

## MORPHOGENESIS OF THE PROSTATE GLAND IN HUMAN FOETUSES

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### ABSTRACT

**Background:** To study the morphogenesis of the prostate gland in human foetuses. 'Prostates' is a Greek word which literally means "one who stands before", protector, guardian. It is important from a clinical point of view as it undergoes benign enlargement from the fifth decade, hence attracting the attention of males around this age and simultaneously the clinicians.

**Materials and Methods:** 112 foetuses of different gestational ages ranging from 14 weeks (85 mm) to 40 weeks (440 mm), products of terminated pregnancies under MTP Act of India, 1971 and stillbirths were collected from the Department of Obstetrics and Gynaecology, RIMS, Imphal and utilised for the present study with permission from the Institutional Ethical Committee.

**Results and observations:** The first time of appearance of the prostate gland to the naked eye till its definitive adult shape is studied. Growth and development at specific age period at different age groups are described.

**Conclusion:** Increase in vertical and transverse dimensions, assumption of adult shape were noted as the age changes. And at term, it has all the three components of the adult tissues although it is not as mature as in adult. This signifies that the growth of the prostate continues postnatally.

**KEY WORDS:** Prostate gland, human foetuses, morphology and development.

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### INTRODUCTION

The accessory genital glands such as: seminal vesicles, prostate and the bulbo-urethral glands produce secretions that are essential for the reproductive function in men [1]. The prostate

gland is the largest of the accessory sex glands found only in mammals that functions to produce a major fraction of the seminal fluid and surrounds the prostatic urethra from the bladder base to the membranous urethra and is

itself surrounded by a connective tissue capsule. All the three accessory glands of the male genital system develop near the junction between the mesonephric ducts and the pelvic urethra. From the dorsal wall of the allantois, the urorectal septum grows downwards to meet the cloacal membrane so dividing the cloaca and membrane into two: the urogenital sinus & urogenital membrane ventrally and dorsally the anorectal canal and the anal membrane. The urogenital sinus (endoderm) has three unequally sized parts. The uppermost and the largest is the vesical (vesicourethral) part. The middle or the pelvic part of the sinus forms the rest of the prostatic urethra, membranous urethra & the prostate (the surrounding mesoderm forming the fibromuscular stroma). By 13<sup>th</sup> to 15<sup>th</sup> week, testosterone concentrations having reached their peak embryonic levels, the prostate begins its secretory activity. Maturation of the gland continues while embryonic testosterone levels are high; however as testosterone level falls during the 3<sup>rd</sup> trimester, the gland enters a quiescent state [2].

The classical description of the prostate gland is that it is a pyramidal shaped fibromuscular exocrine organ that has the size, shape and consistency of a large chestnut [3].

It is traversed by the urethra which passes between its anterior & middle thirds. The ejaculatory ducts pass into the prostatic urethra [4].

### MATERIALS AND METHODS

One hundred and twelve foetuses of different gestational ages ranging from 14 weeks (85 mm) to 40 weeks (440 mm), products of terminated pregnancies under MTP Act of India, 1971 and stillbirths were collected from the Department of Obstetrics and Gynaecology, RIMS, Imphal and utilised for the present study with permission from the Local Ethical Committee. Only those foetuses which were free from any gross anatomical abnormality were selected for the present study. The age of the foetuses were assessed from the obstetrical history, crown rump length (CRL) and gross features before fixation. The foetuses were preserved in 10% formalin for 10 to 15 days.

The specimen were categorised into different age groups as follows for easier study and

observation.

GROUP 1	Upto 14 weeks
GROUP 2	14-18 weeks
GROUP 3	18-22 weeks
GROUP 4	22-26 weeks
GROUP 5	26-30 weeks
GROUP 6	30-34 weeks
GROUP 7	34-40 weeks

After proper fixation (for about 2 weeks) the foetuses were dissected. The abdomen was opened by a left paramedian incision. The incision was extended inferiorly to cut through the cartilage of the hip joint thus exposing urinary bladder and the prostate from the left. The specific age at which the gland first appeared in situ and was visible to the naked eye just below the urinary bladder was noted. The prostate gland was finally removed along with the urinary bladder and was studied for the growth and development and its changing size, shape and volume.

### RESULTS AND OBSERVATIONS

The first time of appearance of the prostate gland to the naked eye till its definitive adult shape (at term) is studied in detail. Growth and development at specific age period at different age groups are described as below. (Fig.1 )

**GROUP I: 14 Weeks:** In the earliest specimen of this series (14weeks; CRL-7.5 cm), the developing prostate is seen as jelly like, slight fusiform bulge along the upper half of the developing urethra i.e. in the lower part of the urogenital sinus (Fig.2). A mild angulation also exists in between the proximal swollen part and the caudal part of the developing urethra. Posteriorly, the anlage of the developing prostate is seen enveloping the cephalic end of the developing urethra.

**GROUP II: 14 -18 Weeks:** Prostate is more prominent and better defined than the previous age group and has gained in size, but still its length is more than its transverse measurement and thus it has an elongated look and is tapered distally (Fig.3). The developing prostate is discernable both on anterolateral as well as on posterior aspects as a swollen fold of tissue enveloping the upper third of the developing

urethra. It can be demarcated from the urinary bladder by a shallow groove and prostate is seen more prominent on the posterior aspect (Fig.4). Viewed from the side, the developing prostate looks like a tongue shaped structure merging with each other on the anterior aspect of the urethra.

**GROUP III: 18 -22 Weeks:** At subsequent age groups i.e. at 20 weeks it increases in size. It is seen developing transversely across the cephalic end of the urethra and at this stage, it assumes a more rounded appearance with increase in both transverse and anteroposterior measurements (Fig.5). There is increase in prostatic volume throughout from all its aspects. The prostatic swelling appears harder and more fibrous at this stage than the earlier stages. It is seen as a greyish white structure which could be the beginning of the formation of the capsule. Posterolaterally, the prostate is more prominent and the posterior surface looks flat.

**GROUP IV: 22-26 Weeks:** At this stage, there is overall increase in size in all dimensions, thus gain in anteroposterior and transverse measurements resulting as lateral convexities (Fig.6). The convexity on lateral aspect is more prominent as two protruding bodies whilst the dorsal surface is nearly flat with a median longitudinal groove.

The prostate at 22 weeks is seen to have a greyish white covering which is glistening and is probably the beginning of the capsule formation. It is firmer in consistency as compared to the previous age group. The development of the prostate is more enhanced posteriorly and is seen as two lateral protruding masses from the cephalic end of the developing prostate. There is a pyramidal shaped elevation on the anterior aspect tapering toward the apex which is recognized as the anterior lobe. From the anterior view, the two lateral lobe rudiments are seen as ovoid swelling jutting out from the main tissue. On lateral view, a small ovoid swelling with rough surface is recognizable as the representative of the lateral lobes. From the posterior aspect, the two lateral swellings got merged with each other behind to form a flattened posterior lobe. The posterior surface is bigger than the lateral and anterior lobes. Thus the growth of the developing prostate is responsible

to give a conical appearance with the broader cephalic end and a narrow tapered end caudally. In this age group, there is overall growth in all dimensions however, the transverse dimension is more than the vertical and anteroposterior dimensions.

**GROUP V: 26-30 Weeks:** It acquires a greyish white appearance losing its glistening jelly like appearance of previous age group at 26 weeks and the posterior surface shows reddish brown spots which are just visible to the naked eye and these spots may probably be the developing blood vessels (venous plexus). The prostate shows more increase in its transverse dimension. The posterior aspect seems to support the lower part of the urinary bladder in a funnel shaped way. The two lateral lobes are bulkier than the earlier age group and they fuse and get merged with each other in the same plane and form the posterior lobe (Fig.7). From the lateral view, the two lateral lobes are slightly higher than the anterior lobe and extend upwards and laterally. The capsule formation goes on and is well enhanced by 27 weeks as evidenced by a clear cut lower margin of the developing prostate and well separated out by a groove which demarcates it from the developing urethra. The upper margin is also separated out from the developing urinary bladder. The prostate thus acquires a firm looking appearance and is more fibrous and looks pinkish grey (because of the blood supply). The anterior lobe is represented by a band of fibromuscular tissue connecting the two lateral lobes in front of the urethra adjoining the neck of the bladder and is continuous posterolaterally to become the well formed lateral lobes. Thus at around 30 weeks, it has a stunted pear shape appearance whose anterior portion has developed into a firm mass.

**GROUP VI: 30-34 weeks:** With further increase in size in all dimensions, the shape of the prostate assumes as that of an adult. The prostate is well formed and has a firm nodular look and is reddish brown in color. As viewed from the front, the gland is firm due to more condensation of the fibres and muscles. The anterior lobe can be visualized which has a fibromuscular texture and is rougher than the other lobes. The two lateral lobes, as two eminences show

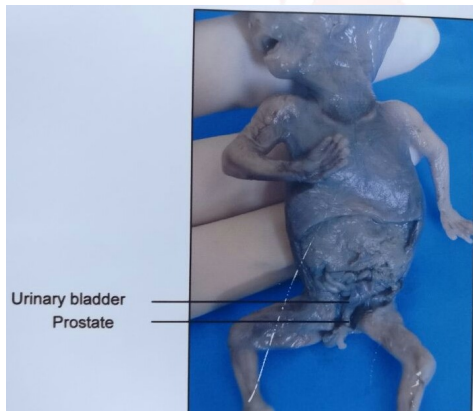


more growth posterolaterally and are extended on the posterior lobe and are seen merging with each other on the posterior aspect. The inferior margin of the prostate is well developed and surrounds the urethra all around it delineating it thus showing the well developed capsule. On the posterior surface is seen a network of blood vessels by 30 weeks visible to the naked eye (Fig.8).

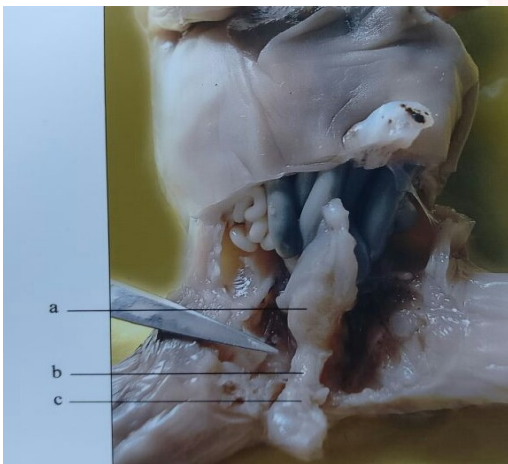
**Fig. 1:** Prostates at different weeks.



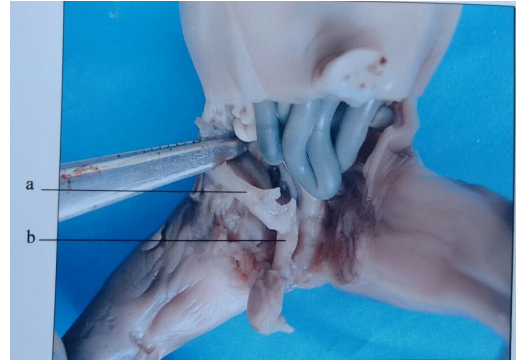
**Fig. 2:** Photograph of early developing prostate insitu seen as a fusiform bulge overlying on upper half of the developing urethra at 14wks visible to naked eye at first time.



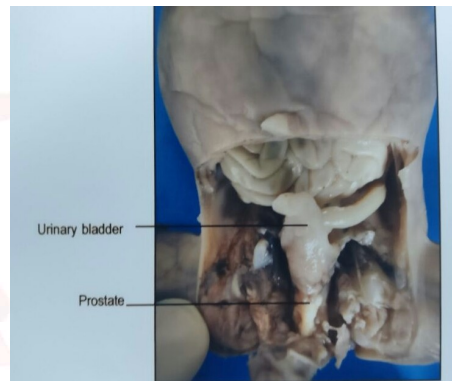
**Fig. 3:** Photograph showing anterior view of the developing prostate at 16wks old foetus. Note: a) urinary bladder b) prostate c) urethra.



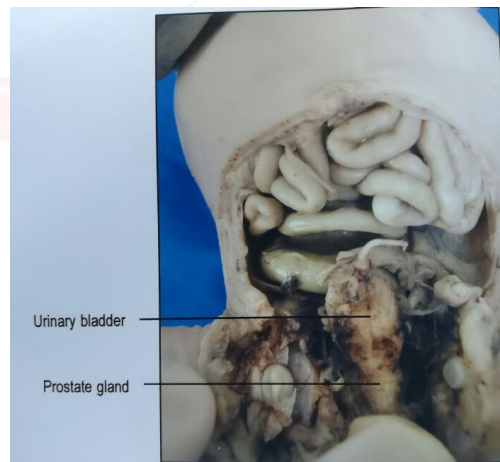
**Fig. 4:** Photograph showing the lateral view of the developing prostate at 16wks old foetus showing the demarcation from the urinary bladder by a shallow groove. a) bladder b) prostate.



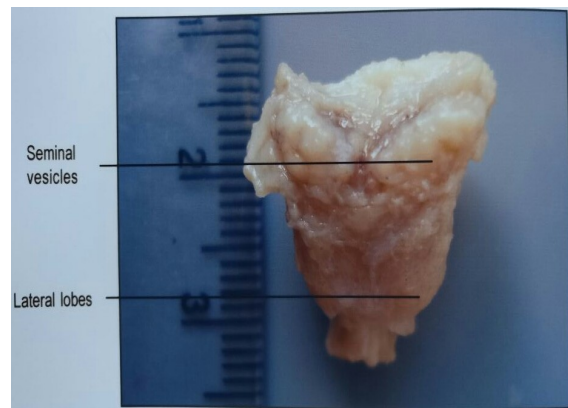
**Fig.5:** Photograph showing the anterior view of the developing prostate at 20wks.



**Fig.6:** Photograph of the anterior view of 26wks prostate.



**Fig. 7:** Posterior view of the prostate gland at 30wks.



**Fig. 8:** Photograph of 30wks prostate (posterior view) assuming the adult shape.



**Fig. 9:** Photograph showing the anterior view of the developing prostate at 40wks.



**GROUP VII: 34-40 weeks:** With further growth in this age group, there is overall increase in all dimensions (fig. 9). The prostate is firmer in consistency, harder than before and by 36 weeks, the two lateral protrusions are well developed and have a nodular rounded look presenting as two lobes with the central area flattened. Firm anterior lobe of the prostate is seen as a pyramidal shaped strip of prostatic tissue in front of the upper part of the urethra and neck of the bladder. The two lateral lobes seem to extend anterolaterally more or less assuming the adult shape. The false capsule is more distinct on the lateral lobes and is better defined as compared to the earlier age group. The vertical dimension decreases as seen from the anterior view and is about half of the horizontal dimension. The anterior lobe has a ratio of about one third of the lateral lobes. The posterior surface is flattened and the upper and lateral surface is occupied by the two lateral lobes.

## DISCUSSION

Development of the prostate begins with the growth of prostatic buds from the urogenital sinus at about 10<sup>th</sup> week of fetal development in

Majority of the workers have laid more emphasis on histogenesis. The prostate gland begins to develop in the 10<sup>th</sup> week as a cluster of endodermal evaginations that bud from the pelvic urethra and grow into the surrounding mesenchyme [9]. The prostate gland develops during the third month from interactions between the urogenital sinus mesenchyme and the endoderm of the proximal part of the urethra [10,11]. The multiple endodermal outgrowths arise from the prostatic part of the urethra and grow into the surrounding mesenchyme. The glandular epithelium of the prostate differentiates from these endodermal cells [7].

In males the inferior portion of the primitive urogenital sinus gives rise to the pelvic urethra and to penile urethra. By 6<sup>th</sup> week, the germ cells migrating from the yolk sac begin to arrive in the mesenchyme of the posterior body wall. The arrival of germ cells in the area just medial to the mesonephroi at the 10<sup>th</sup> thoracic segment induces cells of the mesonephroi and adjacent coelomic epithelium to aggregate into somatic sex cords that invest the germ cells. The sexual differentiation of genetic males begins at the end of the 6<sup>th</sup> week when a specific gene on the Y chromosome (SRY) is expressed in the sex cords cells. The product of this gene called the SRY protein initiates a development cascade that leads to the formation of the testes, the male external genitalia and the entire constellation of male secondary sex characteristics. The paramesonephric ducts degenerate. During the 3<sup>rd</sup> month the prostate and bulb urethral glands grow from the adjacent pelvic urethra [6].

The present workers while studying on developmental gross anatomy notice a gradual change in the size of the prostate as age advances. In the present study, the youngest prostate studied is of a 14 week old foetus and the prostate is visible as a longitudinal strip of mesenchymal induration which is pearly white and jelly like surrounding the upper half of the developing urethra.

The prostate is described to have four surfaces— anterior, posterior and two inferolateral surfaces [10]. The posterior surface is flat transversely and convex vertically. It is characterised by a midline groove that is wider toward the base of the gland and serves to partially separate the



gland posterior into right and left lateral lobes [3]. The posterior surface superiorly presents an annular groove through which the two ejaculatory ducts enter into the prostate. This annular groove subdivides this surface into upper smaller and lower larger parts. The latter is again subdivided by a median vertical groove into two lateral right and left lobes. The upper or superior part belongs to the median lobe and intervenes between the two ejaculatory ducts posteriorly and the prostatic urethra anteriorly. The lateral lobes are connected together anteriorly in front of the urethra by a fibromuscular band known as the isthmus which has no glandular tissue [10]. The posterior surface is described as nearly flat [12]. According to our present study, the posterior surface is seen as a merging surface from the two bulge of lateral lobes in a funnel shape with a V shaped trough toward upper end in which seminal vesicles are also converged towards the midline.

The numbers 4,3,2 indicate the transverse, vertical and anteroposterior dimensions of the gland in cms [10]. In our present study, the anteroposterior, transverse and vertical dimensions are measured. The mean transverse dimension of 38-42 week old foetus is 14.75mm, that of vertical is 10.17mm, and the anteroposterior dimension is 7.17mm.

## CONCLUSION

From the above results and observations in detail, the conclusions are drawn:

At 14 weeks, the prostate is visible to the naked eye as pyramidal shaped, jelly like tissue surrounding upper half of the urethra, the vertical dimension more than both anteroposterior and transverse dimensions. The capsule is also well noticeable at this stage.

Subsequently, around 18-20 weeks, the prostatic height reduces with simultaneous increase in anteroposterior and transverse dimensions.

Development of the two lateral lobes is attributed to the increase in transverse dimension. The anterior lobe is well defined as a fibromuscular band connecting the two lateral lobes.

By 26<sup>th</sup>-30<sup>th</sup> weeks, the prostate becomes firmer and loses its jelly like appearance gradually assuming the morphology of adult prostate i.e,

transverse dimension more than the anteroposterior dimension. From 34<sup>th</sup> week onwards, to the naked eye, the prostatic shape is almost like adult i.e, different lobes are visible to the naked eye. The two lateral lobes are the most recognisable lobes on either side, the anterior lobe as a band in front of the urethra, the posterior lobe as a flattened surface behind the urethra with the seminal vesicles above it as in adult.

**Conflicts of Interests: None**

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