### Before, Between and Beyond Pregnancy: From Concept to Practice

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### **Life-Course Perspective**

A way of looking at life not as disconnected stages, but as an integrated continuum

### **Life Course Perspective**



Lu MC, Halfon N. Racial and ethnic disparities in birth outcomes: a life-course perspective. Matern Child Health J. 2003; 7:13-30.

### **Life Course Perspective**

- Early programming
- Cumulative pathways
- Improving MCH: A life-course perspective

Early Programming



#### Barker Hypothesis Birth Weight and Coronary Heart Disease



Rich-Edwards JW, Stampfer MJ, Manson JE, Rosner B, Hankinson SE, Colditz GA et al. Birth weight and risk of cardiovascular disease in a cohort of women followed up since 1976. Br Med Jr 1997;315:396-400.

#### Barker Hypothesis Birth Weight and Hypertension



Law CM, de Swiet M, Osmond C, Fayers PM, Barker DJP, Cruddas AM, et al. Initiation of hypertension in utero and its amplification throughout life. Br Med J 1993; 306:24-27.

#### Barker Hypothesis Birth Weight and Insulin Resistance Syndrome



Barker DJP, Hales CN, Fall CHD, Osmond C, Phipps K, Clark PMS. Type 2 (non-insulin-dependent) diabetes mellitus, hypertension and hyperlipidaemia (Syndrome X): Relation to reduced fetal growth. Diabetologia 1993; 36:62-67.

#### **Maternal Stress & Fetal Programming**



### Prenatal Stress & Programming of the Brain

#### Prenatal stress (animal model)

- Hippocampus
  - Site of learning & memory formation
  - Stress down-regulates glucocorticoid receptors
  - □ Loss of negative feedback; overactive HPA axis

#### Amygdala

- Site of anxiety and fear
- □ Stress up-regulates glucocorticoid receptors
- Accentuated positive feedback; overactive HPA axis

Welberg LAM, SeckI JR. Prenatal stress, glucocorticoids and the programming of the brain. J Neuroendocrinol 2001;13:113-28.

#### Prenatal Programming of the Hypothalamic-Pituitary-Adrenal Axis



Fig. 1. Schematic representation of the hypothalamic-pituitary-adrenal (HPA) axis. GR, glucocorticoid receptor; MR, mineralocorticoid receptor; PVN, paraventricular nucleus; CRH, corticotropin-releasing hormone; AVP, arginine vasopressin; ACTH, adrenocorticotropic hormone.

Welberg LAM, SeckI JR. Prenatal stress, glucocorticoids and the programming of the brain. J Neuroendocrinol 2001;13:113-28.

## **Epigenetics**

#### VOLUME CONTROLS FOR GENES

THE DNA SEQUENCE is not the only code stored in the chromosomes. So-called epigenetic phenomena of several kinds can act like volume knobs to amplify or mute the effect of genes. Epigenetic information is encoded as chemical attachments to

Gibbs WW The Unseen Genome

the DNA or to the histone proteins that control its shape within the chromosomes. Among their many functions, the epigenetic volume controls muffle parasitic genetic elements, called transposons, that riddle the genome.

Scientific American 200



Bevond

#### **Epigenetics** Same Genome, Different Epigenome



R.A. Waterland, R.A. Jirtle, "Transposable elements: targets for early nutritional effects on epigenetic gene regulation," *Mol Cell Biol*, 23:5293-300, 2003. Reprinted in the New Scientist 2004



#### **Epidemic of Childhood Overweight & Obesity**



Source: National Center for Health Statistics, National Health and Nutrition Examination Survey

Note: Estimate not available for 1976-1980 for Hispanic; overweight defined as BMI at or above the 95<sup>th</sup> percentile ofr the CDC BMI-for-age growth charts

#### Prenatal Programming of Childhood Overweight & Obesity

#### Matern Child Health J DOI 10.1007/s10995-006-0141-8

ORIGINAL PAPER

#### Prenatal Programming of Childhood Overweight and Obesity

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Abstract Objective: To review the scientific evidence for prenatal programming of childhood overweight and obesity, and discuss its implications for MCH research, practice, and policy.

Methods: A systematic review of observational studies examining the relationship between prenatal exposures and childhood overweight and obesity was conducted using MOOSE guidelines. The review included literature posted on PubMed and MDConsult and published between January 1975 and December 2005. Prenatal exposures to maternal diabetes, malnutrition, and cigarette smoking were examined, and primary study outcome was childhood overweight or obesity as measured by body mass index (BMI) for children ages 5 to 21.

Results: Four of six included studies of prenatal exposure to maternal diabetes found higher prevalence of childhood overweight or obesity among offspring of diabetic mothers, with the highest quality study reporting an odds ratio of adolescent overweight of 1.4 (95% CI 1.0–1.9). The Dutch famine study found that exposure to maternal malnutrition in early, but not late, gestation was associated with increased

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Department of Community Health Sciences and the Center for Healthier Children, Families and Communities, UCLA School of Public Health, Box 951772, Los Angeles, CA 90095–1772, USA e-mail: meluwela.edu odds of childhood obesity (OR 1.9, 95% CI 1.5–2.4). All eight included studies of prenatal exposure to maternal smoking showed significantly increased odds of childhood overweight and obesity, with most odds ratios clustering around 1.5 to 2.0. The biological mechanisms mediating these relationships are unknown but may be partially related to programming of insulin, leptin, and glucocorticoid resistance in utero.

Conclusion: Our review supports prenatal programming of childhood overweight and obesity. MCH research, practice, and policy need to consider the prenatal period a window of opportunity for obesity prevention.

Keywords Prenatal programming · Childhood obesity · Overweight · Developmental programming · Fetal programming · Gestational diabetes · Matemal malnutrition · Cigarette smoking

Childhood overweight and obesity is a growing problem in the United States and worldwide. The prevalence of childhood overweight in the U.S. tripled between 1980 and 2000 [1]. Today approximately 1 in 6 (16%) U.S. children are overweight with significant racial-ethnic disparities. For example, nearly 1 in 4 (23%) non-Hispanic black girls ages 6 to 19 are overweight, a prevalence almost twice that of non-Hispanic white girls [1].

Overweight and obesity has significant lifelong consequences on the health and well-being of children [2, 3]. Childhood obesity is associated with early-onset Type II diabetes mellitus, hypertension, metabolic syndrome, and sleep apnea. It is also associated with cognitive or intellectual impairment and social exclusion and stigmatization as parts of a vicious cycle including school avoidance [3]. Childhood obesity tracks strongly into adulthood [4, 5]; obesity beyond

### Prenatal Programming of Childhood Obesity



## **Cumulative Pathways**

Photo: http://www.lam.mus.ca.us/cats/encyclo/smilodon/

### Allostasis: Maintain Stability through Change



McEwen BS. Protective and damaging effects of stress mediators. N Eng J Med. 1998; 338: 171-9.

### Allostastic Load: Wear and Tear from Chronic Stress



McEwen BS. Protective and damaging effects of stress mediators. N Eng J Med. 1998; 338: 171-9.



### **Stressed vs. Stressed Out**

#### Stressed

- Increased cardiac output
- Increased available glucose
- Enhanced immune functions
- Growth of neurons in hippocampus & prefrontal cortex

#### Stressed Out

- Hypertension & cardiovascular diseases
- Glucose intolerance & insulin resistance
- Infection & inflammation
  - Atrophy & death of neurons in hippocampus & prefrontal cortex

### **Allostasis & Allostatic Load**



McEwen BS, Lasley EN. The end of stress: As we know it. Washington DC: John Henry Press. 2002

## **Rethinking Preterm Birth**



### **Sequelae of Preterm Birth**



### Racial & Ethnic Disparities Infant Mortality



### **Racial & Ethnic Disparities Preterm Births < 37 Weeks**



### **Racial & Ethnic Disparities Very Preterm Births < 32 Weeks**



### **Rethinking Preterm Birth**

Vulnerability to preterm delivery may be traced to not only exposure to stress & infection during pregnancy, but host response to stress & infection (e.g. stress reactivity & inflammatory dysregulation) patterned over the life course (early programming & cumulative allostatic load)

### Preterm Birth & Maternal Ischemic Heart Disease



Kaplan-Meier plots of cumulative probability of survival without admission or death from ischemic heart disease after first pregnancy in relation to preterm birth

Improving MCH: A Life-Course Perspective





### Women's Health: A Life-Course Perspective













#### Improving MCH: A Life-Course Perspective

The life-course perspective suggests a need for an expanded approach to improve perinatal health, one that emphasizes not only risk reduction during pregnancy, but also health promotion and optimization before and between pregnancies and, indeed, across the life course. The approach needs to be both clinical and population-based, addressing individual factors as well as social determinants of perinatal health.



#### Improve Healthcare Access Before, Between, Beyond Pregnancy

#### Preconception & interconception care

- Mandated coverage & provision to override exclusions based on preexisting conditions
- Covered service under FEHBP
- Medicaid waiver (e.g. Illinois Healthy Women Initiative)

#### Comprehensive women's healthcare over the life course.

# **Interconception Care**

Quality

Enhance service integration Before, Between, Beyond Pregnancy

Medical home for pregnancy

California's Comprehensive Perinatal Services Program (CPSP)

Wellness homes across the life course

- Hope Street Family Center in Los Angeles
- DC Developing Families Center



#### <u>Early Childhood</u> •Early Head Start •Head Start •UPK •Choir Academy

<u>Child Welfare</u> •Preventive Services •Foster Care Service •Parenting Workshop •Newborn Home Visiting COPS Waiver <u>Legislative Agenda</u> Reauthorize Healthy Start SCHIP Minimum Wage Legislatio Women's Health Financin

> MCH Life-Course Organization

<u>Housing</u> •Home Ownership •Affordable Housing •Base Building- St. Nicks Economic Opportunities •Harlem Works •Financial Literacy •LPN RN Training Program •Union Employment •Micro Lending Savings •Empowerment Zone

<u>Health System</u> Case Management - Title V Funds Health Education - Regionalization Outreach -Harlem Hospital Perinatal Mood Disorders-Birthing Center Interconceptional Care



Wellness

#### **Invest in Wellness**

#### Before, Between, Beyond Pregnancy

#### Wellness Trust

- Fund prevention (VAT, sin tax, etc)
- Set national priorities on prevention (USPSTF)
- effective delivery systems
- develop an information technology backbone
- Incentivize prevention

#### Children's Trust

Family & Community

**Strengthen Families & Communites** *Before, Between, Beyond Pregnancy* 

Fatherhood initiatives

Best start zones

### **Reproductive Social Capital**

Features of social organization (e.g. networks, norms, and social trust) that facilitate coordination and cooperation to promote reproductive health within a community.

### **Reproductive Social Capital**

100 Intentional Acts of Kindness toward a Pregnant Woman

# Education

### **Educational Development**

- 1. Preconception and prenatal care
- 2. Parenting education
- 3. Child care
- 4. Universal Preschool
- 5. Early Head Start and Head Start
- 6. K-12 small class size, teacher quality, standards
- 7. After school and summer programs
- 8. Youth development
- 9. Health education/physical education
- 10. Comprehensive school health clinics

### **Educational Development**

Early Childhood programs After-school programs Summer Programs + Comprehensive school health clinics \$ 156 billion

# Environment

"Now that you're here, the word of the Lorax seems perfectly clear. **UNLESS** someone like you Cares a whole awful lot, Nothing is going to get better. It's not. "SO... Catch!" calls the Once-ler. He lets something fall. "It's a Truffula Seed. It's the last one of all! You're in charge of the last of the Truffula Seeds. And Truffula Trees are what everyone needs. Plant a new Truffula. Treat it with care. Give it clean water. And feed it fresh air. Grow a forest. Protect it from axes that hack. Then the Lorax And all of his friends May come back."

Policy

## **Maternity Leave**

| Country        | Weeks before<br>delivery | Weeks after<br>delivery | Salary |
|----------------|--------------------------|-------------------------|--------|
| USA            | 4                        | 6                       | No     |
| UK             | 10 + 5 days              | 25 + 5 days             | Yes    |
| France         | 6 + 3 days               | 8 + 5 days              | Yes    |
| Germany        | 7 + 3 days               | 8                       | Yes    |
| Italy          | 8 + 5 days               | 12 + 6 days             | Yes    |
| Spain          | 16                       | 6                       | Yes    |
| Finland        | 4                        | 37                      | Yes    |
| Czech Republic | 6                        | 22                      | Yes    |

#### **Expand Parental Leave**

Only 45% of parents working in private sector have guaranteed unpaid parental leave through FMLA

Only 5% have paid parental leave

#### Expand paid leave

- Unemployment insurance
- Temporary disability insurance
- New social insurance
- New cash benefits program

# We Can Do Better

#### The definition of insanity is doing the same thing over and over and expecting different results



#### □ "We must become the change we want to see."

#### - MOHANDAS GANDHI

