



## **Prof. Atul Singhal**

University College London, Institute of Child Health, UK

---

Atul Singhal is Great Ormond Street Hospital Children's Charity Professor of Paediatric Nutrition at University College London, Institute of Child Health, and Honorary Consultant Paediatrician at Great Ormond Street Hospital. He graduated in Medicine from the Royal Free Hospital, London in 1986 and has been a consultant in paediatrics since 1998. Previously, he was the Director and Deputy Director of the Childhood Nutrition Research Centre, UCL Institute of Child Health.

He has broad interests in paediatric nutrition, but his current research focuses on the influence of early nutrition for long-term health, the effects of nutritional interventions to reduce long-term cardiovascular risk, and nutritional interventions for obesity.

# Developmental origins of health and disease; where are we now

The idea that nutrition may act during a critical window early in development to permanently affect, or 'program'<sup>1</sup>, long-term health first emerged from studies in animals<sup>2</sup> but is now strongly supported in humans. Nutrition throughout the life course, including fetal life, infancy, the preschool, or toddlers years, and in adolescence has been shown to impact on long-term health, a hypothesis known as the developmental origins of health and disease<sup>3</sup>.

Some of the earliest evidence for nutritional programming in humans was obtained from observational studies from the late 1980's linking low birth weight with adult cardiovascular disease<sup>4</sup>. Now, both under and over-nutrition during pregnancy has been associated with adverse outcomes in the offspring such as neuro-cognitive impairment, obesity and an increased risk of cardiovascular disease<sup>4</sup>. Importantly, recent experimental (randomised) evidence supports a causal link between over-nutrition during pregnancy and an increased risk of obesity in the offspring<sup>5</sup>.

Postnatally, the strongest evidence for nutritional programming has been obtained for the long-term benefits of breast-feeding. Breast-feeding, not only has benefits for short-term health, but has been shown to have major advantages for long-term cognitive function<sup>6,7</sup>, atopic disease<sup>8</sup>, bone health<sup>9</sup>, and risk of obesity and cardiovascular disease<sup>10,11</sup>. There is particularly strong evidence that breast-feeding can improve later cognitive development, a hypothesis supported by several systematic reviews, evidence of a dose-response association<sup>12</sup>, data from a cluster randomised trial<sup>13</sup>, as well as evidence of benefits of breast-feeding on visual development<sup>13</sup> and structure of the brain<sup>14,15</sup>.

The mechanisms underlying the cognitive benefits of breast feeding are uncertain, but previous studies have investigated differences between human milk and formula in concentrations of biologically active factors such as nucleotides, lipids, and the milk fat globular membrane. More recent research focuses on the impact of breast milk nutrients (e.g. tryptophan, nucleotides and long-chain fatty acids) on sleep modulation and early brain development in infancy<sup>16</sup>.

This presentation will give an update on the development origins of health and disease<sup>11</sup> including an overview of the role of human milk intake on long-term cognitive function and the possible mechanisms involved. It will highlight the key role of promoting exclusive breast-feeding<sup>13</sup>, the potential impact of specific breast milk nutrients, and the importance of

experimental (randomised) studies in interpreting the effects of early nutrition on later health. Finally, it will consider the implications of nutritional programming for nutritional, clinical and public health practice.

In interpreting the effects of early nutrition on later health. Finally, it will consider the implications of nutritional programming for nutritional, clinical and public health practice.

## References

- Lucas A. Programming by early nutrition in man. The childhood environment and adult disease. CIBA Foundation symposium 156. Bock GR and Whelan J. Wiley, Chichester; 1991; 38-55.
- McCay CM. Is longevity compatible with optimum growth? *Science* 1933; 77: 410-411.
- Bateson P, et al. Developmental plasticity and human health. *Nature* 2004; 430: 419-21.
- Barker DJP. Fetal origins of coronary heart disease. *BMJ* 1995; 311: 171-4.
- Patel N, et al. Infant adiposity following a randomised controlled trial of a behavioural intervention in obese pregnancy. *Int J Obes (Lond)*. 2017 Jul;41(7):1018-1026.
- Isaacs EB, et al. Early diet and general cognitive outcome at adolescence in children born at or below 30 weeks gestation. *J Pediatr* 2009; 155: 229-234.
- Belfort MB, et al. Infant feeding and childhood cognition at ages 3 and 7 years: effects of breastfeeding duration and exclusivity. *JAMA Pediatr* 2013; 455: 836-888.
- Kramer MS, et al. PROBIT Study Group (Promotion of Breastfeeding Intervention Trial). Promotion of Breastfeeding Intervention Trial (PROBIT): a randomized trial in the Republic of Belarus. *JAMA*. 2001;285:413-420.
- Fewtrell MS, et al. Early diet and peak bone mass: 20 year follow-up of a randomized trial of early diet in infants born preterm. *Bone* 2009; 45:142-9.
- Patro-Golab B, et al. Nutritional interventions or exposures in infants and children aged up to 3 years and their effects on subsequent risk of overweight, obesity and body fat: a systematic review of systematic reviews. *Obes Rev*. 2016 ; 17:1245-1257.
- Singhal A. Early Life Origins of Obesity and Related Complications. *Indian J Pediatr*. 2017 Dec 16.
- Isaacs EB, et al. Impact of breast milk on intelligence quotient, brain size, and white matter development. *Pediatr Res* 2010;67: 357-62.
- Kramer MS, et al. Breastfeeding and child cognitive development: new evidence from a large randomized trial. *Arch Gen Psychiatry*. 2008 May;65(5):578-84.
- Singhal A, et al. Infant nutrition and stereoacuity at age 4-6 years. *Am J Clin Nutr* 2007; 85: 152 -159.
- Sean CL, et al. Breastfeeding and early white matter development: A cross-sectional study. *NeuroImage* 2013; 82: 77-86.
- Schneider N, et al. Diet and nutrients in the modulation of infant sleep: A review of the literature. *Nutr Neurosci*. 2016 Nov 21:1-11. follow-up of a randomized trial of early diet in infants born preterm. *Bone* 2009; 45:142-9.
- Patro-Golab B, et al. Nutritional interventions or exposures in infants and children aged up to 3 years and their effects on subsequent risk of overweight, obesity and body fat: a systematic review of systematic reviews. *Obes Rev*. 2016 ; 17:1245-1257.
- Woo Baidal JA, et al. Risk factors for childhood obesity in the first 1,000 days; a systematic review. *Am J Prev Med* 2016; 50: 761-779.
- Singhal A. The role of infant nutrition in the global epidemic of non-communicable disease. *Proc Nutr Soc* 2016; 75: 162-168.
- Isaacs EB, et al. Impact of breast milk on intelligence quotient, brain size, and white matter development. *Pediatr Res* 2010;67: 357-62.
- Kramer MS, et al. PROBIT Study Group (Promotion of Breastfeeding Intervention Trial). Promotion of Breastfeeding Intervention Trial (PROBIT): a randomized trial in the Republic of Belarus. *JAMA*. 2001;285:413-420.
- Singhal A, et al. Infant nutrition and stereoacuity at age 4-6 years. *Am J Clin Nutr* 2007; 85: 152 -159.
- Sean CL, et al. Breastfeeding and early white matter development: A cross-sectional study. *NeuroImage* 2013; 82: 77-86.
- Singhal A, et al. Early origins of cardiovascular disease; is there a unifying hypothesis? *Lancet* 2004; 363:1642-5.