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## Gestational weight gain adequacy and intrapartum oxytocin and cesarean section use: Observational population-based study in France

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#### Abstract

**Introduction:** Research on maternal prepregnancy weight suggests adiposity is associated with dysfunctional labor, but knowledge about how gestational weight gain (GWG) affects labor is sparse. Our objective was to evaluate associations between GWG adequacy and intrapartum obstetric interventions (oxytocin administration; cesarean section) necessitated by labor dysfunction.

**Material and methods:** Using national, population-based French National Perinatal Survey 2016 data, we included term cephalic singleton pregnancies involving trial of labor (n = 9724). For the intrapartum oxytocin administration analysis, we included only women with spontaneous labor (n = 7352). GWG was calculated as the difference between end of pregnancy and prepregnancy weight (both self-reported) and categorized as insufficient, adequate (reference group), or excessive by prepregnancy body mass index (BMI; underweight <18.5, normal weight 18.5–24.9, overweight 25–29.9, obese  $\geq 30 \text{ kg/m}^2$ ) using the 2009 Institute of Medicine thresholds. Multilevel generalized estimating equation logistic regression models, unadjusted and adjusted for a priori confounders, evaluated intervention-GWG adequacy associations within BMI categories (under/normal weight combined), stratified by parity (primiparas; multiparas).

**Results:** GWG adequacy was associated with oxytocin use among under/normal weight women (primiparas: insufficient 57.3%, adequate 60.8%, excessive 65.0%, p = 0.014; multiparas: insufficient 27.2%, adequate 29.1%, excessive 36.2%, p < 0.001) and overweight primiparas (insufficient 56.0%, adequate 58.7%, excessive 72.5%, p = 0.002). In unadjusted and adjusted models, trends of increased odds of oxytocin administration among women with excessive GWG were found regardless of parity and prepregnancy BMI. Similarly, among under/normal weight women, GWG adequacy was associated with intrapartum cesarean section (primiparas: insufficient 10.7%, adequate 12.7%, excessive 15.3%, p = 0.014; multiparas: insufficient 3.1%,

Abbreviations: aOR, adjusted odds ratio; BMI, body mass index; CI, confidence interval; CS, cesarean section; GA, gestational age; GWG, gestational weight gain; IOM, Institute of Medicine; NPS, French National Perinatal Survey; OR, odds ratio.

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adequate 3.5%, excessive 6.3%, p < 0.001) with increased cesarean section among multiparas with excessive GWG persisting in adjusted models (adjusted odds ratio 1.9, 95% confidence interval 1.3–2.7). However, intrapartum cesarean section was reduced among multiparas with overweight and obese prepregnancy BMI and excessive GWG.

**Conclusions:** Excessive GWG was associated with intrapartum oxytocin administration, regardless of parity or prepregnancy BMI, and cesarean section among women with under/normal weight prepregnancy BMI, providing evidence for benefits of healthy GWG for normal labor progression. Additional research is needed to verify our findings and understand differences by BMI.

#### KEYWORDS

body mass index, gestational weight gain, Institute of Medicine guidelines, obstetric interventions, pregnancy, weight gain

#### 1 | INTRODUCTION

Worldwide, the obstetric population is increasing in body mass index (BMI)<sup>1</sup> and gestational weight gain (GWG),<sup>2,3</sup> posing numerous risks to mothers (for example, gestational diabetes and pre-eclampsia) and fetuses (for example, macrosomia and preterm delivery).<sup>4</sup> Further, increased maternal prepregnancy weight/adiposity/BMI may be associated with intrapartum obstetric interventions related to dysfunctional labor (for example, cesarean section [CS] or oxy-tocin administration) through increased adiposity causing uterine contraction impairment resulting in prolonged labor,<sup>5</sup> a blunting of the effects of oxytocin,<sup>6</sup> or hormonal factors (excess adipose tissue producing elevated levels of estrogen).<sup>6,7</sup>

Although these mechanisms could reasonably be extended to excess GWG as well, few studies have explored the association between GWG and intrapartum interventions due to dysfunctional labor. In particular, no studies have investigated the association between GWG and intrapartum oxytocin administration. Previous studies have found evidence that excessive/high GWG may be associated with dystocia,<sup>8</sup> shoulder dystocia,<sup>9</sup> CS due to dystocia,<sup>10</sup> or dysfunctional labor,<sup>11</sup> and emergency/intrapartum/non-elective CS,<sup>9,12-18</sup> but the applicability of these studies to contemporary French or European obstetric populations is limited. Specifically, many of these studies were conducted in Asian<sup>8,10,11</sup> or other non-European<sup>9,13-15,19</sup> populations. Although other studies of the association between GWG and intrapartum CS were conducted in European populations, important differences, such as GWG adequacy not based on the 2009 Institute of Medicine (IOM) recommendations,<sup>16</sup> older cohorts before (fully or partially) the IOM recommendations,<sup>16-18</sup> or non-general obstetric populations used (two previous CS attempting trial of labor,<sup>15</sup> obese prepregnancy BMI<sup>19</sup>) and a different research focus (risks of gestational weight loss),<sup>12</sup> limits their direct applicability to current French or European obstetric populations.

Both BMI and GWG are modifiable risk factors for intrapartum obstetric interventions, with improving GWG a particularly relevant,

#### Key message

Gestational weight gain is potentially associated with labor dysfunction. Excessive gestational weight gain was associated with intrapartum oxytocin use (overall) and cesarean section (among women with lower body mass index). Healthy gestational weight gain should be encouraged as it might reduce intrapartum interventions.

but understudied, target for prenatal intervention. Therefore, the objective of this study was to evaluate the association between GWG adequacy and intrapartum obstetric interventions necessitated by labor dysfunction in women with term singleton pregnancies undergoing trial of labor within a large, prospective population-based French cohort.

#### 2 | MATERIAL AND METHODS

#### 2.1 | Study population

We used data from the 2016 French National Perinatal Survey (NPS),<sup>2</sup> a routine, nationally representative study including all live and stillbirths in all maternity units in metropolitan, mainland France (n = 493 hospitals) during 1 week. Trained study personnel, generally midwives, performed data collection. Face-to-face interviews with women 2–3 days after birth were conducted using a standardized questionnaire to collect sociodemographic characteristics and antenatal care information and chart abstraction was performed to collect information related to maternal and neonatal health and delivery.

A total of 13147 women were included in the 2016 NPS (Figure 1). We excluded women with non-singleton births (n = 249),

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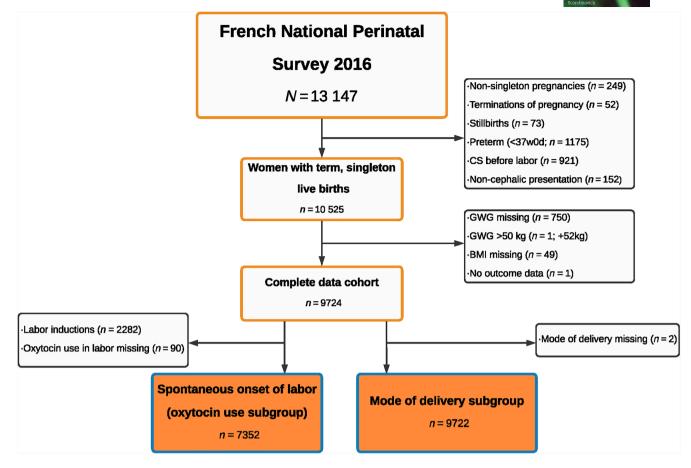


FIGURE 1 Participant flowchart. BMI, body mass index; CS, cesarean section; GWG, gestational weight gain.

pregnancy terminations (n = 52), and stillbirths (n = 73), as these women may have distinct GWG patterns. Additionally, we excluded preterm births ( $<37^{+0}$  weeks' gestational age [GA] or missing GA at birth; n = 1175) and pregnancies with prelabor CS (or missing onset of labor; n = 921) or non-cephalic presentation (n = 152), as our research interest was intrapartum obstetric interventions in term pregnancies. We also excluded women with missing (n = 750) or implausible (n = 1; gain >50 kg or loss >30 kg) GWG, missing prepregnancy BMI (n = 49), or missing outcome data (n = 1). As a result, 9724 women were included in the present study.

#### 2.2 | Study variables

Observed GWG was calculated as a woman's end of pregnancy weight minus her prepregnancy weight (both self-reported). The main exposure of interest was GWG adequacy (insufficient, adequate, excessive) based on the 2009 IOM guidelines, which provide specific guidelines based on maternal prepregnancy BMI (underweight, normal weight, overweight, obese).<sup>3</sup> We used a previously described method to account for the inherent correlation between GA at birth and GWG, as detailed in Box 1.<sup>20</sup> Briefly, the IOM recommendations for total GWG at 40 weeks were converted to ranges

of proportions for each BMI category by determining expected GWG at 40 weeks (based on the assumptions underlying the IOM recommendations) and then dividing IOM recommended ranges by the IOM expected GWG. Women's individual proportions of recommended GWG achieved were calculated as observed GWG/ expected GWG, and GWG adequacy for each woman was classified as insufficient (below lower bound), adequate (within recommended range), or excessive (above upper bound of the converted IOM ranges of proportions) GWG.

In line with the IOM recommendations, which provide distinct thresholds depending on prepregnancy BMI, GWG adequacy is reported and analyzed by BMI category. Maternal prepregnancy BMI (kg/m<sup>2</sup>) was determined by self-reported height and self-reported prepregnancy weight and classified as: underweight less than 18.5, normal weight 18.5–24.9, overweight 25–29.9, or obese 30 kg/m<sup>2</sup> or more. For analysis, women with underweight and normal weight BMI were combined because of the small number of underweight women.

French guidelines (2007 French National Nutrition and Health Program [*Programme National Nutrition et Santé*]) recommend that women with normal prepregnancy BMI gain 12kg, and higher GWG for women with underweight prepregnancy BMI but lower GWG (not under 7 kg) for women with overweight or obese prepregnancy

# BOX 1 Determination of gestational weight gain adequacy based on the 2009 Institute of Medicine guidelines and assumptions using a method described previously to account for gestational age at birth<sup>20</sup>

Prepregnancy BM	1	Recommendations	for GWG <sup>3</sup>			o recommended on proportions
Category	BMI (kg/m <sup>2</sup> )	Total first- trimester (<13 weeks; kg)	Rate of second- and third-trimester GWG (kg/week)	Total at 40 weeks (kg)	Expected GWG (40 weeks)	Range of proportion of GWG
Underweight	<18.5	2.0	0.51	12.5-18.0	15.77	0.79-1.14
Normal weight	18.5-24.9	2.0	0.42	11.5-16.0	13.34	0.86-1.20
Overweight	25-29.9	1.0	0.28	7.0-11.5	8.56	0.81-1.34
Obese	≥30	0.5	0.22	5.0-9.0	6.44	0.78-1.41

Abbreviations: BMI, body mass index; GWG, gestational weight gain; IOM, Institute of Medicine.

1. Conversion of IOM recommendations to ranges of proportions for each BMI category

a. Calculation of Expected gestational weight gain (GWG) at 40 weeks based on IOM recommendations
 Expected GWG = Total first-trimester GWG (kg)+([gestational age [GA] at birth - 13 weeks] \* Rate of second- and third-trimester GWG).

Example for overweight BMI: Expected GWG at 40 weeks =  $(1 + [40 - 13]^* 0.28) = 8.56$  kg.

b. Ranges of proportions of GWG calculated from recommended ranges of total GWG at 40 weeks: lower and upper bounds of recommended range divided by expected GWG at 40 weeks

Example for underweight BMI: (12.5/15.77; 18/15.77) = 0.79-1.14.

2. Women's GWG compared with ranges of proportions

a. Observed GWG (kg) calculated as: self-reported end of pregnancy minus prepregnancy weight

b. Individual expected GWG based on gestational age at delivery (kg) calculated using the formula from step 1a

Example for a woman with normal weight BMI and GA at birth 35 weeks:

Individual Expected GWG = (2 + [35-13]\*0.42) = 11.24 kg.

c. Women's individual proportion of recommended GWG achieved calculated as: observed GWG/expected GWG

Example for woman with normal weight BMI and GA at birth 35 weeks with GWG 12.5 kg (continued from 2b):

Individual proportion of recommended GWG achieved = 12.5 kg/11.24 kg = 1.11.

- d. GWG adequacy determined by comparing women's individual proportion of recommended GWG achieved with the recommended ranges of proportions for her BMI category
- Insufficient GWG: less than lower limit
- Adequate GWG: within recommended range of proportions
- Excessive GWG: more than upper limit

Example for woman with normal weight BMI and GA at birth 35 weeks with GWG 12.5 kg (continued from 2b and 2c): given that the individual proportion of recommended GWG achieved was 1.11 and range of proportion of GWG was 0.86–1.20, GWG was considered adequate.

BMI.<sup>21</sup> Though generally similar to the IOM guidelines, the French guidelines lack clear ranges of appropriate GWG to define GWG adequacy and furthermore the IOM guidelines are routinely used in clinical practice and research internationally and are better known to French clinicians.<sup>15,17,22,23</sup> Hence, we used the IOM guidelines in our analysis.

Our main outcomes, collected by chart abstraction, are oxytocin administration during labor (yes, no) and CS. We excluded women with prelabor CS from all analyses (Figure 1) and for the analysis of oxytocin administration, only included women with spontaneous onset of labor (n = 7352), excluding induction of labor.

Maternal characteristics, collected by interview, reported are: parity (primiparous, multiparous), age (<25, 25–34, ≥35 years), country/region of birth (France, not France [includes Europe, northern Africa, sub-Saharan Africa, other]), education (less than high school or high school completed; any post-high school graduation education), and for multiparas, previous mode of delivery (vaginal delivery only, vaginal delivery and CS, CS only). Pregnancy complications and delivery characteristics, obtained from chart abstraction, included gestational diabetes (yes, no) and hypertension during pregnancy (blood pressure: systolic >140 mm Hg or diastolic >90 mm Hg; yes [with or without proteinuria], no), birthweight (grams; ≤2499, 2500–2999, 3000–3499, 3500–3999,  $\geq$ 4000), and GA at birth (continuously [days] and categorically: early term [37–38 weeks], full term [39–40 weeks], late term [41+weeks]).

#### 2.3 | Statistical analyses

Overall maternal prepregnancy BMI, GWG, and intrapartum intervention use were determined. Maternal characteristics, pregnancy complications, and delivery characteristics were compared by GWG adequacy, for all eligible women and within the spontaneous onset of labor (oxytocin analysis) subgroup. Percentages of women receiving oxytocin or CS within maternal prepregnancy BMI categories were compared by GWG adequacy. For these descriptive/crude analyses, categorical variables were assessed using the chi-squared test and continuous variables using analysis of variance (normally distributed) or the Kruskal-Wallis test (non-normally distributed).

Then, multilevel generalized estimating equation logistic regression models, accounting for hospital clustering, were used to calculate unadjusted and adjusted odds ratios (OR and aOR) and 95% confidence intervals (CIs) for the association between GWG adequacy and oxytocin administration and CS. Confounders chosen a priori based on literature review and constructed directed acyclic graphs (not shown) included in adjusted models were: maternal country/region of birth, age, education, prepregnancy BMI (to account for potential residual confounding within BMI categories), GA at birth (days), and birthweight (g; continuous variables). Adjusted ORs and 95% CIs for these covariates are also reported for insight into potential relationships underlying associations.

To verify that combining women with underweight and normal weight prepregnancy BMI did not impact our results, we performed sensitivity analyses to compare the GWG and BMI of these two groups and to repeat the analyses of the association between GWG adequacy and oxytocin administration and CS with women with underweight prepregnancy BMI excluded.

Due to differences in GWG and prepregnancy BMI by parity and potential differences in labor progression and management,<sup>24,25</sup> all analyses were stratified by parity. SAS software version 9.4 for Windows (SAS Institute Inc.) was used for statistical analyses.

#### 2.4 | Ethics statement

The 2016 NPS was performed in accordance with the Declaration of Helsinki and with all relevant national and institutional guidelines and regulations. The 2016 NPS was approved by the National Council on Statistical Information (Comité du Label; approval number 2016X703SA), the French Commission on Information Technology and Liberties ([CNIL]; registration number 915197; CNIL 2016-004, approved January 14, 2016), and the Inserm ethics committee (approval IRB00003888 no. 14-191, approved December 9, 2014). Participants provided verbal consent for survey participation before being interviewed, as approved by the Inserm ethics committee. 305

#### 3 | RESULTS

#### 3.1 | Population characteristics

Overall, 8.4% of primiparas were underweight, 64.7% were normal weight, 17.8% were overweight, and 9.1% were obese. For multiparas, 6.8% were underweight, 59.3% were normal weight, 21.2% were overweight, and 12.7% were obese by their prepregnancy BMI. Among primiparas, 25.9% had insufficient, 36.4% had adequate, and 37.7% had excessive GWG, while among multiparas, 29.7% had insufficient, 37.2% had adequate, and 33.1% had excessive GWG.

Among both primiparas (Table 1) and multiparas (Table 2), differences by GWG adequacy were noted for most maternal, pregnancy, and delivery characteristics (p < 0.05). Results were generally similar within the oxytocin subgroups of primiparas (Table S1) and multiparas (Table S2).

## 3.2 | Association between intrapartum oxytocin administration and GWG adequacy

Overall, 62.6% of primiparas and 32.8% of multiparas received intrapartum oxytocin. Among primiparas and multiparas with underweight and normal weight prepregnancy BMI, intrapartum oxytocin administration increased across insufficient, adequate, and excessive GWG (primiparas: p = 0.014; multiparas: p < 0.001; Figure 2), a trend also evident among primiparas with overweight prepregnancy BMI (p = 0.002). In unadjusted and adjusted regression models (Table 3), similar results were noted among women with underweight and normal weight prepregnancy BMI. Additionally, regardless of parity or prepregnancy BMI, a trend of increased oxytocin administration among women with excessive GWG was apparent. For primiparas with obese prepregnancy BMI, oxytocin administration was also increased among those with insufficient GWG (aOR 2.6, 95% CI 1.1–6.3). Some associations were noted between oxytocin administration and other covariates of interest (Table S3).

# 3.3 | Association between intrapartum CS and GWG adequacy

Overall, 15.8% of primiparas and 5.7% of multiparas received an intrapartum CS. Among primiparas and multiparas with underweight and normal weight prepregnancy BMI, intrapartum CS increased across insufficient, adequate, and excessive GWG (primiparas: p = 0.014; multiparas: p < 0.001; Figure 3). Similar trends were found for primiparas with overweight (p = 0.527) and obese (p = 0.872) prepregnancy BMI. However, intrapartum CS was reduced among multiparas with overweight and obese prepregnancy BMI and excessive GWG. In unadjusted and adjusted regression models (Table 3), similar results were noted, though statistically significant associations for the association between GWG adequacy and intrapartum CS in adjusted models were only noted for multiparas with underweight and normal weight 
 TABLE 1
 Maternal and pregnancy and delivery characteristics by GWG adequacy among primiparous women.

	Insufficien n = 1094	t GWG <sup>a</sup>	Adequate n = 1542	GWG <sup>a</sup>	Excessive n = 1595	GWG <sup>a</sup>	р
	n	(%)	n	(%)	n	(%)	value <sup>b</sup>
Maternal characteristics							
BMI <sup>c</sup>							
Underweight + normal weight	884	80.8	1287	83.5	920	57.7	<0.001
Overweight	113	10.3	179	11.6	462	29.0	
Obese	97	8.9	76	4.9	213	13.4	
Age							
<25 years	236	21.6	298	19.3	380	23.8	0.011
25-34 years	719	65.7	1059	68.7	1055	66.1	
≥35 years	139	12.7	185	12.0	160	10.0	
Country or region of birth							
France	904	82.7	1307	84.8	1372	86.0	0.064
Not France	189	17.3	235	15.2	223	14.0	
Education							
Less than high school/high school	391	36.1	526	34.4	719	45.4	<0.001
Any post-high school graduation	693	63.9	1003	65.6	864	54.6	
Pregnancy and delivery characteristic	:S						
Hypertension during pregnancy <sup>d</sup>							
No	1062	97.3	1480	96.5	1464	92.2	< 0.001
Yes (with or without proteinuria)	29	2.7	54	3.5	123	7.8	
Gestational diabetes							
No	966	88.4	1407	91.5	1470	92.6	< 0.001
Yes	127	11.6	131	8.5	117	7.4	
GWG (kg), median (IQR)	9	6-10	13	12-15	18	16-20	< 0.001
Birthweight (g), mean (SD)	3151.7	400.7	3273.7	420.7	3376.0	439.8	< 0.001
≤2499	44	4.0	30	1.9	36	2.3	< 0.001
2500-2999	338	30.9	363	23.5	273	17.1	
3000-3499	501	45.8	706	45.8	666	41.8	
3500-3999	187	17.1	364	23.6	492	30.9	
≥4000	23	2.1	79	5.1	127	8.0	
Gestational age at birth (days), median (IQR)	279	274-285	280	274-286	281	274-287	<0.001
Early term (37–38 weeks)	216	19.7	335	21.7	304	19.1	<0.001
Full term (39-40 weeks)	675	61.7	863	56.0	873	54.7	
Late term (41+ weeks)	203	18.6	344	22.3	418	26.2	

Abbreviations: BMI, body mass index; GWG, gestational weight gain; IQR, interquartile range; SD, standard deviation.

<sup>a</sup>Based on 2009 Institute of Medicine thresholds,<sup>3</sup> accounting for gestational age at birth;<sup>20</sup> considered insufficient GWG if below recommendation, adequate if within recommendation, or excessive if above recommendation.

<sup>b</sup>Categorical variables: chi-squared test; normally distributed continuous variables: analysis of variance; non-normally distributed continuous variables: Kruskal–Wallis test.

<sup>c</sup>BMI: underweight: <18.5; normal weight: 18.5–24.9; overweight: 25–29.9; obese: ≥30 kg/m<sup>2</sup>.

 $^{d}$ Hypertension defined as blood pressure: systolic >140 mm Hg or diastolic >90 mm Hg.

TABLE 2 Maternal and pregnancy and delivery characteristics by GWG adequacy among multiparous women.

	Insufficie n = 1632	nt GWG <sup>ª</sup>	Adequate o n = 2044	GWGª	Excessive n = 1817	GWGª	р
	n	(%)	n	(%)	n	(%)	value <sup>b</sup>
Maternal characteristics							
BMI <sup>c</sup>							
Underweight + normal weight	1215	74.4	1486	72.7	931	51.2	<0.001
Overweight	204	12.5	393	19.2	566	31.2	
Obese	213	13.1	165	8.1	320	17.6	
Age							
<25 years	144	8.8	122	6.0	118	6.5	0.011
25–34 years	1066	65.3	1356	66.3	1216	66.9	
≥35 years	422	25.9	566	27.7	483	26.6	
Country or region of birth							
France	1283	78.7	1682	82.3	1473	81.1	0.020
Not France	348	21.3	362	17.7	344	18.9	
Education							
Less than high school/high school	787	48.7	835	41.2	943	52.4	<0.001
Any post-high school graduation	830	51.3	1190	58.8	857	47.6	
Previous delivery mode(s)							
VD only	1436	88.0	1792	87.7	1577	86.8	0.219
VD and CS	72	4.4	69	3.4	80	4.4	
CS only	124	7.6	183	9.0	159	8.8	
Pregnancy and delivery characteristics							
Hypertension during pregnancy <sup>d</sup>							
No	1600	98.4	1994	98.2	1742	96.3	<0.001
Yes (with or without proteinuria)	26	1.6	37	1.8	67	3.7	
Gestational diabetes							
No	1408	86.5	1844	90.7	1635	90.2	<0.001
Yes	220	13.5	188	9.3	178	9.8	
GWG (kg), median (IQR)	8	5-10	13	11-14	17	15-20	<0.001
Birthweight (g), mean (SD)	3269.8	425.8	3404.8	425.1	3504.6	453.1	<0.001
≤2499	54	3.3	33	1.6	34	1.9	<0.001
2500-2999	382	23.4	291	14.2	185	10.2	
3000-3499	728	44.6	895	43.8	668	36.8	
3500-3999	388	23.8	661	32.3	679	37.4	
≥4000	80	4.9	164	8.0	251	13.8	
Gestational age at birth (days), median (IQR)	278.5	273-284	280	273-285	279	273-285	<0.001
Early term (37-38 weeks)	393	24.1	461	22.6	400	22.0	0.008
Full term (39-40 weeks)	994	60.9	1198	58.6	1063	58.5	
Late term (41+ weeks)	245	15.0	385	18.8	354	19.5	

Abbreviations: BMI, body mass index; CS, cesarean section; GWG, gestational weight gain; IQR, interquartile range; SD, standard deviation; VD, vaginal delivery.

<sup>a</sup>Based on 2009 Institute of Medicine thresholds,<sup>3</sup> accounting for gestational age at birth;<sup>20</sup> considered insufficient GWG if below recommendation, adequate if within recommendation, or excessive if above recommendation.

<sup>b</sup>Categorical variables: chi-squared test; continuous variables: analysis of variance (ANOVA); normally distributed continuous variables: ANOVA; nonnormally distributed continuous variables: Kruskal-Wallis test.

<sup>c</sup>BMI: underweight: <18.5; normal weight: 18.5–24.9; overweight: 25–29.9; obese: ≥30 kg/m<sup>2</sup>.

<sup>d</sup>Hypertension defined as blood pressure: systolic >140 mm Hg or diastolic >90 mm Hg.

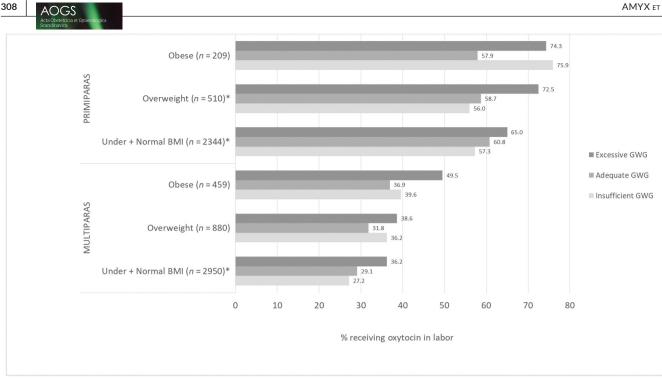


FIGURE 2 Oxytocin use among women with spontaneous labor according to gestational weight gain (GWG) adequacy<sup>a</sup> by maternal prepregnancy body mass index (BMI)<sup>b</sup>. \*Statistically significant difference (p < 0.05). <sup>a</sup>Based on 2009 Institute of Medicine thresholds,<sup>3</sup> accounting for gestational age at birth;<sup>20</sup> considered insufficient GWG if below recommendation, adequate if within recommendation, or excessive if above recommendation. <sup>b</sup>BMI: underweight: <18.5; normal weight: 18.5-24.9; overweight: 25-29.9; obese: ≥30 kg/m<sup>2</sup>.

prepregnancy BMI and excessive GWG (aOR 1.9, 95% CI 1.3-2.7) and multiparas with overweight prepregnancy BMI and excessive GWG (aOR 0.6, 95% CI 0.4-0.9). Some associations were noted between intrapartum CS and other covariates of interest (Table S3).

#### 3.4 Sensitivity analysis

GWG was similar between women with underweight and normal weight prepregnancy BMI (primiparas: underweight: median 13kg, interguartile range (IQR) 10-17 kg; normal weight: median 14 kg, IQR 11-17 kg; multiparas: underweight: median 13 kg, IQR 10-17 kg; normal weight: median 13 kg, IQR 10-16 kg) and the underweight group was fairly close in BMI (primiparas: median: 17.7 kg/m<sup>2</sup>, IQR 17.2- $18.2 \text{ kg/m}^2$ ; multiparas: median: 17.9 kg/m<sup>2</sup>, IQR 17.3–18.3 kg/m<sup>2</sup>) to the normal weight group (lower cut-point 18.5 kg/m<sup>2</sup>). Results were similar when underweight women were excluded from the analyses of the association between GWG adequacy and intrapartum interventions (results not shown).

#### 4 DISCUSSION

Among women with term singleton live births, GWG adequacy was associated with intrapartum oxytocin administration and CS. Specifically, excessive GWG was associated with intrapartum oxytocin administration, regardless of parity or prepregnancy BMI, suggesting an association between excessive GWG and labor dysfunction. In contrast, excessive GWG was significantly associated with intrapartum CS only among multiparas with underweight and normal weight prepregnancy BMI. Similar trends were found for primiparas with underweight and normal weight prepregnancy BMI, but CS was reduced among multiparas with overweight and obese prepregnancy BMI and excessive GWG.

We found that oxytocin administration was increased with excessive GWG regardless of prepregnancy BMI or parity, in line with previous studies, which found higher oxytocin administration with increased BMI,<sup>24-26</sup> including a previous NPS study.<sup>27</sup> Though we did not identify any other studies specifically investigating the association between GWG adequacy and intrapartum oxytocin administration, the association is plausible because increased BMI/adiposity may cause uterine contraction impairment,<sup>5</sup> which could lead to increased use of oxytocin in labor. Given the general lack of evidence, additional research is needed to clarify the relation between GWG adequacy and intrapartum oxytocin administration necessitated by labor dysfunction.

We found that GWG adequacy was associated with intrapartum CS among underweight and normal weight prepregnancy BMI regardless of parity. Specifically, in line with previous literature, insufficient GWG was associated with decreased  $\text{CS}^{9,11,15,16}$  and excessive GWG with increased CS.<sup>9,11,14-19</sup> However, in contrast, a trend of decreased CS with excessive GWG was found for multiparas with overweight or obese prepregnancy BMI. Although one previous study also found reduced CS due to dystocia among women

	Primiparas	aras							Multiparas	aras						
	Oxyto	Oxytocin administration <sup>a</sup>	ation <sup>a</sup>		Intrap	ntrapartum CS			Oxyto	Oxytocin administration <sup>a</sup>	ation <sup>a</sup>		Intrap	Intrapartum CS		
	Unadjusted <sup>b</sup>	isted <sup>b</sup>	Adjusted <sup>c</sup>	ed <sup>c</sup>	Unadji	Unadjusted <sup>b</sup>	Adjusted <sup>c</sup>	edc	Unadjusted <sup>b</sup>	ısted <sup>b</sup>	Adjusted <sup>c</sup>	ed <sup>c</sup>	Unadju	Unadjusted <sup>b</sup>	Adjusted <sup>c</sup>	sd <sup>c</sup>
	OR	95% CI	aOR	95% CI	OR	95% CI	aOR	95% CI	OR	95% CI	aOR	95% CI	OR	95% CI	aOR	95% CI
Underweight+ Normal weight <sup>d</sup>	I weight <sup>d</sup>															
Insufficient GWG <sup>e</sup>	0.9*	0.7-1.1	0.9	0.8-1.2	0.8*	0.6 -1.1	0.9	0.6-1.1	0.9*	0.7-1.1	0.9*	0.8 -1.1	0.9*	0.6-1.4	•6.0	0.6 -1.3
Excessive GWG <sup>e</sup>	1.2*	1.0-1.5	1.2	0.9-1.5	1.3*	1.0 -1.6	1.2	0.9-1.5	1.4*	1.2 -1.7	1.3*	1.1-1.6	1.8*	1.3-2.6	1.9*	1.3-2.7
Overweight <sup>d</sup>																
Insufficient GWG <sup>e</sup>	0.9*	0.5-1.5	$1.1^{*}$	0.6 -1.9	0.8	0.4-1.6	0.8	0.4-1.7	1.2	0.8-1.9	1.2	0.8-1.9	0.9	0.5-1.6	0.8	0.4-1.5
Excessive GWG <sup>e</sup>	1.8*	1.2-2.8	1.8*	1.1 -2.8	1.1	0.7-1.7	1.0	0.6-1.6	1.4	1.0-1.8	1.2	0.9-1.7	9.0	0.4-1.0	0.6	0.4-0.9
Obese <sup>d</sup>																
Insufficient GWG <sup>e</sup>	2.4	1.0-5.8	2.6	1.1-6.3	0.9	0.5-1.8	0.9	0.5-1.8	1.1	0.7-1.9	1.0	0.6-1.8	0.9	0.5-1.6	0.7	0.4-1.4
Excessive GWG <sup>e</sup>	2.1	1.0-4.5	2.5	1.1-5.5	1.0	0.6-1.8	1.1	0.6-2.0	1.7	1.0-2.8	1.7	1.0-2.8	0.8	0.5-1.4	0.8	0.5-1.4
Note: OR/aOR (95% CI) in bold type had a statistically significant association (does not cross null)	in bold ty	/pe had a sta	tistically :	significant ass	ociation (	does not cross	s null).	-		:						
Abbreviations: aUK, adjusted odds fatio; Li, confidence intervai; LS, cesarean section; uwu, gestational weight gain; UK, odds fatio. a Ammentiumer with compression and effektions.	justed od	ds ratio; CI, C	contidency	e Interval; CS,	cesarean	section; שעו	ى, gestatic	onal weight g	ain; UK, c	odds ratio.						
Among women wurispontaneous onset of labor only. <sup>b</sup> Associations estimated usine multi-level generalized estimating equations models accounting for clustering by hospital.	lusing m	us oriset or is ulti-level gen	eralized e	stimating equ	ations mo	idels accounti	ne for clus	stering by ho	snital							
	,	)	-		:	-	-		-			1				
Associations estimated using multi-level generalized estimating equations models accounting for clustering by hospital and adjusted for maternal age, country/region of birth, education, birthweight (g),	d using m	ulti-level gen	eralized e	stimating equati	lations mc	odels accounti	ing tor clu	stering by ho	spital anc	adjusted tor	maternal	age, country/i	region of	birth, educat	ion, birth	veight (g),

gestational age at birth (days), prepregnancy body mass index ( $kg/m^2$ ). Š

<sup>d</sup>Body mass index: underweight: <18.5; normal weight: 18.5–24.9; overweight: 25–29.9; obese: ≥30kg/m<sup>2</sup>.

<sup>e</sup>Based on 2009 Institute of Medicine thresholds,<sup>3</sup> accounting for gestational age at birth;<sup>20</sup> considered insufficient GWG if below recommendation, adequate if within recommendation (reference group), or excessive if above recommendation.

\*GWG adequacy variable statistically significant (p < 0.05) Type 3 statistic in model.

AOGS

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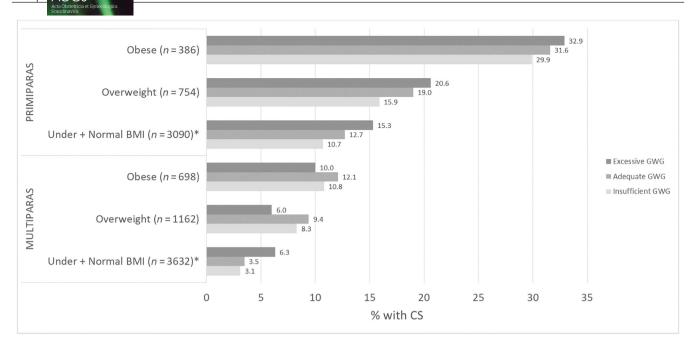


FIGURE 3 Cesarean section (CS) among women undergoing trial of labor according to gestational weight gain (GWG) adequacy<sup>a</sup> by maternal prepregnancy body mass index (BMI)<sup>b</sup>. \*Statistically significant difference (p < 0.05). <sup>a</sup>Based on 2009 Institute of Medicine thresholds,<sup>3</sup> accounting for gestational age at birth;<sup>20</sup> considered insufficient GWG if below recommendation, adequate if within recommendation, or excessive if above recommendation. <sup>b</sup>BMI: underweight: <18.5; normal weight: 18.5–24.9; overweight: 25–29.9; obese:  $\geq$ 30kg/m<sup>2</sup>.

with obese prepregnancy BMI and excessive GWG,<sup>10</sup> most found associations between excessive/increased GWG and CS regardless of BMI,<sup>9,13,15-17</sup> albeit some did not report results by BMI category.<sup>11,14,18</sup> Differences in findings between studies by BMI could be a result of different classifications for GWG (*z*-scores),<sup>16</sup> GWG categories,<sup>13,18</sup> focus on weight loss,<sup>12</sup> or study population (women with two previous CS attempting trial of labor,<sup>15</sup> only primiparas<sup>9</sup>). Additionally, some differences in classifications of intrapartum CS are likely, as definitions included unplanned/emergency,<sup>13,16</sup> during labor/emergency,<sup>18</sup> in labor,<sup>14</sup> following trial of labor/among women who had labored,<sup>9,15,19</sup> emergency/acute,<sup>17</sup> non-elective,<sup>12</sup> or primary CS due to dysfunctional labor/dystocia.<sup>10,11</sup> Though GWG adequacy within some populations was relatively similar to our cohort,<sup>10,12,17</sup> in others, excessive GWG was more<sup>9,14</sup> or less<sup>11</sup> common.

Nonetheless, the specific association between inadequate GWG and intrapartum CS necessitated by labor dysfunction among underweight and normal weight women is plausible. Excessive GWG among women with higher prepregnancy BMI may not substantially alter the pre-existing underlying physiology leading to intrapartum CS to the same extent as among women entering pregnancy with a lower BMI. For example, excessive GWG could have a greater impact on uterine contraction impairment and hormone and lipid levels among underweight and normal weight women given their relatively lower prepregnancy risk profile, whereas the impact of excessive GWG could be less among overweight and obese women given their relatively elevated prepregnancy risk profile. Additionally, mechanical risks of excess adiposity associated with labor dysfunction in the first stage of labor may be beneficial in the second stage of labor, as evidenced by findings of a shorter second stage of labor among overweight and obese women, potentially related to increased abdominal pressure and strength when pushing.<sup>25</sup>

The lack of association between GWG and intrapartum interventions among women with obese prepregnancy BMI could also reflect selection biases. Obstetricians may prefer to schedule these women for a planned CS (given their higher risk for intrapartum CS due to prolonged labor),<sup>7</sup> meaning that only the healthiest obese women with a higher likelihood of achieving a vaginal delivery attempt a trial of labor. Given that additional maternal characteristics were associated with intrapartum interventions, the potential mechanisms/ relation between BMI, GWG, and intrapartum interventions and other factors are complex. Additional research is needed to verify our finding of differences by BMI and to further explore the potential mechanisms for these associations.

The NPS provide extensive, rigorous data, obtained by specially trained study personnel within a large, nationally representative sample (based on comparisons of selected perinatal indicators, eg, maternal age and GA, available from birth certificate and hospital discharge statistics in the corresponding years).<sup>2</sup> To limit biases due to the correlation between GWG and length of gestation, we accounted for GA at birth in our definition of GWG adequacy,<sup>28</sup> as done in previous studies.<sup>20,23</sup> Additional methodological strengths are the large sample size and low level of missing data (~1.5% in multivariable analyses; >90% of women participating in NPS with necessary data on BMI, GWG, and outcomes for analysis). Given the lack of studies examining intrapartum CS or oxytocin administration, our study provides important information for providers, in

Limitations of our study include measurement error due to the inclusion of some self-reported (in particular for both prepregnancy and end of pregnancy weight) and retrospective data. Though selfreported weight may be underestimated<sup>29</sup> and the precision of timing of maternal weight estimates (for example, how close the end of pregnancy weight measure was to delivery) is unclear, because GWG is not reported in medical records in France in a standardized manner, this information was obtained by self-report to ensure complete and consistently reported data. Additionally, longitudinal GWG information was not available, preventing the evaluation of GWG trajectories or timing. Finally, although the NPS includes a large cohort of women, after stratifying by BMI and parity, we had low power to detect differences, particularly among women with obese prepregnancy BMI. Similar studies should be performed in countries with higher obesity rates to provide additional evidence within this group. While institutional differences in obstetric interventions are likely, the multi-level models used in our adjusted analysis accounted for these differences. As we have performed many statistical tests, results, particularly for other covariates included in the adjusted models, should be interpreted with caution and are only included for hypothesis generation and to understand the various factors that impact the associations.

### 5 | CONCLUSION

Among a nationally representative sample of women giving birth to singletons in France, excessive GWG was associated with increased oxytocin administration regardless of prepregnancy BMI and parity and intrapartum CS among women with underweight and normal weight prepregnancy BMI, providing further evidence for the benefits of healthy GWG for normal labor progression. Additional research in larger cohorts is needed to verify our findings and to further explore the potential mechanisms, in particular the role of BMI, for the associations evaluated.

#### AUTHOR CONTRIBUTIONS

Béatrice Blondel and Camille Le Ray contributed substantially to the design and data acquisition of the French National Perinatal Surveys. Melissa Amyx, Camille Le Ray, and Jennifer Zeitlin contributed substantially to the design of the current study. Melissa Amyx conducted data analysis, interpreted the results, and developed the draft manuscript under the supervision of Camille Le Ray and Jennifer Zeitlin and with input from Béatrice Blondel. All authors critically reviewed and approved the final manuscript.

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### CONFLICT OF INTEREST

The authors have stated explicitly that there are no conflicts of interest in connection with this article.

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#### SUPPORTING INFORMATION

Additional supporting information can be found online in the Supporting Information section at the end of this article.

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