#### **EFFECTS OF DIFFERENT PHASES OF MENSTRUAL CYCLE ON RESPIRATION** Hemalatha N. R<sup>1</sup>

#### HOW TO CITE THIS ARTICLE:

Hemalatha N. R. "Effects of Different Phases of Menstrual Cycle on Respiration". Journal of Evolution of Medical and Dental Sciences 2014; Vol. 3, Issue 65, November 27; Page: 14183-14188, DOI: 10.14260/jemds/2014/3899

**ABSTRACT: BACKGROUND:** Menstrual cycle is an integral part of life in women. The characteristic rhythmic changes in the rate of secretion of ovarian hormones produce physiological changes in different organ systems, in addition to changes in the reproductive system. **AIM:** The present study is done to elucidate a possible correlative changes in respiration during the different (menstrual, luteal and follicular) phases of menstrual cycle in healthy young girls. **MATERIAL AND METHODS:** Volume and rate of resting ventilation were measured serially in 40 healthy female student's volunteers, during menstrual, follicular and luteal phases of menstrual cycle. **STATISTICAL ANALYSES:** ANOVA, 'T' test and 'P' value are applied to find out statistical significance and conclusions are drawn based on this statistical treatment. **RESULTS:** Respiratory Rate was significantly higher (P<0.006) in luteal phase than in menstrual and follicular phases. The Minute ventilation though showed slight increase in LP, was not statistically significant. The Tidal Volume showed fall during LP. **CONCLUSION:** The luteal increase in ventilation suggests a possible role for progesterone in stimulating the respiratory drive, either centrally or through the peripheral chemoreceptors or by both, thus suggesting a possible beneficial role of progesterone in the management of respiratory illnesses.

**KEYWORDS:** Female, Follicular phase, Luteal phase, Menstrual cycle, Progesterone, Respiration, Students.

**INTRODUCTION:** Menstrual cycle is an integral part of life in women. The characteristic rhythmic changes in the rate of secretion of ovarian hormones produce physiological changes in the brain, in the musculoskeletal system, cardiovascular and pulmonary functions, in addition to changes in the reproductive system. Ovulation is followed by the formation of a corpus luteum, which secretes progesterone.

Though the effect of cyclical changes in hormonal secretion on reproductive system is wellknown, that on other organ systems has not been well established. The information on the variability of respiration which is relative to the menstrual phases is insufficient. The objective of the present study is to elucidate a possible correlative changes in respiration during the different (menstrual, luteal and follicular) phases of menstrual cycle in healthy young Indian girls.

**MATERIALS AND METHODS:** Forty healthy female student volunteers from Bangalore Medical College, Bangalore were studied throughout their menstrual cycle after taking written informed consent from them. Thorough history was taken, including menstrual history and complete clinical examination done before accepting them as subjects.

The selection of the subjects was based on fairly normal and regular menstrual history ( $28 \pm 2$  days cycle). Menstrual histories including regularity, associated dysmenorrhea, history of premenstrual symptoms like headache, irritability, edema and age of menarche were collected for screening the subjects. Female subjects with irregular menstrual cycles and/or using medications for excess bleeding of the uterus were excluded.

The respiratory rate (RR), tidal volume (TV) and minute ventilation (MV) were recorded using Benedict – Roth apparatus (for 6 minutes) along with basal oxygen consumption during each phase of menstrual cycle–1-5<sup>th</sup> day during menstrual phase (MP), 9-12<sup>th</sup> day during follicular phase (FP) and 19–22<sup>nd</sup> day during luteal phase (LP). The phases were corroborated with the period of ovulation, as judged from daily basal oral temperature of subjects throughout cycle.

Emotional and physiological distress was avoided by allowing a few trials of breathing through the apparatus before the day of actual test. The experiment was carried out under almost similar environmental conditions at the basal state in morning hours. To minimize the effect of training, if at all, the subjects were randomly allocated to be studied first in any of the three phases 12(MP), 16(FP) and 12(FP).

The RR (breaths/min) and TV (ml) were obtained from the recording chart, the product of which was taken as MV (L/min).

The procedure repeated during next two phases of menstrual cycle in each subject.

**STATISTICAL ANALYSES:** ANOVA, 'T' test and 'P' value are applied to find out statistical significance and conclusions are drawn based on this statistical treatment.

**RESULTS:** 40 female students were selected based on purposive sampling basis for the present study. The RR (breaths/min) and TV (ml) recorded during 3 phases of Menstrual Cycle.

The descriptive statistics (table 1a) and ANOVA (Table 1 b) of RR shows  $18.5\pm2.66$ ,  $17.85\pm2.55$  and  $19.73\pm2.65$  in MP, FP and LP, respectively with F- ratio =5.269 and p<0.006 suggestive of significant increase during luteal phase.

The Tidal Volume descriptive statistics (table 2 a) and ANOVA (Table 2 b) application has showed fall during LP.

The MV (RR×TV) though showed slight increase in LP, was not statistically significant. (Table 3a and 3b).

**DISCUSSION:** Functional lungs are able to adapt to the metabolic need of the body and accordingly the lung function varies during different conditions. Normal ventilation is a complex process involving several interactions. Behavioural, chemical and central neuromuscular drives influence breathing profoundly. The altered state of breathing during menstrual cycle probably involves all or any such mechanisms. Since hormonal levels vary as per the metabolism, any variation in the female gonadal hormone during different phases of menstruation exerts corresponding changes in the lung functions.

Our study showed slight increase in RR, MV and fall in TV during LP which can be explained as follows:

Increased levels of both estrogen and progesterone, with predominant effect of progesterone, characterize LP. These hormones have varied physiological actions, which can influence metabolism. Progesterone increases the sensitivity of respiratory Centres to  $CO_2$ , thus causing hyperventilation,  $\uparrow O_2$  intake and  $\downarrow$  alveolar and arterial  $CO_2$ .<sup>1</sup> Also, because of hyperventilation, breath becomes shorter, leading to decrease in tidal volume.

The increase in MV during LP has been shown in studies done by Milne et al,<sup>2</sup> Das and Jana.<sup>3</sup> Earlier studies indicate that hyperventilation during luteal phase of menstrual cycle and during pregnancy are associated with increased progesterone concentration.<sup>3,4</sup>

J of Evolution of Med and Dent Sci/ eISSN- 2278-4802, pISSN- 2278-4748/ Vol. 3/ Issue 65/Nov 27, 2014 Page 14184

Further, administration of progesterone to normal men and post-menopausal women produces hyperventilation.<sup>5,6</sup> Progesterone has also been used for treating hypoventilating patients.<sup>7,8</sup>

Increase in brain adrenergic receptors during pregnancy is associated with behavioural changes which can modify ventilation.<sup>9</sup> Recently it has been reported that luteal phase of menstrual cycle is also associated with increased adrenoreceptor density.<sup>10</sup>

Evidences suggest that both luteal phase of menstrual cycle and pregnancy are associated with central ventilatory drive as indicated by increase in mouth occlusion pressure.<sup>11,12</sup>

The physiologic changes observed during the luteal phase of menstrual cycle are known to mimic early pregnancy.<sup>13</sup> A significant increase in minute ventilation in the luteal phase was observed when compared to the menstrual and follicular phases.<sup>3</sup> This may be a result of increased oxygen consumption<sup>12</sup> or due to thyroid hormone over activity during the luteal phase.<sup>14</sup> Earlier studies indicate that an increase in progesterone secretion was a possible cause for the hyperventilation in the luteal phase.<sup>15</sup>

Study done by Das showed significant increase (P<0.0l) in minute ventilation was noted during luteal phase compared to both menstrual and follicular phases. Ventilation during the latter two phases was almost equal. Hyperventilation during the luteal phase was significantly high in VT only, without any appreciable change in respiratory rate.<sup>16</sup>

The most possible cause for an increase in lung function parameters during the luteal phase of menstruation and pregnancy is the hyperventilation associated with increased progesterone secretion.<sup>17,18</sup>

Periodic hyperventilation like during exercise improves respiratory muscle strength and lung capacity. Progesterone induces hyperventilation through both the central medullary and peripheral receptors and the sensitivity of respiratory receptors during the luteal phase and pregnancy.<sup>19</sup>

The fluctuation in respiration during different phases of menstrual cycle in healthy young women, highest in luteal phase, may be an important factor to be taken into consideration especially in women with predisposition to respiratory allergies. They may be prone to develop symptoms of asthma during premenstrual and menstrual phases when the progesterone levels fall.

**CONCLUSION:** In summary, the Minute Ventilation raised during Luteal Phase, along with increase in Respiratory Rate and decrease in Tidal Volume during Luteal Phase of menstrual cycle. However, this study was done in a small group of subjects and may not represent the whole Indian women. Further study needs to be performed.

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	Minimum	Maximum	Mean	Std. deviation	Std. error	95% confidence interval for Mean		
						Lower bound	Upper bound	
MP	15	24	18.5	2.66	.421	17.65	19.35	
FP	15	24	17.85	2.558	.404	17.03	18.67	
LP	15	26	19.73	2.65	.419	18.88	20.57	
Total	15	26	18.69	2.716	.248	18.20	19.18	
	Table 1a: Descriptive Respiratory Rate/min							

	Sum of squares	Df	Mean square	F-ratio	p-value		
Between Groups	72.517	2	36.258	5.269	< 0.006		
Within Groups	805.075	117	6.881				
Total 877.592 119							
Table 1b: ANOVA of Respiratory Rate/min							

	Minimum	Maximum	Mean	Std. deviation	Std. error	95% confidence interval for Mean			
						Lower bound	Upper bound		
MP	450	750	622.5	79.219	12.526	597.16	647.84		
FP	450	750	647.5	80.024	12.653	622.71	672.29		
LP	400	900	615.0	92.126	14.566	585.54	644.46		
Total	400	900	620.0	83.112	7.587	604.98	635.02		
	Table 2a: Descriptive statistics of tidal volume								

	Sum of squares	Df	Mean square	F-ratio	p-value			
Between Groups	1500	2	750	.107	>0.899			
Within Groups	820500	117	7012.821					
Total 822000 119								
Table 2b: ANOVA of tidal volume								

	Sum of squares	Df	Mean square	F-ratio	p-value
Between Groups	6.700	2	3.350	1.012	>0.367
Within Groups	387.277	117	3.310		
Total	393.977	119			

Pair		Mean	Std. deviation	Std. error mean	t-value	df	p-value	
1	MP	11.460	1.8842	.2979	0.171	39	>0.865	
	FP	11.517	1.9964	.3157				
2	MP	11.460	1.8842	.2979	1.547	39	>0.130	
	LP	11.988	1.5474	.2447				
3	FP	11.517	1.9964	.3157	1.489	39	>0.145	
	LP	11.988	1.5474	.2447				
Table 3b: Statistical Inference based on paired samples student's t-test on								

Minute Ventilation (L/min).

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> Date of Submission: 08/11/2014. Date of Peer Review: 09/11/2014. Date of Acceptance: 21/11/2014. Date of Publishing: 26/11/2014.