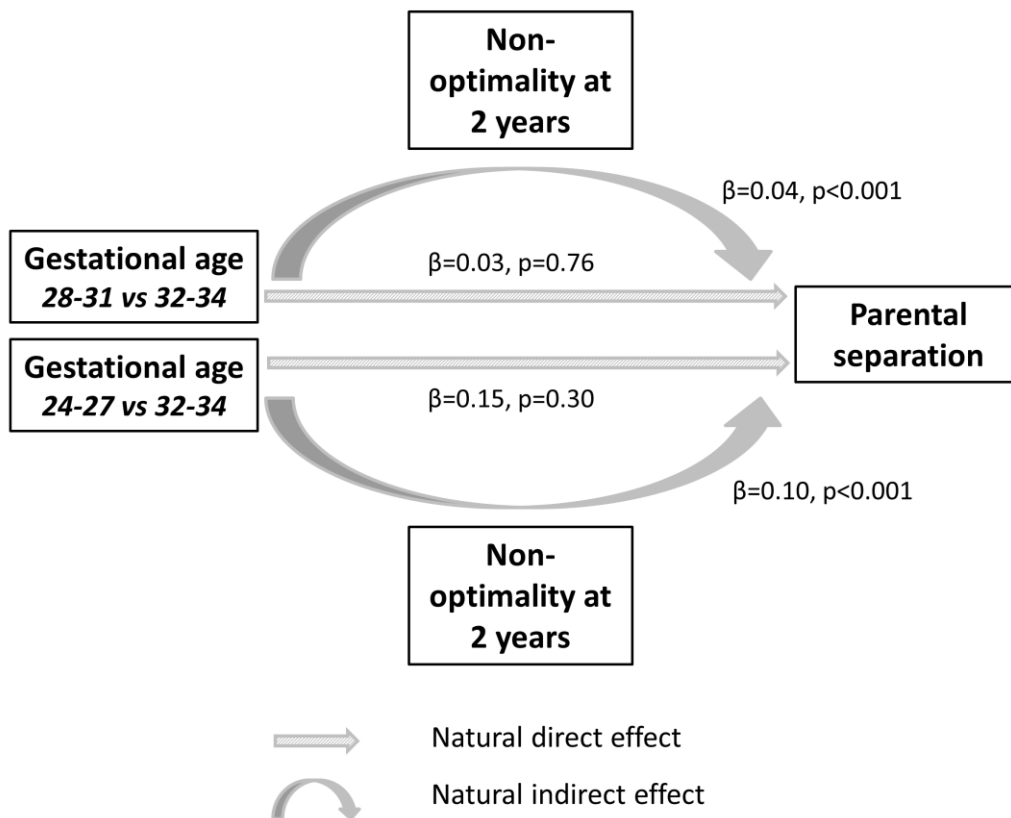


Supplementary materials

Supplementary Figure 1. Decomposition of the effect of low gestational age on the risk of parental separation into a direct effect and an indirect effect mediated by the neurodevelopment of preterm infants (non-optimality at two years). This mediation model was adjusted for gender, multiple pregnancy, Z-score of birthweight, socioeconomic level, social security benefits for those with low incomes, and urbanicity of the residential municipality.



Supplementary Table 1. Incidence risks and incidence rates of parental separation in preterm infants population according to the perinatal characteristics of the child, the socio-economic level of the family and the urbanicity of the residential municipality (n=5,732).

Variable	Category	Incidence risk of parental separation (N events/N at risk) x 100	Incidence rate of parental separation for 1000 children-year	P value**
Gestational age (weeks)	32–34	9.6	22.6	0.030
	28–31	9.9	24.1	
	24–27	13.0	31.6	
Optimality at 2 years*	Yes	9.2	21.5	< 0.001
	No	13.4	34.2	
Gender	Female	9.6	23.0	0.450
	Male	10.3	24.5	
Multiple pregnancy	No	10.2	24.3	0.460
	Yes	9.6	22.9	
Z score of birthweight	<-1	10.3	24.3	0.570
	-1 to 0	10.5	25.3	
	0 to 1	9.2	21.8	
	>1	9.6	23.5	
Socio-economic level	Intermediate	10.8	26.7	< 0.001
	High	7.5	16.4	
Social security benefits due to low income	No	7.6	17.7	< 0.001
	Yes	27.4	73.6	
Urbanicity	Rural	6.6	15.2	< 0.001
	Urban	12.0	29.1	

*Children with a non-optimal neuromotor and/or psychomotor assessment and/or sensorial disability at two years were considered as non-optimal.

**log-rank test

Supplementary Table 2. Adjusted associations between the risk of non-optimal neurodevelopment at two years and gestational age with adjustment variables for preterm infants born between 2005 and 2013 followed in the LIFT cohort (n=5,732).

Variable	Category	N (%)	Adjusted OR [95%CI]
Gestational age (weeks)	32–34	3,494 (61.0)	1
	28–31	1,730 (30.2)	2.06 [1.78, 2.39]
	24–27	508 (8.9)	4.23 [3.43, 5.20]
Gender	Female	2,640 (46.1)	1
	Male	3,092 (53.9)	1.38 [1.2, 1.59]
Multiple pregnancy	No	3,617 (63.1)	1
	Yes	2,115 (36.9)	0.94 [0.82, 1.09]
Z score of birthweight	<-1	1,378 (24.0)	1
	-1 to 0	2,044 (35.7)	0.73 [0.61, 0.87]
	0 to 1	1,787 (31.2)	0.72 [0.6, 0.87]
	>1	523 (9.1)	0.99 [0.77, 1.26]
Socio-economic level	Intermediate	4,254 (74.2)	1
	High	1,478 (25.8)	0.72 [0.61, 0.85]
Social security benefits (SSB) due to low income	No	5,031 (87.8)	1
	Yes	701 (12.2)	1.31 [1.07, 1.59]
Urbanicity	Rural	2,104 (36.7)	1
	Urban	3,628 (63.3)	0.90 [0.78, 1.04]

OR: odds ratio; CI: confidence interval.

Supplementary Table 3. Crude and adjusted associations between the risk of parental separation and neurodevelopment of preterm infants (optimality at two years) with adjustment variables and interaction term between optimality at two years and social security benefits due to low income (n=5,732).

Variable	Category	N (%)	Adjusted HR [95% CI]
Optimality at 2 years*	Yes	4,636 (80.9)	1
	No	1,096 (19.1)	1.27 [0.99, 1.63]
Gender	Female	2,640 (46.1)	1
	Male	3,092 (53.9)	1.07 [0.91, 1.26]
Multiple pregnancy	No	3,617 (63.1)	1
	Yes	2,115 (36.9)	0.96 [0.81, 1.14]
Z score of birthweight	<-1	1,378 (24.0)	1
	-1 to 0	2,044 (35.7)	1.10 [0.89, 1.36]
	0 to 1	1,787 (31.2)	0.97 [0.77, 1.21]
	>1	523 (9.1)	1.04 [0.76, 1.44]
Socio-economic level	Intermediate	4,254 (74.2)	1
	High	1,478 (25.8)	0.64 [0.52, 0.79]
Social security benefits (SSB) due to low income	No	5,031 (87.8)	1
	Yes	701 (12.2)	3.27 [2.65, 4.04]
Urbanicity	Rural	2,104 (36.7)	1
	Urban	3,628 (63.3)	1.80 [1.49, 2.19]
Optimality at 2 years * SSB	Yes * No	4,102 (71.6)	1
	No * Yes	167 (2.9)	1.52 [1.03, 2.23]

HR: hazards ratio; SSB: social security benefits; CI: confidence interval.

*Children with a non-optimal neuromotor and/or psychomotor assessment and/or sensorial disability at two years were considered as non-optimal.

Supplementary Table 4. Adjusted associations between the risk of parental separation occurring from the 24-month visit and neurodevelopment of preterm infants (optimality at two years) with adjustment variables (n=5,511). In this analysis, the separations occurring between the 6-month and the 18-month visit were excluded.

	Category	N (%)	Adjusted HR [95% CI]
Optimality at 2 years*	Yes	4,477 (81.2)	1
	No	1,034 (18.8)	1.49 [1.17, 1.91]
Gender	Female	2,549 (46.3)	1
	Male	2,962 (53.7)	0.98 [0.79, 1.21]
Multiple pregnancy	No	3,479 (63.1)	1
	Yes	2,032 (36.9)	0.90 [0.72, 1.13]
Z score of birthweight	<-1	1,326 (24.1)	1
	-1 to 0	1,952 (35.4)	1.01 [0.77, 1.32]
	0 to 1	1,729 (31.4)	0.98 [0.74, 1.30]
	>1	504 (9.1)	1.02 [0.68, 1.54]
Socio-economic level	Intermediate	4,068 (73.8)	1
	High	1,443 (26.2)	0.67 [0.52, 0.87]
Social security benefits (SSB) due to low income	No	4,903 (89.0)	1
	Yes	608 (11.0)	3.01 [2.38, 3.81]
Urbanicity	Rural	2,054 (37.3)	1
	Urban	3,457 (62.7)	1.82 [1.43, 2.32]

HR: hazards ratio; CI: confidence interval.

*Children with a non-optimal neuromotor and/or psychomotor assessment and/or sensorial disability at two years were considered as non-optimal.

Supplementary Table 5. Adjusted associations between the risk of parental separation and neurodevelopment of preterm infants (optimality at two years) with adjustment variables after imputation of missing values (13 infants with weight at birth missing and 990 infants with neurodevelopmental outcome at two years missing) using the multiple imputation method (n=6,732).

Variable	Category	Adjusted HR [95% CI]
Optimality at 2 years*	Yes	1
	No	1.45 [1.21, 1.73]
Gender	Female	1
	Male	1.06 [0.91, 1.24]
Multiple pregnancy	No	1
	Yes	0.97 [0.83, 1.14]
Z score of birthweight	<-1	1
	-1 to 0	1.06 [0.87, 1.29]
	0 to 1	0.92 [0.74, 1.13]
	>1	0.98 [0.72, 1.33]
Socio-economic level	Intermediate	1
	High	0.68 [0.56, 0.82]
Social security benefits due to low income	No	1
	Yes	3.91 [3.32, 4.60]
Urbanicity	Rural	1
	Urban	1.81 [1.51, 2.18]

HR: hazards ratio; CI: confidence interval.

*Children with a non-optimal neuromotor and/or psychomotor assessment and/or sensorial disability at two years were considered as non-optimal.

Supplementary Table 6. Crude and adjusted association between the neurodevelopment of preterm infants and the risk of parental separation. Adjustment was made on perinatal characteristics of the infants, the socio-economic level of the family, and the urbanicity of the residential municipality. Only one infant from each twins' pair was kept in the analyses (n=3,654).

	Category	N (%)	Adjusted HR [95% CI]
Optimality at 2 years*	Yes	2,919 (79.9)	1
	No	735 (20.1)	1.39 [1.10, 1.74]
Gender	Female	1,676 (45.9)	1
	Male	1,978 (54.1)	1.09 [0.89, 1.33]
Multiple pregnancy	No	2,763 (75.6)	1
	Yes	891 (24.4)	0.90 [0.71, 1.15]
Z score of birthweight	<-1	869 (23.8)	1
	-1 to 0	1,283 (35.1)	1.08 [0.84, 1.39]
	0 to 1	1,147 (31.4)	0.84 [0.64, 1.10]
	>1	355 (9.7)	0.95 [0.64, 1.40]
Socio-economic level	Intermediate	2,705 (74)	1
	High	949 (26)	0.72 [0.56, 0.92]
Social security benefits (SSB) due to low income	No	3,153 (86.3)	1
	Yes	501 (13.7)	3.48 [2.82, 4.29]
Urbanicity	Rural	1,297 (35.5)	1
	Urban	2,357 (64.5)	1.81 [1.43, 2.31]

HR: hazards ratio; CI: confidence interval

* Infants with a non-optimal neuromotor and/or psychomotor assessment and/or sensorial disability at two years were considered as non-optimal.

Supplementary Table 7. Comparison of the infants born <35 weeks between 2005 and 2011 still followed at the 60-month visit (n=3,295) and those lost to follow-up between the 24-month and the 60-month visit (n=1,518).

Variable	Category	Children still followed at the 60-month visit (n=3295)	Children lost to follow-up between the 24-month and the 60-month visit (n=1518)	P value
Parental separation	Living together	2,868 (87.0)	1,309 (86.2)	0.469
	Separated	427 (13.0)	209 (13.8)	
Gestational age (weeks)	32–34	2,025 (61.5)	949 (62.5)	0.772
	28–31	978 (29.7)	440 (29.0)	
	24–27	292 (8.9)	129 (8.5)	
Optimality at 2 years*	Yes	2,719 (82.5)	1,190 (78.4)	< 0.001
	No	576 (17.5)	328 (21.6)	
Gender	Female	1,507 (45.7)	693 (45.7)	0.982
	Male	1,788 (54.3)	825 (54.3)	
Multiple pregnancy	No	2,063 (62.6)	949 (62.5)	0.976
	Yes	1,232 (37.4)	569 (37.5)	
Z score of birthweight	<-1	816 (24.8)	357 (23.6)	0.192
	-1 to 0	1,156 (35.1)	579 (38.2)	
	0 to 1	1,030 (31.3)	444 (29.3)	
	>1	291 (8.8)	135 (8.9)	
Socio-economic level	Intermediate	2,318 (70.3)	1,191 (78.5)	< 0.001
	High	977 (29.7)	327 (21.5)	
Social security benefits due to low income (SSB)	No	2,864 (86.9)	1,292 (85.1)	0.099
	Yes	431 (13.1)	226 (14.9)	
Urbanicity	Rural	1,221 (37.1)	550 (36.2)	0.604
	Urban	2,074 (62.9)	968 (63.8)	

*Children with a non-optimal neuromotor and/or psychomotor assessment and/or sensorial disability at two years were considered as non-optimal.