Original articles

Dynamic changes in the zone of apposition and diaphragm length during maximal respiratory efforts

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Abstract

Background – Clinical tests of diaphragmatic strength are limited by the wide normal variation in maximal pressure which result, in part, from changes in diaphragmatic length. During relaxation at different lung volumes diaphragmatic length (LDI) can be estimated from the length of the zone of apposition (Lzapp) and the transverse diameter of the rib cage (Drc). A study was carried out in two subjects using sequential digital radiography at six frames/second to determine whether these relations apply during maximal respiratory efforts which distort the rib cage and diaphragm.

Methods – The length of the anteroposterior contour of the diaphragm and Drc were determined by curve fitting. Lzapp was measured with a millimetre rule.

Results – A significant correlation was found between LDI and Lzapp during both maximal inspiratory and expulsive manoeuvres ($R^2=0.98$ and 0.52). LDI was estimated from the measurements of Lzapp and Drc using a multiple regression equation derived from measurements during static relaxation. Despite the complex dynamic events at the onset of these “static” manoeuvres, actual LDI correlated strongly with derived LDI using all data for the two manoeuvres in each subject ($R^2=0.95$ and 0.84).

Measurements with ultrasonography (12 cm linear probe) and magnetometers confirmed the changes in Lzapp and Drc during inspiratory and expulsive efforts.

Conclusions – Non-invasive measurements of Lzapp and Drc can be used to derive an accurate estimate of diaphragmatic length under dynamic conditions.

(Maximal pressures developed by the diaphragm depend, not only on its neural drive and muscle strength, but also on its force-length and force-velocity relations. Other factors, including the cross sectional area of the lower rib cage and diaphragmatic shape, may also be important. Using sequential digital radiography at six frames/second we observed progressive muscle shortening at the onset of maximal “static” inspiratory efforts which move the diaphragm to a less advantageous position on the force-length curve. By contrast, the “extra” pressure developed during expulsive manoeuvres with the glottis held open was associated with transient lengthening of the diaphragm produced by the abdominal muscles. The changes in length and velocity of diaphragmatic muscle explained much of the difference in maximal voluntary transdiaphragmatic pressure between the manoeuvres. These observations suggest that measurements of the diaphragmatic muscle length and velocity might improve the predictive power of clinical tests of respiratory muscle strength.

Because diaphragmatic length (LDI) is difficult to measure in vivo, interest has focused on the relation between LDI and the width of the zone of apposition between the diaphragm and chest wall (Lzapp). A linear relation exists between LDI and Lzapp (measured from radiographs) during relaxation at lung volumes between residual volume (RV) and total lung capacity (TLC). During an “isovolume manoeuvre”, however, in which LDI is essentially constant, Lzapp decreases as the diameter of the rib cage (Drc) increases because the diaphragm “peels away” from the chest wall. Rochester and coworkers examined radiographs taken during relaxation at different lung volumes and found that 95% of the variance of Lzapp can be accounted for by measurements of LDI and the transverse diameter of the rib cage (Drc). Further studies in anaesthetised dogs showed that Lzapp correlated well with LDI during unloaded respiration. The authors concluded that measurement of Lzapp may be invaluable in the study of breathing mechanics. However, these measurements have not been reported for the strong respiratory efforts used to test diaphragmatic strength and endurance in human subjects. There are substantial distortions of...
the rib cage and diaphragm during these efforts, and these might influence the relations between Lzapp, LDi, and DRC.

The present study was designed to examine the correlated changes in Lzapp and DRC during maximal voluntary efforts. Measurements were made with sequential radiography and confirmed with ultrasonography.

**Methods**

Because the experiment involved exposure to radiation of more than 10 millisieverts (mSv), only two subjects (authors) were used (DM, subject 1, 39 years, 189 cm, 76 kg; SG, subject 2, 36 years, 172 cm, 62 kg). Both were trained in the respiratory manoeuvres so that the lowest exposures could be achieved (see below). The recording procedures were approved by the institutional ethics committee. All radiological procedures were supervised by the radiation protection physicist from our institution (GS), and informed consent was obtained. The methodology has been reported in detail elsewhere and is described briefly here.

**RECORDING PROCEDURES**

Oesophageal, gastric, and transdiaphragmatic pressures were separately recorded with a single catheter. Mouth pressure was also measured proximal to a shutter. An X-ray detector on the abdominal wall provided a timing signal to allow precise temporal correlation between the radiographic images and physiological data. All analogue signals and the timing signal were recorded on tape (DC, 1-25 kHz).

The maximal inspiratory efforts were performed against a shutter with minimal elevation of abdominal pressure. During expulsive efforts the subject attempted to keep the glottis open. Each manoeuvre began at a relaxed end expiratory position and was repeated six times.

Radiographs were obtained using a digital angiography unit (Phillips DVI) equipped with a 35 cm image intensifier. The subject was seated and markers were taped over the spine and the ninth interspace in the mid axillary line. Images were obtained at six frames/second with anteroposterior and lateral beams (11-16 frames in each contraction). Only the anteroposterior images were used for this study. In subject 2 anteroposterior images enclosed the full width of the lower chest and upper abdomen. In subject 1 the field enclosed only the left half of the chest and abdomen so measurements were doubled to give approximate LDi and DRC. Images were printed with minimal distortion on 10 x 13.5 cm film.

Values for the pressures were measured at the onset of each radiation pulse (duration 20 ms). To digitise the diaphragm profile a grid was placed on each radiograph overlaying a spinal marker and six points along each hemidiaphragm were resolved into cartesian coordinates. Coordinates for the lateral costal origins of the diaphragm were measured from anatomical landmarks defined at TLC. Lzapp was assumed to be zero at TLC. A curve was fitted by computer so that the total length of the diaphragm (or hemidiaphragm) could be calculated. Coordinates were adjusted for display and imaging scaling. We assumed that one quarter of the resting length comprised an inextensible tendon. The length of Lzapp was measured from the same images using a transparent ruler graduated in millimetres. The diameter of the rib cage (DRC) was derived for each frame as the widest horizontal distance encompassed by the curve fitted to the diaphragm coordinates.

Ultrasonography was used to assess changes in Lzapp in the same subjects in separate studies conducted after the radiographic imaging (Accuson XP-128). DRC was measured with linearised magnetometers attached to the skin in the mid axillary line by double-sided tape. Measurements were made using a 12 cm linear array probe (3-5 MHz) held firmly against the rib cage in the longitudinal axis just anterior to the magnetometer in the right mid axillary line. All images were stored on video tape together with a simultaneous signal of Pdi. Oesophageal and gastric pressures and Poi and DRC were also recorded on magnetic tape. The length of the probe ensured that the full extent of Lzapp could be followed throughout each manoeuvre (at 18-28 frames/second). Before each maximal manoeuvre the subject inhaled to TLC so that the costal insertion of the diaphragm could be visualised. Other methods were similar to those of the radiographic study. Images were replayed and

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**Figure 1** Representative profiles of left hemidiaphragm (anteroposterior projection) of subject 1 showing changes in shape and position during expulsive and inspiratory manoeuvres. Horizontal arrow heads indicate length of zone of apposition from insertion (lowest symbols). Diagonal arrows show direction of motion of diaphragm costal insertion during manoeuvres.
measured on a high resolution screen. Approximately one in four images was measured so that the overall sampling rate was comparable to that for the sequential radiography (six frames/second).

**Results**

The radiographic study showed that both diaphragmatic manoeuvres were associated with complex changes in Lzapp, LDI, DRC, and dome shape (fig 1). The manoeuvres differed in the direction of motion of the origin of the diaphragm, in the shape of the dome, and in the changes in Lzapp. The lateral costal margins moved outwards and cephalad at the onset of the inspiratory manoeuvre whereas, during the expulsive effort, they moved inwards and caudad. The dome became flattened with greater radii of curvature during the development of maximal inspiratory pressure, whereas the expulsive manoeuvre produced a more rounded shape with a reduction in radii of curvature.

All data for Pdi, Lzapp, and DRC are shown for subject 1 in fig 2. There was progressive shortening of the diaphragm at the onset of the inspiratory contraction with a considerable decrease in Lzapp. The absolute decrease in Lzapp (22 mm to the peak pressure) was more than twice the reduction in length of the hemidiaphragm. This was due to the outward motion of the rib cage (increase in DRC) which resulted in "peeling away" of the diaphragm from the chest wall independent of LDI (see also fig 1). During inspiratory efforts in subject 2, however, the decrease in Lzapp was almost identical to the decrease in hemidiaphragm length (25 mm) because there was minimal change in DRC in two of the three trials.

Diaphragmatic motion during pressure development in the expulsive manoeuvre was biphasic. During the initial phase (to about 70% Pdimax in the expulsive effort) there was progressive shortening of Lzapp and LDI in both subjects. The subsequent pressure development (above that in the inspiratory manoeuvre) was associated with an increase in Lzapp and LDI. In subject 1 the increase in Lzapp preceded that of LDI whereas, in subject 2, both parameters changed simultaneously. This may reflect differences in the timing of abdominal muscle recruitment or in rib cage compliance.

**CORRELATIONS BETWEEN Lzapp, LDI AND DRC**

The regression equations relating Lzapp to LDI and DRC in each subject are given in the table. There was a significant correlation between LDI and Lzapp in both subjects. In the dynamic phase of these manoeuvres, however, the relation between DRC and Lzapp was variable, both between subjects and the manoeuvres. In subject 1:

\[
\text{Lzapp} = -0.01 + 0.006 \times \text{LDI} + 0.0025 \times \text{DRC} 
\]

\[
R^2 = 0.624**
\]

Subject 2:

\[
\text{Lzapp} = -0.024 + 0.007 \times \text{LDI} + 0.002 \times \text{DRC} 
\]

\[
R^2 = 0.532**
\]

**Regression equations relating Lzapp to LDI and DRC**

\[
\begin{align*}
\text{Expulsive} & \\
\text{Subject 1} & \quad \text{Lzapp} = -0.01 + 0.006 \times \text{LDI} + 0.0025 \times \text{DRC} \\
& \quad R^2 = 0.624** \\
\text{Subject 2} & \quad \text{Lzapp} = -0.024 + 0.007 \times \text{LDI} + 0.002 \times \text{DRC} \\
& \quad R^2 = 0.532** \\
\text{Inspiratory} & \\
\text{Subject 1} & \quad \text{Lzapp} = -0.031 + 1.004 \times \text{LDI} + 0.002 \times \text{DRC} \\
& \quad R^2 = 0.887** \\
\text{Subject 2} & \quad \text{Lzapp} = -0.0123 + 0.400 \times \text{LDI} + 0.002 \times \text{DRC} \\
& \quad R^2 = 0.780** \\
\end{align*}
\]

Lzapp: length of the zone of apposition of the diaphragm; LDI: length of the diaphragm (anteroposterior projection); DRC: maximal diameter of the rib cage (anteroposterior projection). All values in mm.

* p < 0.05; ** p < 0.001.
ject 1 there was an inverse correlation during the inspiratory manoeuvre but no significant relation during the expulsive manoeuvre. In subject 2 the inverse correlation was better for the expulsive than the inspiratory manoeuvre. This reflected the small changes in DRC in this subject, especially for the inspiratory manoeuvre.

To determine whether LDI during maximal respiratory manoeuvres could be reliably estimated from measurements of Lzapp and DRC we substituted the values obtained in this study in the multiple regression equation of Rochester and coworkers. This equation \(\text{LDI} = 2 \times (0.984 \times \text{Lzapp} + 0.462 \times \text{DRC} + 34.8)\) in mm; \(R^2 = 0.94\) was based on data from 16 subjects who had static radiographs taken during relaxation at various lung volumes over the full range of vital capacity. It gives the length of the right hemidiaphragm and we have doubled it for total diaphragm length. Figure 3 shows the relation between LDI measured directly from each radiograph and LDI derived from this equation by substituting Lzapp and DRC measured from the same radiographs. The derived values lie within the 95% confidence intervals for the equation with a relatively small scatter of points around the regression lines \((R^2 = 0.95\) and \(0.84\) for subjects 1 and 2 respectively).

MEASUREMENTS WITH ULTRASONOGRAPHY

To determine whether the same pattern of change in Lzapp could be observed with a technique which did not require radiation exposure, the inspiratory and expulsive manoeuvres were repeated while Lzapp was continually monitored by ultrasonography (up to 28 images/second). A 12 cm linear array probe was oriented longitudinally in the mid axillary line. In both subjects an overt reduction was observed in Lzapp in the dynamic phase of strong inspiratory efforts performed near FRC. There was also a secondary increase in Lzapp during expulsive efforts at or before the peak Pdi. Typical responses are compared for the two methods in fig 4. A formal comparison of the radiographic measurements of DRC and those obtained by magnetometers was not undertaken.

Discussion

Several studies have emphasised the likelihood of diaphragm length changes during the quasistatic manoeuvres used to test diaphragmatic strength. In a previous study we have shown that the diaphragm muscle fibres may shorten by 25% of their resting length during pressure development in a maximal inspiratory manoeuvre, while length changes at the onset of expulsive manoeuvres are biphasic. Because it has recently been shown that LDI can be predicted from measurements of Lzapp and DRC during relaxation at different lung volumes, the present study assessed this prediction during two of the voluntary manoeuvres commonly used to test respiratory muscle strength. A relation which holds during relaxation at different lung volumes, or during ventilation in an anaesthetised animal, may not apply to these maximal quasistatic manoeuvres which are associated with substantial distortions of rib cage and diaphragm shape.

Our results show that the relation between true LDI (measured) and LDI derived from measurements of Lzapp and DRC using the equation developed by Rochester and colleagues held in spite of differences between the two subjects in the maximal pressures achieved and in the rib cage movements associated with the manoeuvres.

The absolute values obtained for LDI, Lzapp, and DRC were slightly lower than those
reported by Rochester and colleagues. The most likely explanation is a systematic difference in the method of measurement and scaling. The images (10 × 13.5 cm) in the present study were measured by deriving Cartesian coordinates from a grid and application of a scaling factor, while Rochester and colleagues used a flexible tape measure on large films. No mention was made of correction for any magnification. Another potential source of error is the location of the insertion of diaphragmatic fibres: this was derived from anatomical landmarks in both studies. Such an error might influence the absolute values for LDI and Lzapp but not the changes during the dynamic manoeuvres. This problem would be less likely to occur with ultrasonographic measures (provided that a sufficiently long probe is used) because the origin of the costal fibres can be followed to the chest wall and viewed repeatedly during inspiratory efforts at TLC.

The predictive power of the multiple regression equation introduced by Rochester and colleagues was high under static conditions covering the full range of lung volumes. The novel result of our study was the ability of measurements of Lzapp and DRC to predict LDI accurately during dynamic respiratory manoeuvres in which other factors operate. These include rapid transient changes in diaphragmatic length and distortions of the diaphragmatic dome and rib cage. In spite of these limitations, however, the present data, combined with those of Rochester, show that more than 85% of the variance of the estimate of LDI is accounted for by Lzapp and DRC.

Digital sequential radiography was used here initially to provide quality images, although the radiation dose is unacceptable for routine application. Although the frame rate is higher with most videofluoroscopic units, the visual contrast is not sufficient to trace the diaphragm contour with confidence during dynamic manoeuvres (McKenzie and Gandevia, unpublished observations). However, it may be sufficient to obtain accurate measurements of Lzapp. In the latter study, using videofluoroscopy in dogs, it was necessary to use radio-opaque markers sewn onto the costal diaphragmatic surface to obtain LDI. To make the comparison of derived and actual LDI responses reported here it was necessary to have the best possible image quality. We could not justify exposing more healthy subjects than the two authors simply to determine the general applicability of the method. Rochester's work shows that the method is accurate for healthy subjects during relaxation. We have explored the physiological limits of the method in two healthy subjects. That 85% of the variance of LDI could be accounted for by Lzapp and DRC in two subjects who varied substantially in physical build and performance of the manoeuvres suggests that the technique may have broad application. It remains to be shown whether the technique will allow accurate derivation of LDI in patients, particularly in those with considerable hyperinflation.

It is already established that ultrasonography can be used to assess the thickness of the diaphragm. Our results using ultrasonography show that it is possible to follow the dynamic changes in Lzapp with good time resolution and that these changes are quantitatively comparable to those observed radiographically. Thus, in healthy subjects, non-invasive measurements of Lzapp and DRC could be used to provide estimates of diaphragm length over a range of dynamic conditions. It is too early to predict a clinical application for this method of estimating LDI.

It will certainly be possible in healthy subjects to normalise values of Pmax for changes in LDI and diaphragm thickness. Measurements of cross sectional area of the thorax may also help to convert measurements of pressure into estimates of muscle force or tension.

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