

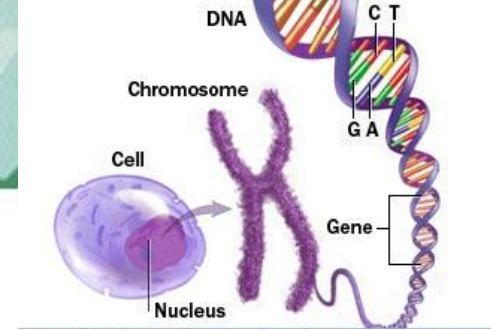
# Obstetrical Outcome among Pregnancies Conceived after Preimplantation Genetic Diagnosis (PGD)

**Raoul Orvieto**  
**Baruch Feldman**  
**Marine Wiesel**

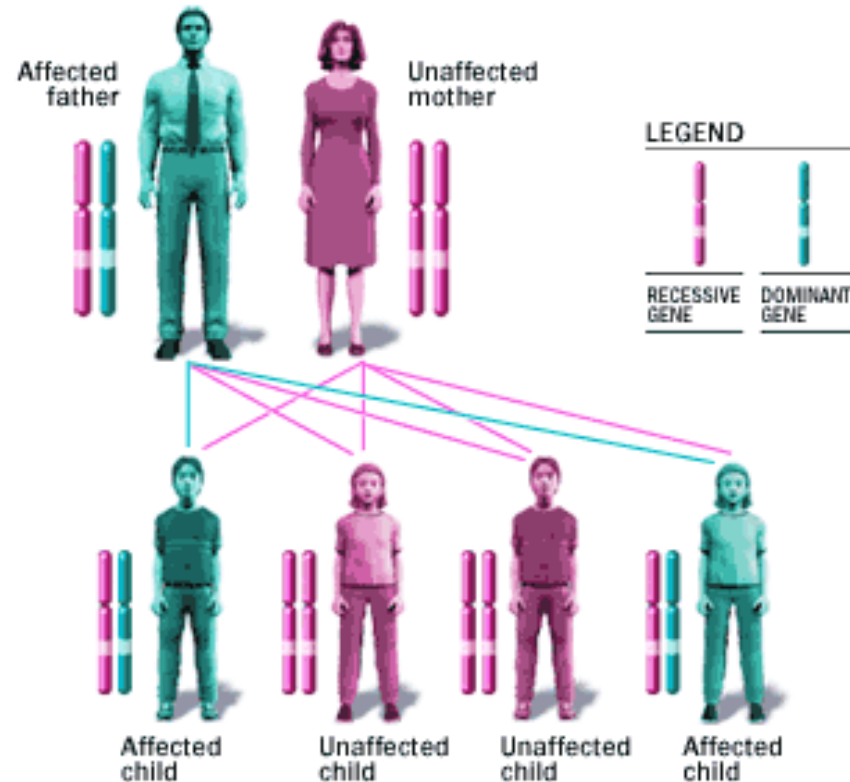


- *Department of Obstetrics and Gynecology,  
Chaim Sheba Medical Center, Ramat Gan, Israel*

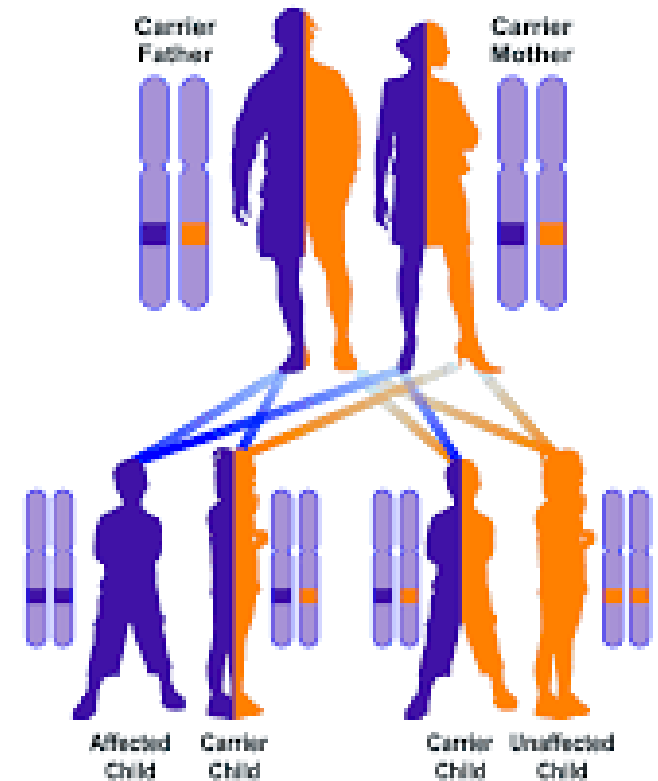
# Genetic disorders



## Autosomal dominant inheritance pattern



## Autosomal Recessive



# **Pre-implantation genetic diagnosis (PGD)**

**PGD allows patients who are carriers of single-gene disorders or carriers of structural chromosome abnormalities (balanced translocations) to select unaffected embryos for transfer, and reduce the transmission of genetic disorders to the offspring.**

# Pre-implantation genetic diagnosis

- **PGD is considered in a similar fashion to prenatal diagnosis.**
- **PGD refers to the genetic profiling of embryos prior to implantation.**
- **PGD is an adjunct to assisted reproductive technology, and requires in vitro fertilization (IVF) to obtain oocytes or embryos for evaluation.**

# IVF cycle



OPU

ET

gonadotropin administration

gonadotropin administration

administration



pre-

treatment cycle

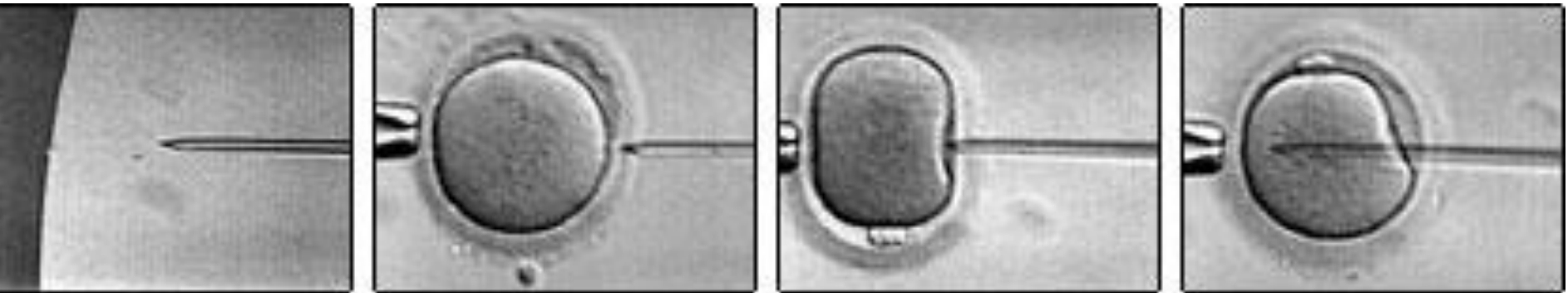


# Fertilization

IVF



ICSI

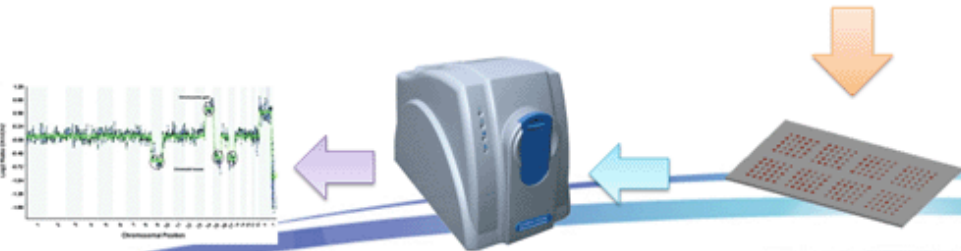
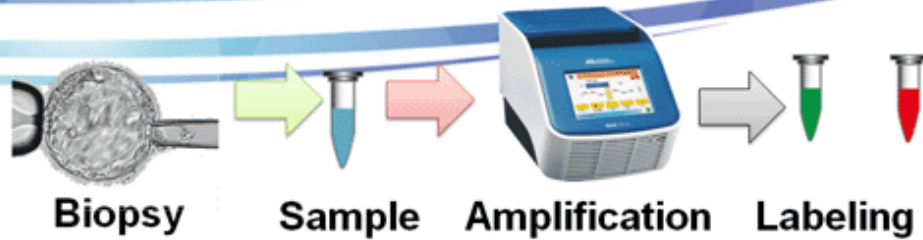


A single sperm being picked up and injected into a mature egg.

# IVE



## PGD-aCGH Workflow





# Pregnancy outcome

□56 Obstetrical outcome after preimplantation genetics

*Ginsberg N, Tur-Kaspa I, Cieslak-Janzen J, Rechitsky S, Pauling D, Horwitz A, Kuliev A, Verlinsky Y*  
*Reproductive Genetic Institute, Chicago, IL, USA*

Abstracts - 6th International Symposium on Preimplantation Genetics 2005

**Conclusions:** The obstetrical outcome after PGD is similar to that of other IVF pregnancies. PGD significantly reduced the incidence of multiple gestations.

995 children

No differences regarding mean term, prematurity (term <32 w and <37 w), mean birthweight, very low birthweight (<1500 g), perinatal death, major malformations and neonatal hospitalizations in singletons and multiples born following PGD versus ICSI were observed.

**Neonatal follow-up of 995 consecutively born children after embryo biopsy for PGD**  
**2012**

**S. Desmyttere<sup>1,\*</sup>, M. De Rycke<sup>1</sup>, C. Staessen<sup>1</sup>, I. Liebaers<sup>1</sup>,  
F. De Schrijver<sup>1</sup>, W. Verpoest<sup>2</sup>, P. Haentjens<sup>3</sup>, and Maryse Bonduelle<sup>1</sup>**

# Neonatal outcome after preimplantation genetic diagnosis

Fertil Steril® 2014

Talia Eldar-Geva, M.D., Ph.D.,<sup>a,b</sup> Naama Srebnik, M.D.,<sup>a,b</sup> Gheona Altarescu, M.D.,<sup>b,c</sup> Irit Varshaver, M.Sc.,<sup>a</sup> Baruch Brooks, Ph.D.,<sup>a</sup> Ephrat Levy-Lahad, M.D.,<sup>b,c</sup> Ruben Bromiker, M.D.,<sup>b,d</sup> and Michael S. Schimmel, M.D.<sup>b,d</sup>

**242 children born after PGD, 242 after ICSI and 733 after a spontaneous conception (SC).**

Birth weight, pregnancy duration, and intrauterine growth for singletons.

	PGD	ICSI	SC	P value <sup>a</sup>
Birth weight (g)	3,238 ± 514	3,062 ± 573	3,204 ± 504	.005
<2,500 (%)	7 (4.4)	19 (12)	27 (5.5)	.01
<1,500 (%)	1 (0.6)	2 (1.3)	7 (1.4)	.7
Pregnancy duration (wk)	38.8 ± 1.8	38.4 ± 2.1	39.2 ± 1.9	.005
<37 (%)	11 (7)	18 (11.4)	28 (5.7)	.05
<34 (%)	2 (1.3)	5 (3.2)	10 (2.0)	.5
<32 (%)	1 (0.6)	2 (1.3)	8 (1.6)	.6
<28 (%)	1 (0.6)	0	1 (0.2)	.5
Intrauterine growth				.001
IUGR (<10th percentile)	8 (5.1)	15 (9.5)	27 (5.5)	
Normal weight (10–90th percentiles)	124 (78.4)	125 (79.1)	421 (85.7)	
LGA (>90th percentile)	18 (16.5)	26 (11.4)	43 (8.8)	

# Pre-term birth and low birth weight following preimplantation genetic diagnosis: analysis of 88 010 singleton live births following PGD and IVF cycles

2017

Sesh Kamal Sunkara<sup>1,\*</sup>, Belavendra Antonisamy<sup>2</sup>, Hepsy Y. Selliah<sup>2</sup>, and Mohan S. Kamath<sup>2</sup>

**Table II** PTB and LBW outcomes following PGD versus autologous IVF.

Outcome	n (%)		OR (95% CI)	aOR <sup>†</sup> (95% CI)
	PGD n = 439	Autologous IVF n = 87 571		
PTB (<37 weeks)	28 (6.4)	7968 (9.1)	*0.68 (0.46–0.99)	*0.66 (0.45–0.98)
Early PTB (<32 weeks)	4 (0.9)	1512 (1.7)	0.52 (0.20–1.40)	0.48 (0.18–1.31)
LBW (<2500 g)	24 (5.5)	8138 (9.3)	*0.56 (0.37–0.85)	*0.58 (0.38–0.88)
Very LBW (<1500 g)	4 (0.9)	1580 (1.8)	0.50 (0.19–1.34)	0.50 (0.18–1.35)

2016

# Preimplantation genetic diagnosis: a national multicenter obstetric and

Obstetrical and neonatal outcomes following preimplantation genetic diagnosis (PGD), in vitro fertilization/intracytoplasmic sperm injection (IVF/ICSI), and spontaneous conception (SC) with or without existing monogenic disorders in the parents, Denmark 1999-2013.

Outcome	PGD <sup>a</sup> vs. IVF/ICSI <sup>b</sup>	PGD monogenic <sup>c</sup> vs. ICSI <sup>d</sup>	PGD monogenic <sup>c</sup> vs. SC <sup>e</sup>	PGD aberration <sup>f</sup> Vs. IVF <sup>g</sup>	PGD aberration <sup>f</sup> vs SC <sup>e</sup>	PGD monogenic <sup>c</sup> vs. SC monogenic <sup>h</sup>
<b>Obstetrical</b>						
Preeclampsia, odds ratio (95% CI)	1.2 (0.6; 2.4)	1.3 (0.5; 3.2)	1.2 (0.5; 3.2)	1.1 (0.4; 3.2)	1.2 (0.4; 3.3)	0.9 (0.4; 2.3)
Preterm primary rupture of membranes, odds ratio (95% CI)	1.3 (0.7; 2.4)	2.0 (0.9; 4.2)	2.1 (1.0; 4.4) <sup>j</sup>	0.6 (0.2; 2.0)	0.6 (0.2; 2.1)	1.4 (0.7; 3.1)
Placenta previa, odds ratio (95% CI)	2.1 (0.8; 5.8)	2.4 (0.6; 9.8)	9.4 (2.3; 38.7) <sup>j</sup>	1.8 (0.4; 7.4)	8.4 (2.0; 34.6) <sup>j</sup>	4.2 (1.1; 15.8) <sup>j</sup>
Abruption of placenta, odds ratio (95% CI)	1.2 (0.2; 8.3)	2.4 (0.3; 17.8)	3.6 (0.5; 26.4)	na	na	2.2 (0.2; 21.9)
Cesarean section, odds ratio (95% CI)	1.7 (1.1; 2.5) <sup>j</sup>	1.9 (1.1; 3.4) <sup>i</sup>	2.1 (1.1; 3.7) <sup>j</sup>	1.6 (0.9; 2.7)	2.0 (1.1; 3.5) <sup>j</sup>	1.4 (0.9; 2.2)
Induction of labor, odds ratio (95% CI)	0.9 (0.6; 1.5)	1.0 (0.5; 2.0)	1.2 (0.6; 2.2)	0.8 (0.4; 1.7)	1.0 (0.5; 2.0)	1.1 (0.7; 1.9)
<b>Neonatal</b>						
Gestational age (d), mean difference (95% CI)	-1.2 (-4.2; 1.8)	-5.1 (-9.1; -1.1) <sup>j</sup>	-6.3 (-9.5; -3.2) <sup>j</sup>	3.2 (-1.25; 7.7)	0.0 (-3.3; 3.3)	0.2 (-3.4; 3.8)
Preterm birth (<37 wk), odds ratio (95% CI)	1.3 (0.8; 2.1)	1.7 (0.9; 3.2)	1.8 (0.9; 3.5)	0.9 (0.4; 1.9)	1.2 (0.5; 2.8)	1.2 (0.6; 2.4)
Birth weight (g), mean difference (95% CI)	-10 (-115; 96)	-149 (-293; 5.4)	-178 (-311; -45) <sup>j</sup>	155 (1; 310) <sup>j</sup>	50 (-88; 189)	20 (-111; 151)
Low birth weight (<2500 g), odds ratio (95% CI)	1.2 (0.7; 2.0)	2.0 (1.1; 3.9) <sup>j</sup>	2.3 (1.1; 4.5) <sup>j</sup>	0.4 (0.2; 1.2)	0.5 (0.2; 1.5)	0.8 (0.4; 1.8)
Length at birth (cm), mean difference (95% CI)	-0.2 (-0.8; 0.4)	-0.9 (-1.8; -0.1)	-1.1 (-1.7; -0.4) <sup>j</sup>	0.6 (-0.2; 1.5)	0.3 (-0.4; 0.9)	0.6 (-0.2; 1.5)
Apgar score: below 7 (%), odds ratio (95% CI)	1.1 (0.3; 4.3)	1.9 (0.5; 8.0)	1.8 (0.5; 8.0)	na	na	0.5 (0.1; 3.3)
Neonatal admission, odds ratio (95% CI)	0.9 (0.5; 1.5)	1.4 (0.8; 2.6)	1.4 (0.8; 2.7)	0.4 (0.2; 1.1)	0.5 (0.2; 1.3)	1.2 (0.2; 2.4)
Length of neonatal admission (d), mean difference (95% CI)	20 (13; 27) <sup>j</sup>	30 (22; 39) <sup>j</sup>	31 (24; 38) <sup>j</sup>	-9 (-24; 5)	-7 (-19; 5)	13 (-11; 38)
Malformations, odds ratio (95% CI)	1.5 (0.9; 2.5)	1.3 (0.6; 2.8)	1.3 (0.6; 2.8)	1.7 (0.8; 3.6)	1.8 (0.8; 3.8)	1.2 (0.6; 2.2)
Stillbirth, odds ratio (95% CI)	na	na	na	na	na	na
Perinatal death, odds ratio (95% CI)	na	na	na	na	na	na
Infant death, odds ratio (95% CI)	na	na	na	na	na	na
Placenta weight (g), mean difference (95% CI)	13 (-21; 49)	-37 (-86; 12)	-17 (-54; 19)	68 (19; 119)	71 (33; 109) <sup>j</sup>	60 (25; 97) <sup>j</sup>





**700 PGD pregnancies**

**Oocyte- IVF/ICSI-Day3/4/5Bx-  
FISH/PCR/CMA**

**Oocyte/embryo quality**

**Pregnancy (maternal and neonatal)  
outcomes- .....Incomplete**



**THANK YOU**