

#### Intervention and Policy Implications for Behavioral Epigenetic Research

Elisabeth Conradt, Ph.D.

Assistant Professor, Developmental Psychology, University of Utah HARC Collaborative Science of Home Visiting 1/31/2018

### Editing the epigenome

- The prenatal and early postnatal period are formative windows when the environment can produce durable, and perhaps reversible, changes in physiology and behavior
- Sensitive period for epigenetic treatment that may program gene expression?

Michael Lübbert Peter A. Jones Editors Epigenetic Therapy of Cancer Precinical Models and Treatment Approaches

2 Springer

## Role of Epigenetics in Treatment

- As an objective tool for assessing the impact of the environment on the epigenome
  - "Health monitoring marker"
- As an objective measure of the progress of an intervention
  - Implies DNA methylation is a mechanism
- As the basis for pharmacological treatment



Loi, Del Savio, & Stupka, 2013; Szyf, Tang, Hill & Musci, 2016; Park & Kobor, 2015; Weaver et al., 2005

# An objective tool for assessing the impact of the environment on the epigenome

- Low licking and grooming → greater
  DNA methylation of NR3c1
- In humans: poor maternal care related to greater DNA methylation of NR3c1?



Liu et al., 1997; Weaver et al., 2005

# Trauma exposure is related to greater DNA methylation of *NR3c1*



CpG Site

Romens, McDonald, Svaren, & Pollak, 2015

# Depression is related to increased DNA methylation of NR3c1, but only if the mother is less sensitive







# An objective tool for assessing the impact of the environment on the epigenome

- Researchers could examine the impact the environment had on the child methylome
- Tool for assessing who is most at risk for developing a "high stress" phenotype?
- Tool to capture inherited initial disadvantage (from birth?)
- Ethical implications: DNA is not destiny, and neither is the epigenome



Loi, Del Savio, & Stupka, 2013; Szyf, Tang, Hill & Musci, 2016; Weaver et al., 2005

# An objective measure of the progress of an intervention

- Following treatment, examine the child methylome
- Are there differences pre-and post intervention? Does the magnitude of these differences predict changes in child and/or caregiver behavior?
- Does changing parenting behavior alter DNA methylation of *NR3c1*?
- Example: Intervention to enhance selfcontrol in African-American men



# Intervention increased self-control, but at a "cost" seen at the epigenetic level



Miller, Yu, Chen, & Brody, 2015

### **Ethical Implications**

- Treatment to reduce stress reactivity in early childhood: Is this the "ideal" infant phenotype?
- The importance of the "fit" between the infant and caregivers
- Infant may be "programmed" prenatally to adapt to a particular postnatal environment



### Genes as Mirrors of Life Experiences

By BENEDICT CAREY NOV. 8, 2010

## Grandma's Experiences Leave a Mark on Your Genes



The Dutch Hunger Winter of 1944 left health effects on people who were in utero at the time.

utch National Archives

Mom's environment during pregnancy can affect her grandchildren

# Why Your DNA May Not Be Your Destiny

By Denise Chow, Assistant Managing Editor I June 4, 2013 12:52pm ET



How A Pregnant Woman's Choices Could Shape A Child's Health

#### Fathers May Pass Down More Than Just Genes, Study Suggests

#### EPIGENETICS

Don't stress dad — it's bad for your kids' health

Lappé, 2016

# Ethical implications: The burden is on the parent (and typically the mother)

- "Careless discussion of epigenetic research on how early life affects health across generations could harm women by implying the burden is on the mother for her child's wellbeing"
- Could increase surveillance and regulation of pregnant women
- Epigenetic research is preliminary and inconsistent



Richardson, Daniels, Gillman, Golden, Kukla, Kuzawa, & Rich-Edwards, 2014

### **Ethical implications**

- Epigenetics may bring back eugenic thinking
  - "DNA is our destiny" → Our ancestors may also make us "who we are" and we can be "poisoned" by the bad choices of our ancestors
  - Concern: If a population is exposed to a pathogenic environment for multiple generations they could be irreparably damaged
  - Racist groups could use epigenetics to claim the "acquired inferiority" of particular groups



Waggoner & Uller, 2015

### Ethical implications: 4 caveats

- 1. Avoid generalizing animal studies to humans without qualification
- 2. Recognize both paternal and maternal influences to the health of the baby
- 3. Convey complexity
- 4. Recognize the role of society
  - 1. Prenatal stress correlates with class, race, and gender



Richardson, Daniels, Gillman, Golden, Kukla, Kuzawa, & Rich-Edwards, 2014

### **Policy Implications**

- Our DNA is not insensitive to social structures and social class
- Reducing social inequalities already at birth
- Having a deeper explanation of precisely how early life stress can become biologically embedded may provide policy-makers with the evidence needed to support policies that support families and young children
- Example: Research briefing in the UK highlights the importance of epigenetic research in policies to improve the nutrition of pregnant women



Park & Kobor, 2015

### Conclusions

- The science of epigenetics reveals that the environment may be more important than we thought
- Compelling evidence that epigenetic processes might be used as a marker of the effects of the environment, as a marker of treatment progress, and it could be leveraged for pharmacological treatment
- We should be careful about what we expect an "ideal" treatment outcome should be
- The science of epigenetics has important policy implications but concerns remain about epigenetic determinism and epigenetic eugenics





