

A Parity-Based Analysis of Women of Advanced Maternal Age and Delivery

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Abstract

Background: An increasing number of older mothers are a world-wide concern in obstetrical practice. First labor and delivery is quite different from the second one onward. Therefore, this study aimed to examine the differences in labor and delivery features between nulliparous and multiparous women of advanced maternal age (AMA) and their younger counterparts.

Methods: Women who delivered at our clinic between January 2004 and December 2022 were enrolled. The inclusion criteria were term (≥ 37 weeks), cephalic presentation, and singleton. We excluded women with fetal demise before labor onset, those with uncontrolled medical diseases such as hypertension, hyperthyroidism, diabetes mellitus, and those with scarred uteri. Nulliparous and multiparous women aged ≥ 35 years at delivery were regarded as having AMA, and were compared with women in their twenties as younger counterparts.

Results: In this study, 514 nulliparous and 1,116 multiparous women of AMA, and their younger counterparts (1,432 nulliparous and 845 multiparous women) were enrolled. Of the nulliparous women of AMA, pregnancy by assisted reproductive technology (ART) (13.4% vs. 0.8%, $P < 0.001$), labor induction (27.2% vs. 12.8%, $P < 0.001$), prolonged second stage of labor (27.7% vs. 14.8%, $P < 0.001$), cesarean section (7.2% vs. 1.1%, $P < 0.001$), and vacuum extraction (16.0% vs. 9.4%, $P < 0.001$) were significantly more frequent compared to their younger counterparts. Similarly, multiparous women of AMA showed a higher frequency or pregnancies by ART (5.7% vs. 0.1%, $P < 0.001$), labor induction (19.2% vs. 12.5%, $P < 0.001$), prolonged second stage of labor (5.2% vs. 2.1%, $P < 0.001$), and vacuum extraction (4.3% vs. 1.5%, $P < 0.001$) compared to their younger counterparts. However, cesarean section rates (0.2% vs. 0.0%, $P = 0.22$) were similar.

Conclusions: Pregnant women of AMA had different clinical risks due to parity. However, a vast majority had successful vaginal deliveries.

Keywords: Advanced maternal age; Labor and delivery; Duration of labor; Mode of delivery; Parity

Introduction

The increasing rate of cesarean sections (CS) is a global issue in obstetrical practice [1-3]. The higher rate of CS is due to the increasing number of older mothers. In many developed countries, it is common for young couples to delay marriage and pregnancy. In the United States, the age of mothers at first childbirth rose from 21.4 in 1970 to 24.9 in 2000, reaching 26.3 in 2014 and 27.4 in 2022 [4-6]. In Japan, the age at first childbirth increased from 25.7 in 1975 to 30.7 in 2019 [7]. In many European countries, mothers aged ≥ 35 years account for 20% of all births, and over 30% in Spain and Ireland in 2019 [8].

Advanced maternal age (AMA) is one of the dystocia-related factors in nulliparous women [9], and it is responsible for many detrimental effects on pregnancy, such as stillbirth, miscarriage, preterm labor, gestational diabetes mellitus, hypertensive disorders of pregnancy, and preeclampsia [10-12]. Reportedly, AMA is associated with higher CS rates in nulliparous and multiparous women [13-15]. However, we reported a very low primary CS rate in low-risk multiparous women [16]. Therefore, in this study, we aimed to examine the clinical differences in labor and delivery between nulliparous and multiparous women of AMA and their younger counterparts.

Materials and Methods

In this study, we enrolled women who delivered between January 2004 and December 2022 at our clinic in Shizuoka City, a central Japanese city with approximately 700,000 inhabitants. The inclusion criteria were: cephalic presentation, singleton, and term pregnancy (≥ 37 weeks gestation). We excluded women with fetal demise before labor onset, those with uncontrolled medical diseases such as hypertension, hyperthyroidism, and diabetes mellitus and women with scarred uteri. Women of AMA (≥ 35 years at delivery) and their younger counterparts in their twenties were analyzed for parity (only for multiparous women), maternal height, maternal prepregnancy body weight, prepregnancy body mass index (BMI), gestational age, gestational weight gain (GWG), neonatal weight, neonatal head circumference, amount of blood loss, pregnancy by assisted reproductive technology (ART), um-

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Table 1. Demographic and Clinical Characteristics of Women of AMA and Their Younger Counterparts (in Their Twenties)

Nulliparous women	Young, N = 1,432	AMA, N = 514	P
Age (years)	26.5 ± 2.3	37.0 ± 2.0	
Age ≥ 40, N (%), max.	0 (0), 29	67 (13.0), 45	
Maternal height (cm)	158.2 ± 5.1	158.6 ± 5.4	0.096 ^a
Prepregnancy body weight (kg)	50.3 ± 6.5	52.0 ± 6.9	< 0.001 ^a
Body mass index (kg/m ²)	20.1 ± 2.3	20.7 ± 2.5	< 0.001 ^a
Gestational age (days)	277.3 ± 6.7	277.7 ± 6.7	0.21 ^a
Gestational weight gain (kg)	10.9 ± 3.6	10.5 ± 3.4	0.031 ^a
Neonatal weight (g)	2,997 ± 334	3,020 ± 364	0.19 ^a
Neonatal head circumference (cm)	33.2 ± 1.2	33.3 ± 1.3	0.26 ^a
Pregnancy by ART, N (%)	11 (0.8)	69 (13.4)	< 0.001 ^b
Induction of labor, N (%)	184 (12.8)	140 (27.2)	< 0.001 ^b
Umbilical artery pH	7.30 ± 0.07	7.30 ± 0.07	0.24 ^a
Umbilical artery base excess	-3.7 ± 3.1	-4.0 ± 3.3	0.057 ^a
Apgar score at 5 min ≤ 7, N (%)	4 (0.3)	5 (1.0)	0.11 ^b
Blood loss (g)	407 ± 294	457 ± 312	0.002 ^a
Epidural, N (%)	61 (4.3)	44 (8.6)	< 0.001 ^b

Multiparous women	Young, N = 845	AMA, N = 1,116	P
Age (years)	26.9 ± 2.23	37.2 ± 2.0	
Age ≥ 40, N (%), max.	0 (0), 29	150 (13.4), 45	
Parity	2.2 ± 0.4	2.4 ± 0.7	< 0.001 ^a
Maternal height (cm)	158.0 ± 5.1	158.5 ± 5.1	0.034 ^a
Prepregnancy body weight (kg)	50.9 ± 7.5	52.5 ± 7.2	< 0.001 ^a
Body mass index (kg/m ²)	20.4 ± 2.9	20.9 ± 2.8	0.002 ^a
Gestational age (days)	274.7 ± 6.1	275.0 ± 6.5	0.28 ^a
Gestational weight gain (kg)	10.6 ± 3.3	9.7 ± 3.2	< 0.001 ^a
Neonatal weight (g)	3,051 ± 326	3,103 ± 368	0.0013 ^a
Neonatal head circumference (cm)	33.4 ± 1.1	33.5 ± 1.2	0.0025 ^a
Pregnancy by ART, N (%)	1 (0.1)	64 (5.7)	< 0.001 ^b
Induction of labor, N (%)	106 (12.5)	214 (19.2)	< 0.001 ^b
Umbilical artery pH	7.34 ± 0.07	7.32 ± 0.07	< 0.001 ^a
Umbilical artery base excess	-1.3 ± 2.8	-2.0 ± 2.9	< 0.001 ^a
Apgar Score at 5 min ≤ 7, N (%)	1 (0.1)	4 (0.4)	0.55 ^b
Blood loss (g)	333 ± 249	349 ± 259	0.16 ^a
Epidural, N (%)	36 (4.3)	78 (7.2)	0.011 ^b

^aUnpaired *t*-test. ^bFisher's exact test. AMA: advanced maternal age; ART: assisted reproductive technology.

bilical artery (UA) pH, UA base excess, Apgar score at 5 min ≤ 7, labor induction, use or non-use of epidural analgesia, duration of first and second stage of labor, and mode of delivery. Nulliparous and multiparous women were analyzed separately. Notably, forceps delivery was not performed in our clinic.

The local Ethics Committee approved this study (approval no. 23007). SPSS for Windows (version 22.0; IBM, Tokyo, Japan) was used for the statistical analysis. A value < 0.05 was

considered as statistically significant.

Results

We enrolled 514 nulliparous and 1,116 multiparous women of AMA, along with 1,432 nulliparous and 845 multiparous women as younger counterparts. Table 1 presents their demo-

Table 2. Mode of Delivery

Nulliparous women	Young, N = 1,432	AMA, N = 514	P ^a
Normal vaginal	1,292 (90.2)	395 (76.8)	< 0.001
Vacuum extraction	124 (9.4)	82 (16.0)	< 0.001
Cesarean section	16 (1.1)	37 (7.2)	< 0.001
Multiparous women	Young, N = 845	AMA, N = 1,116	P ^a
Normal vaginal	832 (98.5)	1,066 (95.5)	< 0.001
Vacuum extraction	13 (1.5)	48 (4.3)	< 0.001
Cesarean section	0 (0.0)	2 (0.2)	0.218

^aResidual analysis. Young: in their twenties; AMA: advanced maternal age, ≥ 35 .

graphic and clinical characteristics. The mode of delivery, indication for CS, duration of labor, and frequency of prolonged labor are presented in Tables 2-5, respectively.

Compared to their younger counterparts, the nulliparous and multiparous women of AMA exhibited significantly heavier body weight and BMI, lower GWG, higher rate of pregnancy by ART, labor induction, and use of epidurals (Table 1).

Regarding mode of delivery, nulliparous women of AMA had a higher rate of vacuum extraction (16.0% vs. 9.4%, $P < 0.001$) and CS (7.2% vs. 1.1%, $P < 0.001$), compared to their younger counterparts. In contrast, multiparous women of AMA had a higher rate of vacuum extraction (4.3% vs. 1.5%, $P < 0.001$) compared with their younger counterparts. However, they had similar CS rates (0.2% vs. 0.0%, $P = 0.218$) (Table 2). The indication for CS was mainly cephalopelvic disproportion/failure to progress among nulliparous women of AMA and their younger counterparts (Table 3). We enrolled only low-risk pregnant women; hence, no CS was performed before the onset of labor.

Nulliparous (median, 527 vs. 570 min, $P = 0.16$) and multiparous (median, 245 vs. 265 min, $P = 0.052$) women of AMA and their younger counterparts had similar duration of first stage of labor. However, when epidural users were excluded, the duration of first stage of labor in multiparous women of AMA was significantly shorter than that of their younger counterparts (median 245 vs. 265 min, $P = 0.035$) (Table 4).

Nulliparous and multiparous women of AMA had a sig-

nificantly longer duration of second stage of labor than their younger counterparts (nulliparous: median 77 vs. 50 min, $P < 0.001$; multiparous: median 14 vs. 11 min, $P < 0.001$) (Table 4). When epidural users were excluded, the duration of second stage of labor was also significantly longer in nulliparous and multiparous women of AMA than in their younger counterparts (Table 4).

Threshold comparisons showed that AMA group had a similar frequency of prolonged first stage of labor (> 20 h for nulliparous women and > 14 h for multiparous women) with younger counterparts in both nulliparous (12.2% vs. 13.6%, $P = 0.48$) and multiparous women (3.0% vs. 2.6%, $P = 0.68$). However, the frequency of prolonged second stage of labor (> 2 h and > 1 h for nulliparous and multiparous women, respectively; with 1 h added for epidural analgesia) was higher in women of AMA than in their younger counterparts in both nulliparous (27.7% vs. 14.8%, $P < 0.001$) and multiparous (5.2% vs. 2.1%, $P < 0.001$) women (Table 5).

Discussion

Many studies have shown that pregnancy in AMA is associated with adverse outcomes for both mother and baby [10-15]. In this study, we enrolled only women with low-risk pregnancies and aimed to clarify whether “healthy” AMA affected the mode of delivery and duration of labor irrespective of parity. As progress of labor and CS rate among nulliparous and multiparous women are quite different [17, 18], it is reasonable to analyze them separately. It has been shown in many studies that AMA is associated with longer labor durations and higher rates of instrumental delivery and CS [13-15]. However, the relationship between maternal age and labor duration is complex. Greenberg et al reported that the duration of labor and prolonged labor increased with increasing maternal age [19]. However, in their study, the duration of first stage of labor did not differ between young (< 20 years) and older women (20 to above 40 years) without epidural use in either the nulliparous or multiparous groups. The duration of second stage of labor differed between young and older women in the nulliparous and multiparous groups irrespective of epidural use. Zaki et al also reported that first stage of labor progressed more quickly with increasing age among nulliparous women up to 40 years

Table 3. Indication for Cesarean Sections

	Young	AMA
Nulliparous women		
CPD/FTP	12	34
NRFS	4	2
Forelying umbilical cord	0	1
Multiparous women		
CPD/FTP	0	1
Face presentation	0	1

CPD: cephalopelvic disproportion; FTP: failure to progress; NRFS: non-reassuring fetal status; Young: in their twenties; AMA: advanced maternal age, ≥ 35 .

Table 4. Duration of Labor

	Young	AMA	P ^a
Nulliparous women			
Duration of labor, first stage			
Total	570 (216 - 1,340)	527 (210 - 1,290)	0.16
Without epidural	560 (211 - 1,266)	510 (202 - 1,200)	0.069
Duration of labor, second stage			
Total	50 (18 - 145)	77 (25 - 253)	< 0.001
Without epidural	49 (18 - 129)	75 (23 - 177)	< 0.001
Multiparous women			
Duration of labor, first stage			
Total	265 (105 - 572)	245 (95 - 560)	0.052
Without epidural	265 (110 - 571)	245 (95 - 580)	0.035
Duration of labor, second stage			
Total	11 (4 - 31)	14 (4 - 43)	< 0.001
Without epidural	11 (4 - 30)	13 (4 - 42)	< 0.001

Data are presented as median (10 percentile, 90 percentile) (min). ^aMann-Whitney U test. Young: in their twenties; AMA: advanced maternal age, ≥ 35 .

old, and all multiparous women. In contrast, the duration of the second stage of labor increased along with increased maternal age [20].

Wang et al reported that among nulliparous women, AMA was associated with a higher frequency of CS before labor (odds ratio (OR): 2.26, $P < 0.005$) and in labor (OR: 1.44, $P = 0.016$), and more instrumental deliveries (OR: 1.49, $P = 0.004$) compared with their younger counterparts [21]. However, they reported similar CS rate in labor (OR: 1.22, $P = 0.168$), and instrumental delivery (OR: 1.35, $P = 0.058$) in multiparous women of AMA compared with younger multiparous women. Crequit et al also showed that among nulliparous women at term in spontaneous labor, AMA was not associated with an increased intrapartum CS rate [22]. Consistent with these previous studies, we suggest that women of AMA are not always associated with longer labor or more difficult labor.

Declined uterine contractility [23], decreased maternal physical strength [24, 25], and the obstetricians' lower threshold for performing CS in women of AMA [14, 26] could be the reasons why pregnant women of AMA have longer labor and

more frequent dystocia-related CS than their younger counterparts. Until recently, uterine contractility was believed to not decline with aging [27, 28]. However, Punson-Jimenez et al showed that uterine contractility declines with aging [23]. Elmenshawy et al [24] and Chomik and Jacinto [25] reported that female physical performance peaks in their twenties for many sports. These findings may explain the longer duration of labor and more frequent dystocia-related CS in women of AMA. Rydahl et al suggested that obstetrical culture such as early intervention and a higher induction rate for women of AMA may contribute to a higher rate of dystocia-related CS [14]. Zeevi et al postulated that a higher rate of CS in the older group might be due to physician's inclination to opt more easily for CS during labor [26].

Is there an age limit for vaginal delivery [26]? The author has never thought that women could be too old for vaginal delivery. However, if nulliparous women in their late 40s or 50s had visited our clinic for delivery, my opinion may change.

This study had several limitations. First, in both nulliparous and multiparous groups of AMA, the oldest women were

Table 5. Frequency of Prolonged Labor

	Young	AMA	P ^a	OR (95% CI)
First stage of labor				
Nulliparous women, > 20 h	13.6% (192/1,416)	12.2% (58/477)	0.48	0.88 (0.65 - 1.21)
Multiparous women, > 14 h	2.6% (22/845)	3.0% (33/1,109)	0.68	1.15 (0.68 - 1.97)
Second stage of labor				
Nulliparous women ^b	14.8% (210/1,416)	27.7% (132/477)	< 0.001	2.20 (1.72 - 2.82)
Multiparous women ^c	2.1% (18/844)	5.2% (57/1,111)	< 0.001	2.48 (1.46 - 4.22)

^aFisher's exact test. ^b> 2 h without epidural, > 3 h with epidural. ^c> 1 h without epidural, > 2 h with epidural. Young: in their twenties; AMA: advanced maternal age, ≥ 35 ; CI: confidence interval; OR: odds ratio.

45 years old (Table 1). If a significant number of mothers of AMA (≥ 45 years old) were included, our results would be different [29]. Second, in our clinic, epidural analgesia was used in $< 10\%$ of our cases (Table 1). In Japan, epidural use during labor has been low [30]. In 2016, only 6.1% of women used epidural analgesia during labor [30]. If epidural analgesia had been used more frequently, the duration of labor and mode of delivery may have been different. Third, this study was done in a single clinic, which may limit generalizability of the findings.

This study had several strengths. The decision to perform CS was made by a single physician, reducing variability. As CS rate is quite different among doctors and hospitals [31, 32], its indication may be quite subjective [33]. Hence, inter-hospital and inter-physician biases were avoided in this study. Second, a longer time was allowed for first and second stages of labor (Table 3). If CS had been performed quickly for cases with a slightly longer duration of labor, there might have been no difference in the duration of labor between women of AMA and their younger counterparts.

Furthermore, many women of AMA are anxious about their childbirth, and anxiety and fear of childbirth may cause dystocia [34]. This study's findings may relieve their fear of labor and delivery as the vast majority of women had successful vaginal births.

Conclusions

In nulliparous and multiparous women, the duration of second stage of labor was longer in those of AMA than in their younger counterparts. However, that of first stage was not. Multiparous women of AMA had similar CS rate, whereas nulliparous women of AMA had higher CS rate than their younger counterparts. Women of AMA are not homogenous, and their risks should be considered individually.

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Conflict of Interest

None to declare.

Informed Consent

Not applicable.

Author Contributions

KI performed all the research including research design, data collection and analysis, and manuscript writing.

Data Availability

The data of this study are available from the corresponding author upon reasonable request.

Abbreviations

AMA: advanced maternal age; ART: assisted reproductive technology; BMI: body mass index; CPD: cephalopelvic disproportion; CS: cesarean section; FTP: failure to progress; GWG: gestational weight gain; NRFS: non-reassuring fetal status; OR: odds ratio; UA: umbilical artery

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