

Advancing Preconception Care: Moving from Evidence to Implementation



UNICEF photo from Burundi

Helena Pachón, PhD, MPH 22 September 2015 7th International Conference on the Prevention of Birth Defects and Disabilities in the Developing World



Acknowledgements

- Alina Flores
- Cindy Moore
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Main Messages





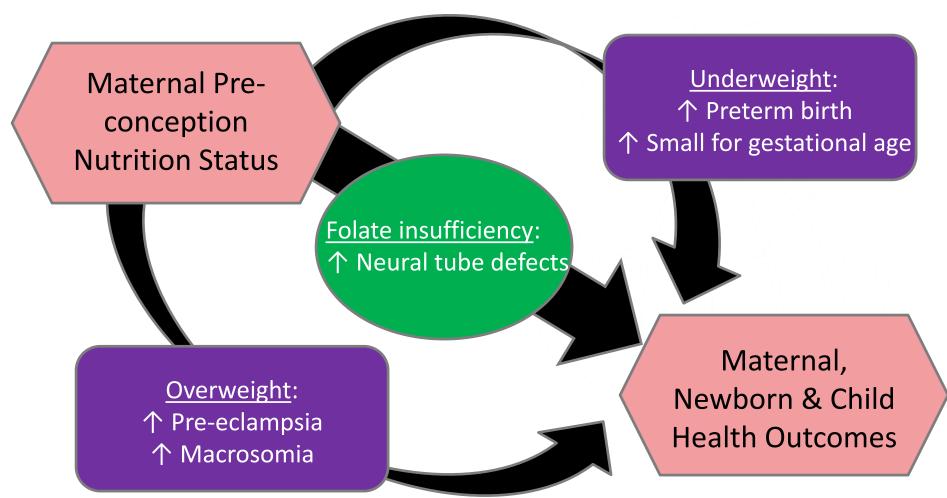


3. With all folic acid interventions, we should aim to reach a birth prevalence of ~6 neural tube defects/10,000





Role of Nutrition in Preconception Health

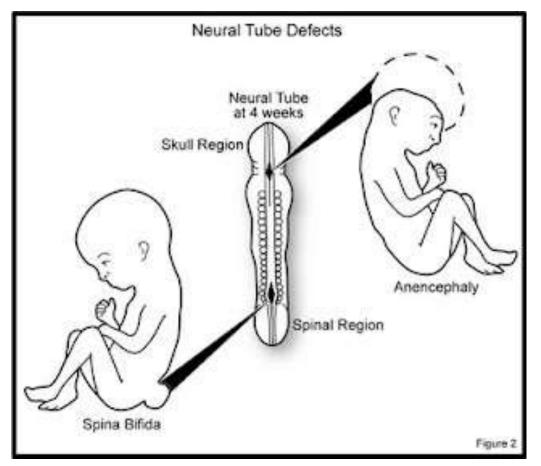


Dean 2014



Neural Tube Defects (NTDs)

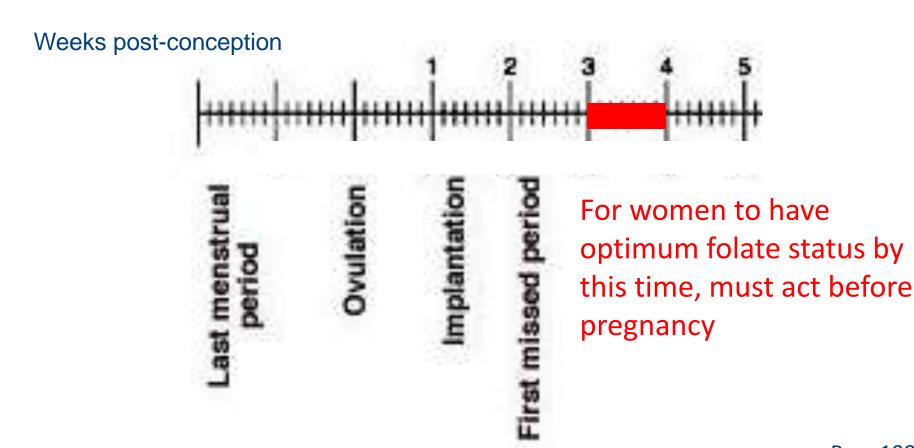
Birth defect affecting the brain and spinal cord



http://www.thescienceofpregnancy.id.au



Neural Tube Closes ~4 Weeks after Conception





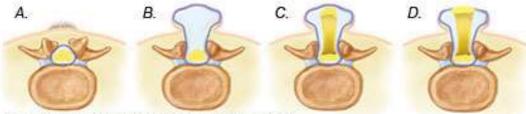
Spina Bifida





Normal newborn vertebra

- A. Spina bifida occulta
- B. Spina bifida with meningocele
- C. Spina bifida with meningomyelocele
- D. Spina bifida with myeloschisis



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Anencephaly

WHO/CDC/ICBDSR 2014



~320,000 Pregnancies Affected with Neural Tube Defects Annually



Global neural tube defects birth prevalence: 24/10,000

March of Dimes 2006





Neural Tube Defects are Preventable with Folic Acid

TABLE IV---PREVALENCE OF NEURAL TUBE DEFECTS (NTD) ACCORDING TO RANDOMISATION GROUP: MAIN ANALYSIS BASED ON ALL WOMEN RANDOMISED WHO HAD AN INFORMATIVE PREGNANCY CLASSIFIED ACCORDING TO RANDOMISATION GROUP (INTENTION-TO-TREAT ANALYSIS)

			AI	women	Women not already pregnant at randomisation*	
Randomisation group			Relative risk: folic acid <i>vs</i>			Relative risk: folic acid vs
	Folic acıd	Other vitamins	NTD/all	non-folic acid (95% CI)	NTD/all	non-folic acid (95% CI)
	+	_	2/298	08())	2/258 2/256 $5/514(1)$	
	+	+	4/295	0.28 (0.12-0.71)	3/256	0.28 (0.11-0.75)
		-	$\frac{13/300}{21/602}$ 21/602 (3-		$\frac{11/260}{7/257}$ 18/517 (3)	
1	- + 8/302 (21/002 (25/0))			7/257		

*First day of last menstrual period was 14 days or more after date of randomisation

72% of neural tube defects were prevented through the administration of folic acid to women



WHO Recommendations

Prevention of neural tube defects

INTEGRATED MANAGEMENT OF PREGNANCY AND CHILDBIRTH (IMPAC)

Standards for Maternal and Neonatal Care

The standard

All women, from the moment they begin trying to conceive until 12 weeks of gestation, should take a folic acid supplement. Women who have had a fetus diagnosed as affected by a neural tube defect (NTD) or have given birth to a baby with NTD should receive information on the risk of recurrence, be advised on the protective effect of periconceptional¹ folate supplementation and be offered high-dose supplementation.

Aim

To prevent NTDs and other congenital malformations in the fetus.

** *

1.5

WHO 2007



WHO Recommendations

- Women trying to conceive need 400 µg folic acid daily, starting two months before the planned pregnancy.
- Pregnant women need to continue taking 400 µg folic acid daily until they are <u>12 weeks pregnant</u>.





WHO Recommendations

 Pregnant women who have previously had a baby with a NTD, have diabetes, or who are under anticonvulsant treatment need 5 mg folic acid daily plus increased food folate intake.



http://www.aquila-style.com/wp-content/uploads/2014/07/shutterstock 21.545cc160715.original-e1406446683307.jpg



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SUSTAINABLE DEVELOPMENT

NOWLEDGE PLATFORM

Ensure healthy lives and promote well-being for all at all ages

HLPF PROCESSES & UN SYSTEM

TARGETS

3.1

By 2030, reduce the global maternal mortality ratio to less than 70 per 100,000 live births

3.2

By 2030, end preventable deaths of newborns and children under 5 years of age, with all countries aiming to reduce neonatal mortality to at least as low as 12 per 1,000 live births and under-5 mortality to at least as low as 25 per 1,000 live births

3.3

By 2030, end the epidemics of AIDS, tuberculosis, malaria and neglected tropical diseases and combat hepatitis, water-borne diseases and other communicable diseases

HOME

SDGS & TOPICS

3.4

By 2030, reduce by one third premature mortality from non-communicable diseases through prevention and treatment and promote mental health and well-being

3.5

Strengthen the prevention and treatment of substance abuse, including narcotic drug abuse and harmful use of alcohol

3.6

By 2020, halve the number of global deaths and injuries from road traffic accidents

3.7

RELEVANT TOPICS

STAKEHOLDER ENGAGEMENT



Health and population

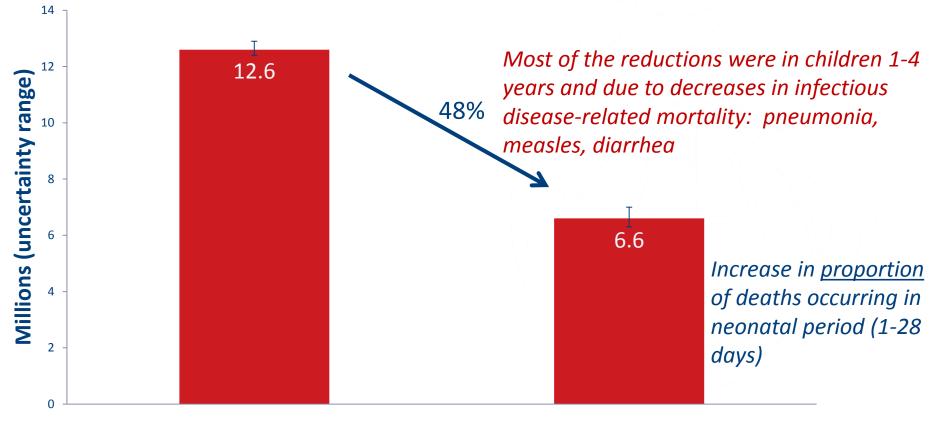


National Sustainable Development Strategies (NSDS)

https://sustainabledevelopment.un.org/topics



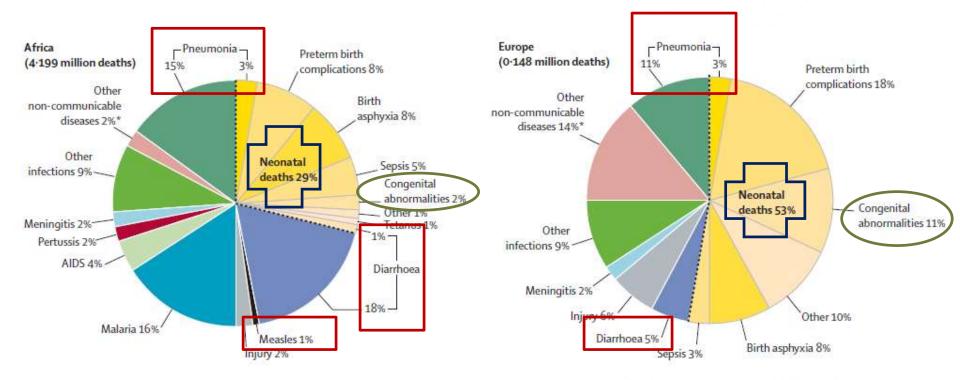
Globally: Annual Deaths Decreased in Children <5 Years



Bhutta 2013, Black 2010



When Mortality from Infections is Lower, Burden from Birth Defects* is Higher



<u>Infection deaths</u> due to pneumonia (15% + 3%) + measles (1%) + diarrhea (18% + 1%) = **38%**

* Birth defects & congenital anomalies are used synonymously

<u>Infection deaths</u> due to pneumonia (11% + 3%) + measles (0%) + diarrhea (5%) = **19%**

Black 2010, Bhutta 2013, WHO 2015

Public Health Strategies to Reduce the Risk of Neural Tube Defects



Provide Vitamin B9



www.breakingmuscle.com

FOOD FOLATE

FOLIC ACID

Suitor 2000



Sources of Food Folate



nttps://ljayhealth.files.wordpress.com/2012/02/spinach





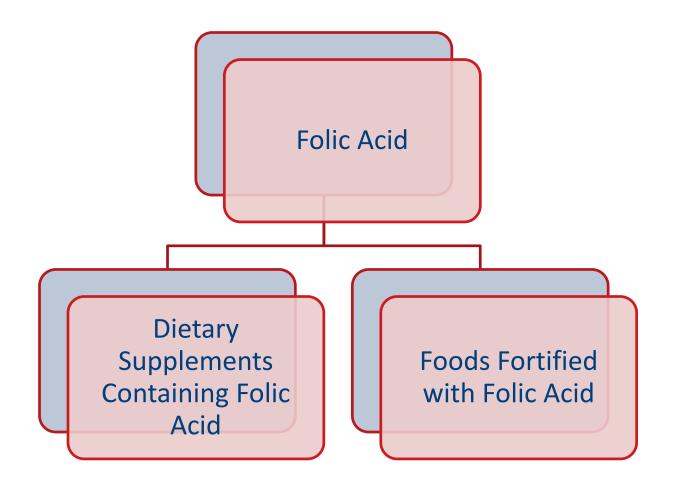


WHO recommendations to prevent neural tube defects: folic acid (primarily)

Suitor 2000, WHO 2007



Sources of Folic Acid



Suitor 2000



Dietary Supplements Containing Folic Acid

Women planning a pregnancy were given daily

- folic acid and other nutrients (in the vitamin group), or
 - other nutrients (in the trace-element group)

000	1.	15
X2	3	5
K	Folic	Seid Y

Table 3. Congenital Malformations, According to Study Group.

MALFORMATION	VITAMIN GROUP	TRACE- ELEMENT GROUP	
	nu	mber	
Neural-tube defect	0	6	P = 0.29

Folic acid supplements given daily to women reduce the first occurrence of a neural tube defect Czeizel 1992

http://www.cdc.gov/ncbddd/folicacid/images/recommend.i



Folic Acid Supplement Use is Low

- Assessed in 49 studies from 22 countries
- Peri-conceptional use ranged from
 0.5% (Italy) - 52%
 (Netherlands)



In practice, folic acid supplement use is low

Ray 2004



Many Pregnancies are Unplanned

"Almost half of all pregnancies worldwide, estimated to be over 100 million annually, are unintended or mistimed."



If women are not planning pregnancy, they may not take supplements in peri-conceptional period

Thurman 2011



Folic Acid Supplementation AND Fortification

- Compared NTD levels before and after recommendations were issued for folic acid supplementation (~1992)
- 13 birth defects registries; 9 European countries + Israel
- "The issuing of recommendations on folic acid [supplementation] was followed by no detectable improvement in the trends of incidence of neural tube defects."

All women capable of becoming pregnant should be encouraged to take folic acid supplements

NTD, neural tube defect

AND...



Food Fortification with Folic Acid

Fortification: the addition of nutrients to foods during processing





Without behavior change, women continue to eat foods, now fortified with folic acid. Extra folic acid will improve women's folate status. Women with unplanned pregnancies will have optimal folate status to prevent NTDs.

NTDs, neural tube defects

WHO/FAO 2006

Countries with Grain Fortification Mandates

84 countries require fortification of wheat flour, maize flour, and/or rice

Wheat flour: 83 August 2015. Source: Food Fortification Initiative. Maize flour: 14 http://www.ffinetwork.org/global_progress/index.php

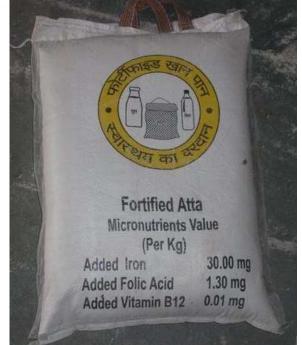
Rice: 6

To request data, e-mail info@ffinetwork.org



Global Experience of Grain Fortification with Folic Acid

- Mandate fortification with <u>folic acid</u>
 - Wheat flour: 79 of 84 countries
 - Maize flour: 12 of 14 countries
 - Rice: 4 of 6 countries
- Folic acid levels required
 - Wheat flour: 0.5-3.3 mg/kg
 - Maize flour: 0.5-2.5 mg/kg
 - Rice: 1-3.08 mg/kg



FFI 2015

mg/kg = parts per million (ppm)



Grain Fortification in Africa

African Countries with Grain Fortification Mandates

Wheat flour (n=26)

- Benin •
- **Burkina Faso** •
- Burundi •
- Cameroon •
- Cape Verde •
- Congo •
- Cote d'Ivoire •

Morocco

Niger

Nigeria

Rwanda

Senegal

Tanzania

Uganda

Togo

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- Djibouti •
- Egypt •
- Ghana •
- Guinea •
- Kenya •
- Liberia •
- Malawi •
- Mali •
- Mauritania •

100 Sierra Leone South Africa

Maize flour (n=8)

- Burundi •
- Kenya •
- Malawi •
- Nigeria •
- Rwanda •
- South Africa •
- Tanzania •
- Uganda •

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Rice (n=0)
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African Experience of Grain Fortification with Folic Acid

Mandate fortification with <u>folic acid</u>

- Wheat flour: 24 of 26 countries (*Exceptions: Congo, Nigeria*)
- Maize flour: 7 of 8 countries (Exception: Nigeria)



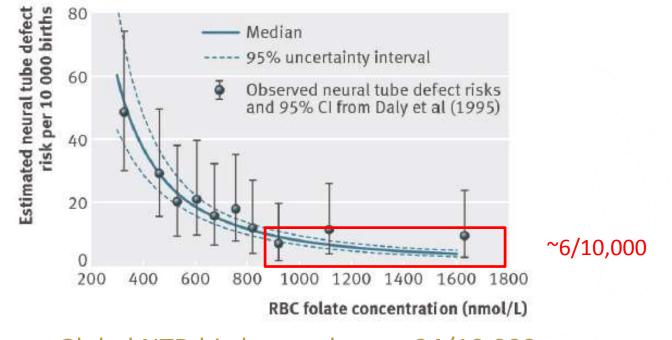
FFI 2015; maps from es.Wikipedia.org





6/10,000: Lowest Birth Prevalence of NTDs with Folic Acid

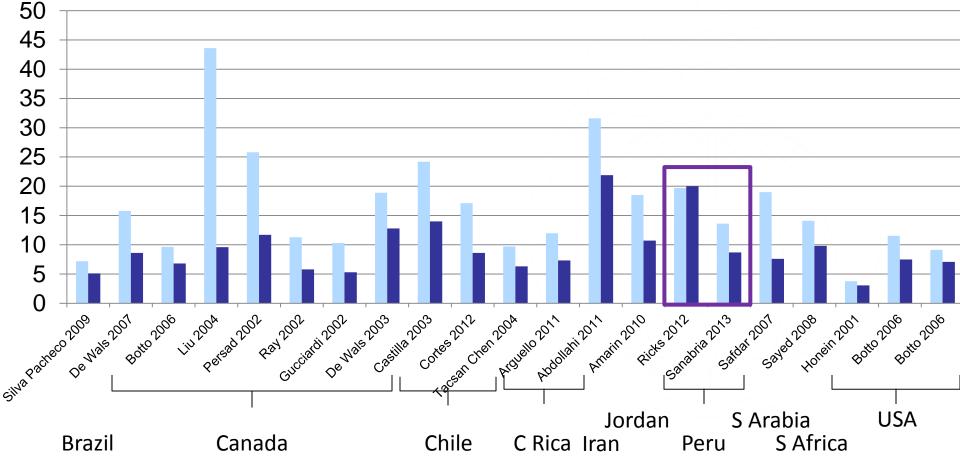
Optimal population red blood cell (RBC) folate concentration to prevent neural tube defects (NTDs)



Global NTD birth prevalence: 24/10,000 Goal: All countries' NTD birth prevalence to reach 6/10,000 Crider 2015, March of Dimes 2006

Reductions in Neural Tube Defects (NTDs) after Flour Fortification with Folic Acid was Initiated

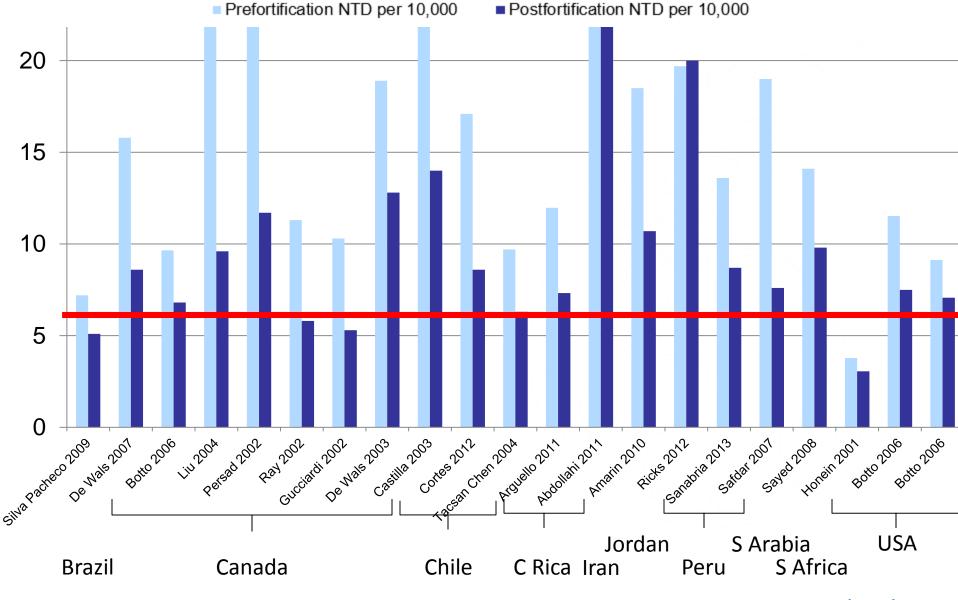
Prefortification NTD per 10,000
Postfortification NTD per 10,000



Fortification of wheat flour <u>+</u> maize flour; folic acid in flour ranged from 1.2-2.2 mg/kg

FFI 2012, updated 2015

Goal: All Countries' NTD Birth Prevalence to Reach 6/10,000



FFI 2012, updated 2015

38,417 Neural Tube Defects Prevented Annually

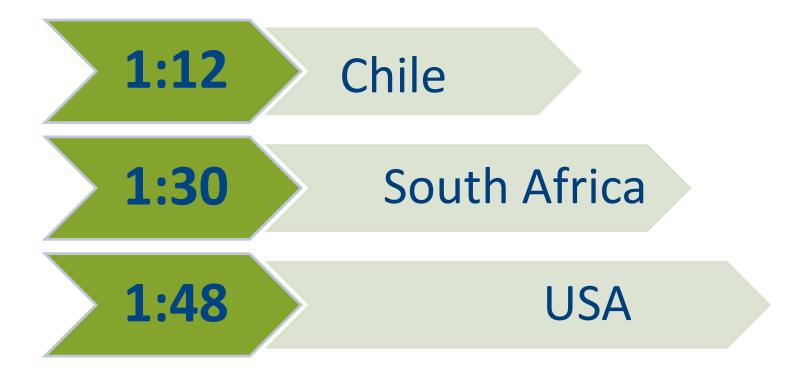
Dr. Vijaya Kancherla to present updated estimates in Symposia 9 (Thursday)

Globally an estimated 38,417 birth defects were prevented in 2012 – an average of 105 a day – where flour was fortified with folic acid

Youngblood 2013



Favorable Cost:Benefit Ratio for Fortification Preventing Spina Bifida

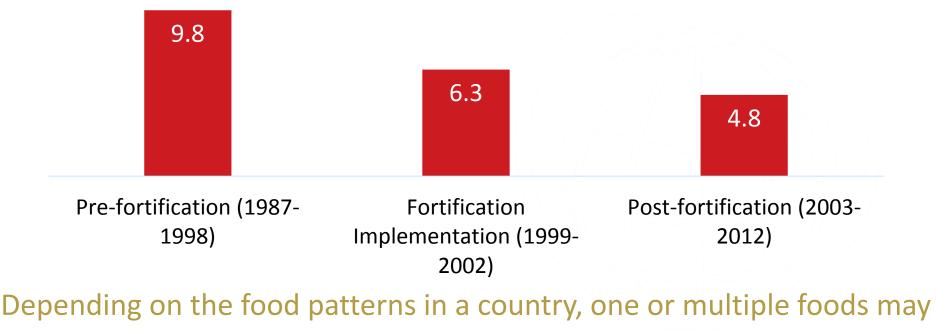


Llanos 2007, Sayed 2008, Grosse 2005



Costa Rica Experience with Multiple Fortified Foods

Neural Tube Defects per 10,000 Live Births



need to be fortified with folic acid to have maximum benefit

Foods fortified with folic acid: wheat flour, maize flour, dairy products, rice

Barboza 2014



Can have Greater Public Health Impact

~25% of FAP SBA is being prevented through flour fortification with folic acid

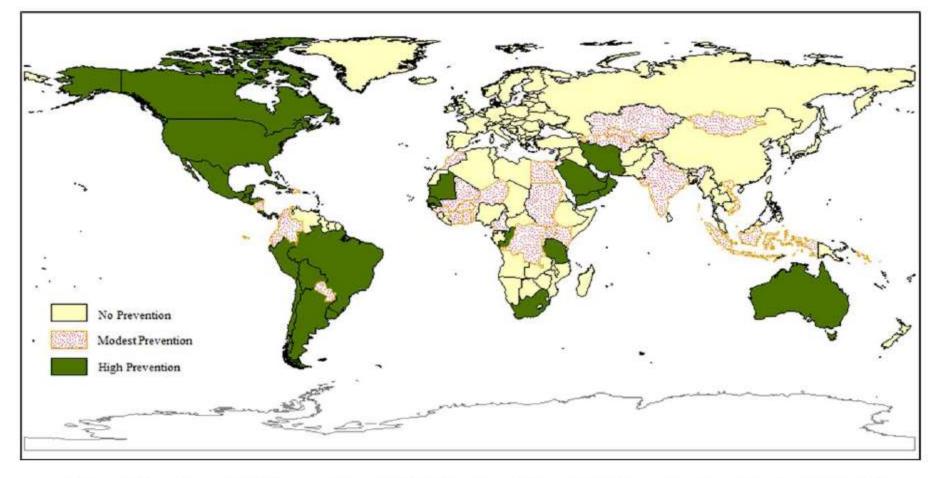


Figure 1. The status of global prevention of folic acid-preventable spina bifida and anencephaly (model II), 2012.

FAP SBA, folic acid-preventable spina bifida and anencephaly (2 types of neural tube defects)

Youngblood 2013



Increasing Price of Folic Acid



www.phillyliving.com



Concerns with Folic Acid

Fortification with folic acid does not

- Increase the incidence of cancer
- Mask vitamin B12 deficiency
- Cause dietary folic acid to exceed the Tolerable Upper Intake Level

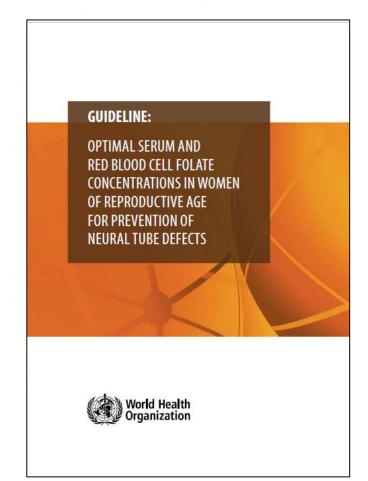


Vollset 2012, Mills 2003, MPI 2012, Yang 2010, Yeung 2011



Limited Laboratory Capacity to Analyze RBC Folate

- WHO guidelines: RBC folate to estimate NTD risk
- ~3 laboratories with capacity to conduct RBC folate analyses using gold standard measure

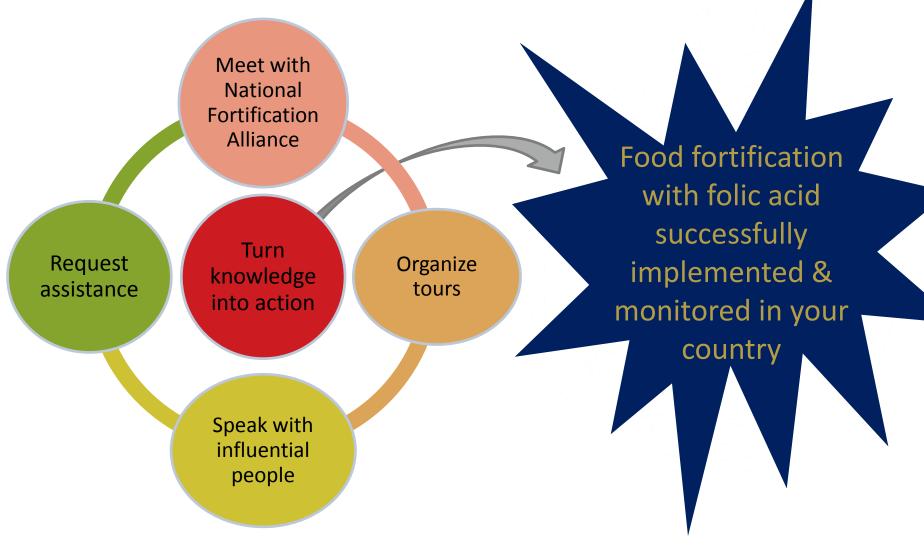




What You Can Do



Your Role





Conclusions (1)

- Neural tube defect risk can be reduced if women have optimum folate status before and in early pregnancy
- 2. We can only meet global infant mortality targets by reducing birth defects
- With folic acid interventions, we should aim to reach a birth prevalence of ~6 neural tube defects/10,000



Conclusions (2)

- Food fortification is the best public health approach to deliver folic acid to women in the peri-conceptional period
- You should act to ensure food fortification with folic acid is implemented & monitored in your country



For More Information

- **Smarter Futures**
- Food Fortification Initiative (FFI)
- International Federation For Spina Bifida and
- Hydrocephalus
- Ronald Afidra, Smarter Futures & FFI
- Helena Pachón, FFI

www.smarterfutures.net/

www.FFInetwork.org

http://www.ifglobal.org/en/

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