

Supplemental Appendix

Effect of folic acid supplementation in pregnancy on preeclampsia - FACT

Shi Wu Wen, Ruth Rennicks White, Natalie Rybak, Laura Gaudet, Stephen Robson, William Hague, Donnette Simms-Stewart, Guillermo Carroli, Graeme Smith, William D. Fraser, George Wells, Sandra T. Davidge, John Kingdom, Doug Coyle, Dean Fergusson, Daniel J Corsi, Josee Champagne, Elham Sabri, Tim Ramsay, Ben Willem J. Mol, Martijn A Oudijk, Mark C Walker, on behalf of the FACT Collaborating Group

Table of Contents

1. Table S1: List of Participating Countries and Sites (Total Recruited)
2. Table S2: Reported adverse events between the two arms
3. Table S3: Comparison of maternal inclusion criteria between the two arms
4. Table S4: Comparison of primary outcome by country

Table S1 – List of Participating Countries and Sites (Total Recruited)**CANADA (1207)**

Centre Hospitalier Universitaire de Québec (225)
Centre Hospitalier Universitaire Sainte-Justine (31)
Children's & Women's Health Centre of British Columbia (16)
Foothills Medical Centre (35)
Health Sciences Centre – General Hospital, Newfoundland (29)
Hôpital Saint-Luc, CHUM (133)
Kingston General Hospital & Queen's University at Kingston (71)
Lois Hole Hospital (31)
London Health Sciences Centre – Victoria Hospital (23)
McMaster University Medical Centre (161)
Regina General Hospital (8)
Royal Victoria Hospital (1)
Saint John Regional Hospital (2)
Sault Area Hospital (2)
St. Boniface General Hospital (16)
St-Mary's Hospital (34)
St-Paul's Hospital (6)
Sunnybrook Research Institute—Sunnybrook Health Sciences Centre (21)
The Dr. Everett Chalmers Regional Hospital (52)
The Moncton Hospital (102)
The Ottawa Hospital—Civic Campus (33)
The Ottawa Hospital—General Campus (96)
The University of Manitoba (79)

UNITED KINGDOM (764)

49 Marine Avenue Surgery (6)
Burnley General Hospital (18)
Darlington Memorial Hospital (7)
Fairfield General Hospital (3)
Gateshead Queen Elizabeth Hospital (28)
Guy's & St Thomas' Hospital (66)
Hinchinbrooke Hospital (8)
James Cook University Hospital (54)
Leighton Hospital (20)
Lincoln County Hospital (9)
Newcastle Upon Tyne Hospital (76)
New Cross Hospital (5)
Norfolk & Norwich (18)
North Cumbria University Hospital (5)
North Manchester General Hospital (4)
North Tyneside General Hospital (21)
Nottingham City Hospital (2)
Queen's Medical Centre (13)
Rochdale Infirmary (1)
Royal Blackburn Hospital (16)
South Tyneside District Hospital (43)
St George's Hospital (50)
Sunderland Royal Hospital (122)
The Hillingdon Hospital (7)
The Royal Oldham Hospital (2)
University Hospital of North Durham (28)
University Hospital of North Tees (33)
Wansbeck General Hospital (31)
Warrington Hospital (11)
West Cumberland Hospital (21)
West Middlesex University Hospital (36)

AUSTRALIA (310)

Ipswich Hospital (92)
Nepean Hospital (7)
Sunshine Hospital (35)
The Royal Women's Hospital (105)
The Townsville Hospital (5)
The Women's and Children's Hospital (66)

ARGENTINA (122)

CEMIC (12)
Hospital Escuela Eva Perón (12)
Hospital Provincial (17)

Hospital Roque Saenz Pena (24)
 Hospital Iturraspe (8)
 Maternidad Martin (47)
 Sanatorio de la Mujer (2)

JAMAICA (61)

Spanish Town Hospital (20)
 The University of West Indies—Mona Campus (40)
 Victoria Jubilee Hospital (1)

Table S2 - Reported adverse events between the two arms

Adverse Events						
	n	%	n	%	RR (95% CI)	p value
Infants						
Any Event	289	20.73	318	22.75	0.89 (0.76 to 1.03)	0.12
0	1105	79.27	1080	77.25		
1-2	268	19.23	279	19.96		
3 and more	21	1.51	39	2.79		
Mothers						
Experiment n=1227		Placebo n=1236				
Any Event	961	78.32	968	78.32	1.00 (0.96 to 1.04)	1.00
0	266	21.68	268	21.68		
1-2	368	29.99	355	28.72		
3 and more	593	48.33	613	49.6		
Severe Adverse Event						
	n	%	n	%	RR (95% CI)	p value
Infants						
Any Event	116	8.33	109	7.8	1.00 (0.75 to 1.32)	0.97
0	1277	91.67	1289	92.2		
1	111	7.97	108	7.73		
2	5	0.36	1	0.07		
Mothers						
Experiment n=1227		Placebo n=1236				
Any Event	227	18.5	195	15.78	1.17 (0.99 to 1.40)	0.07
0	1000	81.5	1041	84.22		
1	186	15.16	156	12.62		
2 and more	41	3.34	39	3.16		

Table S3. Comparison of maternal inclusion criteria between the two arms

Criteria	Folic Acid n=1144		Placebo n=1157		RR (95% CI)	P value
	n	%	n	%		
Pre-existing hypertension	43	22.63	53	22.75	0.99 (0.70 to 1.42)	0.98
Diabetes	30	18.18	22	15.07	1.21 (0.73 to 2.00)	0.46
Twin pregnancy	37	17.21	21	9.86	1.75 (1.06 to 2.88)	0.03
PE in previous pregnancy	63	21.21	66	23.24	0.91 (0.67 to 1.24)	0.56
BMI ≥ 35 kg/m ²	65	11.61	77	12.54	0.93 (0.68 to 1.26)	0.62

Table S4. Comparison of primary outcome by country

Country	Folic Acid n=1144		Placebo n=1157		RR (95% CI)	P value
	n	%	n	%		
Canada	83	14.69	75	12.89	1.14 (0.85 to 1.52)	0.3755
Australia	27	18.12	29	20.86	0.87 (0.54 to 1.39)	0.5568
Argentina	11	19.64	10	17.54	1.12 (0.52 to 2.43)	0.7743
West India	5	20.83	4	13.33	1.56 (0.47 to 5.19)	0.4624
UK	43	12.29	38	10.89	1.13 (0.75 to 1.70)	0.5638