. Alzheimer's disease				2. LGBTI	
3.		1		3. Low income	
4. Blood disorder				4. Military/veterans	
5. Blood pressure				5. Older adults/elderly	
6. Cancer				6. People with disabilities	
7. Chemical/toxin				7. Pregnant and/or post-partum women 8 Rural	
8. Cholesterol	_			d. Third	
9. Diabetes				9. Urban 10. Youth (infants, children, adolescents)	
10. Diet/nutrition				11. Other or unclear	
11. Education/counseling	_			TT. Other of unclear	
12. Firearms 13. Gastrointestinal disease					
13. Gastrointestinar disease				Study design/purpose	
15. Healthcare delivery				1. Analysis of existing data	
16. Heart disease				2. Methods research	
17. HR quality of life	_			3. Non-randomized intervention study	
18. Infectious disease				4. Observational study	
19. Kidney disease				5. Pilot/feasibility/proof-of-concept/safety	
20. Lung disease				6. Randomized intervention study	
21. Maternal/paternal/child				7. Other or unclear	
health				Prevention research category	
22. Medication/device				1. Preventing new health condition, promoting	
23. Mental health 24. Microbiorne				health in the general population, or	
25. Mortality				identifying risk factors for a new health	
25. Mortality 26. Motor vehicle crash				condition	
27. Musculoskeletal disease				2. Screening for risk factor	
28. Neurological disease				3. Screening for early disease	
(not Alzheimer's)				4. Preventing progression of disease	
29. Obesity				preventing recurrence in those with a	
30. Physical activity				known health condition, identifying risk	
31. Policy/built environment				factors for progression or recurrence	
32 Pneumonia/influenza				5. Methods research	
33. Sexual behavior				6. Other or unclear	
34. Stress	_				
35. Stroke					
36. Substance abuse					
37. Suicide					
38. Surgery					
39. Tobacco					
40. Unintentional injuries					
41. Vaccine					
42. Violence					
43. Other or unclear					

### **Team-Based Coding**



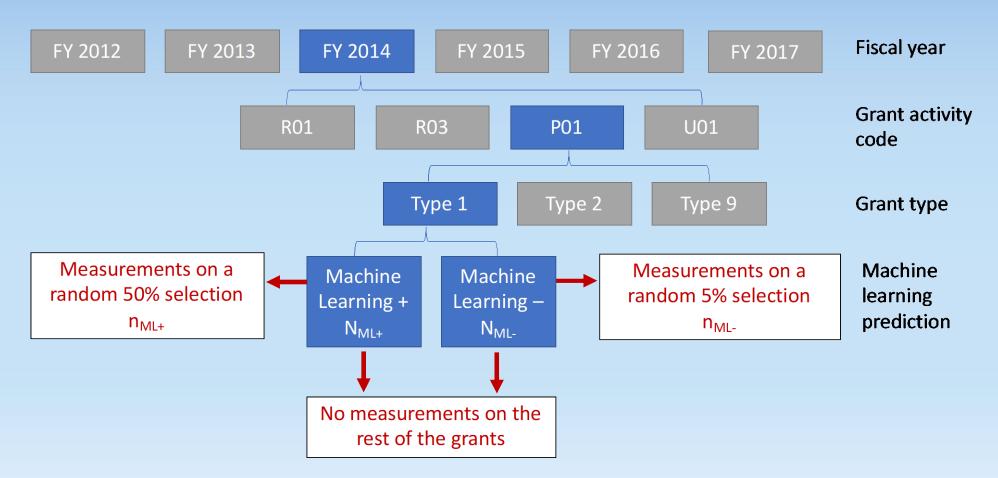
#### Coders

- Contract staff: MPH grads led by a PhD epidemiologist
- 2 months training in groups of 3-4
- Overseen by ODP staff

#### 3-person teams coded abstracts using iPads

- Each person coded independently, then the team resolved disagreements to generate a set of consensus codes for each Application ID
- ODP coded 10-20% of the abstracts weekly for QC using the same methods
- ODP reconciled discrepancies with the contract coding teams
- Average interrater agreement was 0.86.

### Weighting



Weights for FY14, P01, type 1, Machine Learning + =  $N_{ML+}$  /  $n_{ML+}$ 

Weights for FY14, P01, type 1, Machine Learning - =  $N_{ML-} / n_{ML-}$ 

### Primary and Secondary Prevention Research in Humans: FY12-17



### Primary and Secondary Prevention Research in Humans: FY12-17



### American Journal of Preventive Medicine

**RESEARCH ARTICLE** 

### NIH Primary and Secondary Prevention Research in Humans During 2012–2017

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### Follow-Up

- 74% of deaths in the U.S. are attributable to 10 well-known causes.<sup>a</sup>
- 57.3% of deaths and 42.1% of Disability-Adjusted Life Years lost in the U.S. are attributable to 10 well-known risk factors.<sup>b</sup>
- How does NIH prevention research address these causes and risk factors?
  - ODP staff worked through the database of 11,082 coded awards to revise coding for the leading risk factors and causes of death to be consistent with the definitions used by CDC and GBD.
  - ODP then repeated the portfolio analysis, focused on projects that included an exposure or outcome that was a leading risk factor or cause of death.
- <sup>a</sup> Centers for Disease Control (CDC) report Mortality in the United States, 2017. Accessed April 4th, 2019. https://www.cdc.gov/nchs/data/databriefs/db328\_tables-508.pdf#4.
- <sup>b</sup> U. S. Burden of Disease Collaborators, Mokdad AH, Ballestros K, et al. The State of US Health, 1990-2016: Burden of Diseases, Injuries, and Risk Factors Among US States. *JAMA*. 2018;319(14):1444-1472.



**Original Investigation** | Public Health

### Assessment of Prevention Research Measuring Leading Risk Factors and Causes of Mortality and Disability Supported by the US National Institutes of Health

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JAMA Network Open. 2019;2(11):e1914718. doi:10.1001/jamanetworkopen.2019.14718

# NIH Primary and Secondary Prevention Research in Humans vs. Leading Risk Factors for DALYs Lost in the U.S.

Leading Risk Factors - DALYs Lost	NIH prevention research portfolio, % projects (95% CI)	NIH prevention research portfolio, % dollars (95% Cl)	GBD, <sup>a</sup> % attributable DALYs lost
Any Top 10 Risk Factor - DALYs	31.4 (29.6-33.3)	30.3 (26.6-33.9)	42.1
1) High body mass index	5.3 (4.7-6.0)	6.5 (3.7-9.4)	11.6
2) Tobacco	6.6 (5.8-7.6)	5.4 (4.5-6.3)	11.1
3) Dietary risk	7.8 (7.0-8.8)	6.7 (5.7-7.7)	10.4
4) High fasting plasma glucose	4.6 (3.9-5.4)	6.6 (3.7-9.6)	9.7
5) High systolic blood pressure	2.7 (2.2-3.3)	3.1 (2.3-3.9)	8.0
6) Drug use	7.3 (6.4-8.2)	7.6 (6.0-9.2)	6.5
7) Alcohol use	5.6 (4.9-6.4)	4.1 (3.6-4.7)	4.2
8) High LDL cholesterol	1.8 (1.4-2.3)	2.0 (1.3-2.6)	4.0
9) Impaired kidney function	1.6 (1.0-2.3)	1.6 (1.0-2.3)	3.1
10) Occupational risks	0.3 (0.1-0.4)	0.2 (0.1-0.3)	2.5

# NIH Primary and Secondary Prevention Research in Humans vs. Leading Risk Factors for Deaths in the U.S.

Leading Risk Factors for Death	NIH prevention research portfolio, % projects (95% CI)	NIH prevention research portfolio, % dollars (95% Cl)	GBD, <sup>a</sup> % attributable deaths
Any Top 10 Risk Factor for Death	34.0 (32.2-35.9)	32.5 (28.9-36.2)	57.3
1) Dietary risk	7.8 (7.0-8.8)	6.7 (5.7-7.7)	19.1
2) Tobacco	6.6 (5.8-7.6)	5.4 (4.5-6.3)	17.8
3) High systolic blood pressure	2.7 (2.2-3.3)	3.1 (2.3-3.9)	17.4
4) High body mass index	5.3 (4.7-6.0)	6.5 (3.7-9.4)	13.9
5) High fasting plasma glucose	4.6 (3.9-5.4)	6.6 (3.7-9.6)	13.6
6) High total cholesterol	1.8 (1.4-2.3)	2.0 (1.3-2.6)	8.4
7) Impaired kidney function	1.6 (1.0-2.3)	1.6 (1.0-2.3)	6.3
8) Alcohol/drug use	11.2 (10.2-12.4)	10.2 (8.5-11.8)	5.6
9) Air pollution	1.4 (1.1-1.6)	1.4 (0.9-1.8)	3.8
10) Low physical activity	5.0 (4.4-5.7)	4.3 (3.7-4.8)	3.3

# NIH Primary and Secondary Prevention Research in Humans vs. Leading Causes of Deaths in the U.S.

Leading Causes of Death	NIH prevention research portfolio, % projects (95% CI)	NIH prevention research portfolio, % dollars (95% Cl)	CDC, <sup>a</sup> % attributable deaths
Any Top 10 Leading Cause of Death	25.9 (24.0-27.8)	28.2 (24.8-31.5)	74.0
1) Heart disease	4.2 (3.3-5.2)	4.8 (3.2-6.4)	23.0
2) Cancer	11.9 (10.5-13.4)	11.3 (9.2-13.4)	21.3
3) Accidents	1.7 (1.2-2.4)	1.7 (1.1-2.4)	6.0
4) Chronic lower respiratory disease	1.8 (1.4-2.3)	2.0 (1.3-2.7)	5.7
5) Stroke	2.7 (2.2-3.4)	3.3 (2.3-4.3)	5.2
6) Alzheimer's disease	2.0 (1.4-2.6)	3.2 (1.9-4.6)	4.3
7) Diabetes	3.6 (3.0-4.2)	4.5 (3.4-5.6)	3.0
8) Influenza/Pneumonia	0.5 (0.2-1.0)	0.7 (0.1-1.3)	2.0
9) Kidney disease	1.4 (0.9-2.2)	1.4 (0.8-2.0)	1.8
10) Suicide	0.7 (0.5-0.9)	0.7 (0.5-0.9)	1.7

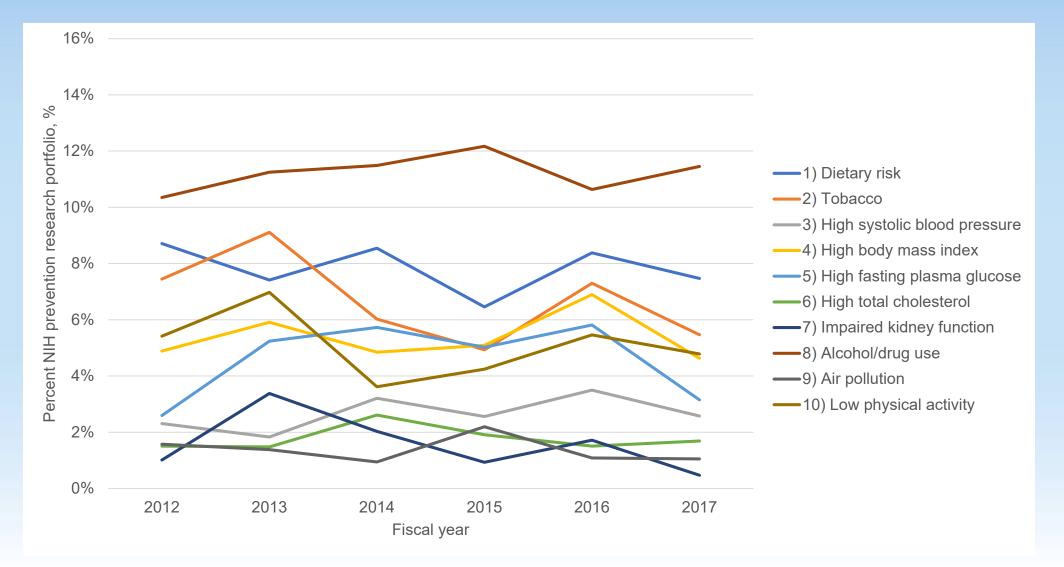
### NIH Primary and Secondary Prevention Research in Humans Involving Multiple Risk Factors or Causes of Death in the U.S.

- 3.3% (95% CI, 2.6-4.1) of prevention research projects measured more than one leading cause of death as an exposure or outcome.
- 8.8% (95% CI, 7.9-9.8) of prevention research projects measured more than one leading risk factor for death as an exposure or outcome.
- 24.6% (95% CI, 22.5-26.9) of prevention research projects included a randomized intervention designed to address a leading risk factor or cause of death.

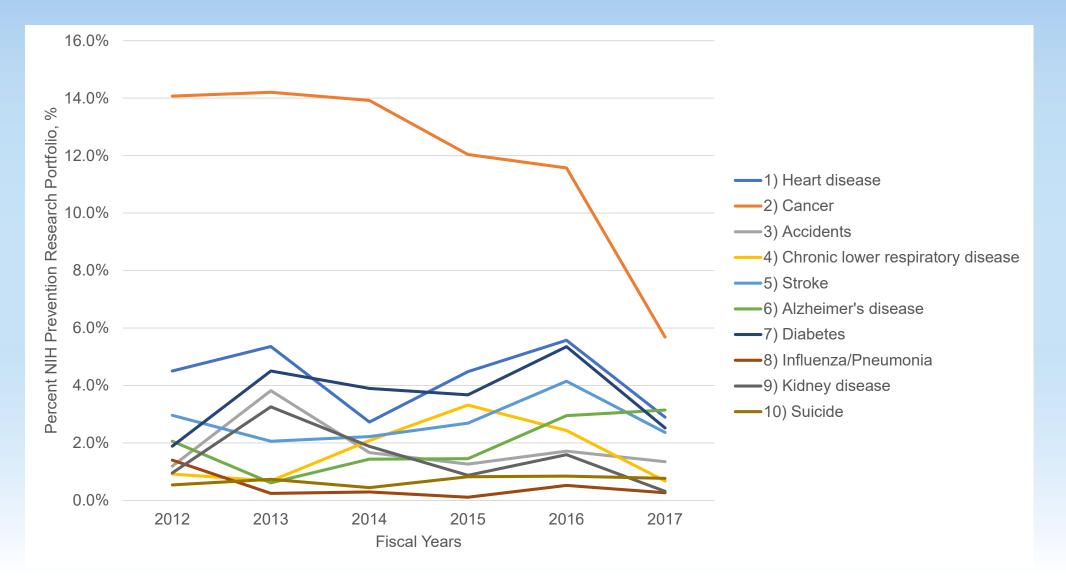
### The Rest of the NIH Prevention Research Portfolio?

Study exposure and/or	Portfolio,	Study exposure and/or	Portfolio,
outcome	% (95% CI)	outcome	% (95% CI)
1) Other	86.0 (83.4-88.4)	14) Sexual behavior	2.8 (2.3-3.4)
2) Genetics	32.0 (28.8-35.4)	15) Chemical/toxin <sup>c</sup>	2.6 (1.9-3.5)
3) Infectious disease <sup>a</sup>	16.7 (14.4-19.2)	16) Violence	2.6 (1.9-3.5)
4) Education/counseling	12.3 (10.9-13.9)	17) Musculoskeletal disease	2.5 (1.6-3.7)
5) Medication/device	11.6 (9.4-14.1)	18) Policy/built environment	2.3 (1.8-2.8)
6) Mental health	10.7 (8.9-12.7)	19) Mortality	2.0 (1.6-2.5)
7) Healthcare delivery	10.0 (8.5-11.6)	20) Kidney disease <sup>d</sup>	1.7 (1.0-2.9)
8) Neurological disease <sup>b</sup>	9.5 (7.6-11.9)	21) Lung disease <sup>e</sup>	1.4 (0.8-2.3)
9) HRQOL	5.6 (4.5-7.0)	22) Heart disease <sup>f</sup>	1.2 (0.9-1.6)
10) Stress	4.0 (3.1-5.2)	23) Surgery	1.2 (0.6-2.2)
11) Vaccine	3.1 (2.0-4.8)	24) Stroke <sup>g</sup>	0.6 (0.4-0.9)
12) Microbiome	3.0 (2.0-4.4)	25) Blood disorder	0.5 (0.3-0.8)
13) GI disease	3.0 (1.8-4.9)	26) Firearms	0.2 (0.1-0.3) <sup>h</sup>

# Trends in prevention research measuring leading risk factors for death in the U.S.



## Trends in prevention research measuring leading causes of death in the U.S.



### Summary of Findings

- During FY12-17, 16.7% of NIH research supported by extramural grants and collaborative agreements focused on primary and secondary prevention in humans, together with methods development to support that research.
  - 51.4% of that portfolio, or 8.6% of the total NIH research portfolio, addressed a leading risk factor <u>or</u> cause of death.
  - 31.4% of that portfolio, or 5.2% of the total NIH research portfolio, addressed a leading risk factor or cause of disability.
  - 3.3% of that portfolio, or 0.6% of the total NIH research portfolio, measured more than one leading cause of death as an exposure or outcome.
  - 8.8% of that portfolio, or 1.5% of the total NIH research portfolio, measured more than one leading risk factor for death as an exposure or outcome
  - 24.6% of that portfolio, or 4.1% of the total NIH research portfolio, included a randomized intervention that addressed a leading risk factor or cause of death.

### **Discussion Questions**

How should NIH respond to these findings?

- Should the prevention research portfolio be reshaped to emphasize projects that address the leading risk factors and causes of death and disability?
- Should the prevention research portfolio be reshaped to emphasize projects that address multiple risk factors or causes of death and disability in the same study?
- Should the prevention research portfolio be reshaped to emphasize the development and testing of preventive interventions to address the leading risk factors and causes of death and disability?
- Other advice?

### Acknowledgments

#### **ODP Team**

Charlene Liggins, Lead Ashley Vargas Luis Ganoza Caballero Erin Ellis Natasha Oyedele

#### Past ODP Team

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