2.1 Ultrasound Modalities

The first US examination of small parts was reported by Howry in 1955. Thyroid sonography (A- and B-scan) was first introduced in 1966–1967 (Fujimoto et al. 1967). It has been widely practiced since the 1970s and is now one of the most popular radiological methods for diagnosing thyroid diseases.

The method is based on differences between the abilities of different tissues to reflect US waves (cyclic sound pressure of an elastic medium with a frequency greater than 20,000 Hz). Modern US scanners permit real-time imaging of organs with constant monitoring of their motion.

Thyroid US has the following advantages:

- It is relatively simple, rapid to perform, and inexpensive.
- It is painless and noninvasive.
- This is no need for any special preparation of the patient before the examination.
- There are no contraindications.
- It is harmless and safe for the patient and staff. US can be used repeatedly in children, pregnant and nursing women, as well as seriously ill patients with severe concomitant pathology.
- Patients can be examined regardless of their medications, including thyroid blocking agents.
- It is a high-resolution technique.
- The differential diagnosis is based on sonographic options, such as Doppler modalities, 3D image reconstruction, and others.
- It supports documentation of video data and static images, as well as easy transmission via modern communication channels with virtual consultations.
- It provides easy guidance for minimally invasive modalities, such as FNAB, PEI, PLA, PGA, and others.

A patient is indicated for thyroid ultrasound in the following cases:

- Complaints that are often a consequence of thyroid pathology: dyspnea, cough, irritability, palpitation, precordial discomfort
- Palpated masses in the anterior neck
- Thyroid pathology detected by other methods
- Cardiovascular pathology, predominantly heart rhythm abnormalities
- Persistent diseases of ENT organs (such as larynx, pharynx, trachea), dysphonia, or aphony
- Dysphagia
- Monitoring of the efficacy of treatment of thyroid diseases
- Postoperative follow-up

Sonography can be utilized as a screening method for thyroid diseases. It permits early detection of patients who are at an increased risk of developing a thyroid disease. Screening is an effective initial stage of evaluation within a target population (Parshin et al. 1999). It helps to pinpoint a possible thyroid abnormality at an early stage, and includes the elements of differential diagnosis that result in subsequent thorough examination and timely treatment in appropriate cases. The advantages of US as a screening method are patient safety, reproducibility, reduced dependence on the quality of the equipment and operator skill, speed, availability, and low cost. The disadvantage of US screening is its comparatively low diagnostic accuracy. A negative screening study does not guarantee the absence of the disease, and sometimes a positive study does not necessarily prove that a thyroid pathology is present. In practice, one example of screening is thyroid US performed by a general practitioner with a simple (e.g., only grayscale) scanner. The exam aims to divide patients into two generalized categories:
those whose thyroids are grossly normal, and those with suspicious abnormalities in their thyroids.

Patients with thyroid abnormalities are subject to further complex qualified US. Complex US assumes the detection and certain differential diagnosis of diffuse changes and focal lesions, which is necessary to determine subsequent tactics.

US options utilized for the diagnosis of thyroid diseases include the following:

1. Grayscale
2. Tissue harmonics
3. Adaptive coloring
4. CDI
5. PDI
6. Grayscale 3D
7. Vascular 3D
8. 4D
9. Panoramic scan
10. Spectral pulsed wave Doppler
11. Others (multislice view, volume CT view, contrast US, US elastography, etc.)

Grayscale (B-mode, 2D mode) is a well-known basic type of scanning that provides an image of the thyroid in typically 256 shades of gray (Fig. 2.1).

The harmonic (the second harmonic, tissue harmonic imaging or THI, tissue harmonic echo) is an algorithm that allocates the harmonic component of fluctuations after the base US impulse has passed though the tissues (Fig. 2.2). It is often available as an option on grayscale scanners with standard probes.

THI enables the diagnosis of 70.8% of patients with thyroid pathologies (Miheeva 2007). THI emphasizes the US signs of thyroid cancer (visualization improves in about 28–30% of cases). It permits a more accurate definition of lesion margins, calcifications, and nodule structure. According to Belashkin et al. (2003), THI improves the visualization quality and defines features of colloid nodules in 80% of cases.

Adaptive coloring utilizes a color map in order to stain a grayscale image. The density of staining depends on the strength of the reflected echo (Fig. 2.3). Color inversion of the image is possible. The option is often added to grayscale scanners with standard probes.

This option improves the subjective perception of an US image. Thus, it helps in the detection of isoechoic thyroid lesions, and in the definition of nodule contours and posterior acoustic changes, especially in small lesions (up to 0.5–0.7 cm in size).

The vascularity of an abnormal thyroid can be characterized by Doppler modalities. The following aspects require special attention:
2.1 Ultrasound Modalities

Blood flow in superior and inferior thyroid arteries
- Vascularity of the parenchyma of the thyroid
- Vascularity of the lesions

Color Doppler imaging (CDI; color flow imaging or CFI; color flow mapping or CFM) is an US technology for visualizing vascular structures. It is based on recording the blood flow velocity and using color encoding to superimpose this velocity onto the grayscale image (Fig. 2.4). This option is incorporated into most modern scanners.

CDI is especially valuable for diagnosing thyroid malignancy. However, some authors consider it to be of limited academic interest and of minor importance for the differential diagnosis of thyroid nodules (Hübsch et al. 1992; Klemens et al. 1997).

The vascular architectonics of the parenchyma (parenchymal blood flow) of the thyroid is usually characterized by the following:
- Vascular pattern intensity
- Symmetry (between the lobes and the segments)
- Regularity of vascular structures within the thyroid parenchyma
- Deformations of the architectonics

The condition of the parenchymal blood flow is an important US indicator of the thyroid status. The vascular pattern in diffuse thyroid diseases is sometimes characterized by the number and the density of color pixels within the parenchyma using the following methods:

1. The color pixel density (CPD) is numerically expressed as the ratio of the area covered by color pixels to the total area of the image (in parts or percent) (Fig. 2.5). Similar calculations can be carried out in three-dimensional US (ratio of volumes). The CPD of a normal thyroid is about 3–15% (Fein et al. 1995; Lelyuk et al. 2007). The CPD is considered to be “increased” if it exceeds 15%, and “decreased” if it is less than 5%.
2. Scoring the number of color cartograms in area units. This is usually performed manually after

Fig. 2.3 (a–d) Thyroid sonograms. Adaptive coloring (a) conventional gray scale mode (b) different varieties of image color
2 Complex Ultrasound Diagnosis of Thyroid Diseases

**Fig. 2.4** (a, b) Thyroid sonograms. CDI (a) hypervascular thyroid nodule in CDI (b) thyroid nodule with peripheral blood flow pattern

**Fig. 2.5** CPD measurement

**Fig. 2.6** (a, b) Measurement of the number of color cartograms in a unit area (1 cm²) (a) transverse scan (b) longitudinal scan

dividing the image into uniform squares with sides of 1 cm (Fig. 2.6). Color spots from different vessels are recorded in every square. The calculation is approximate, because interstitial vessels are not straight; they can appear in various scanning planes, resulting in separate visualization of the fragments. High scanning frequency and high averaging are utilized for accurate calculation. The proposed reference range for vessel density in a normal thyroid is between 0.4 and 2.5 vessels per 1 cm² of thyroid tissue (Fein et al. 1995; Lelyuk et al. 2007).

3. Scoring the amount of color cartograms within the lobe (Fig. 2.7). Only color spots from different vessels are counted. The accepted reference range for a normal thyroid is between five and ten vessels within the lobe (Fein et al. 1995; Lelyuk et al. 2007). The occurrence of more than ten vessels within the lobe at once is interpreted as an increase in parenchymal blood flow. A decrease is characterized by fewer than five vessels within the thyroid lobe.

Sekach et al. (1997) and Laszlo et al. (1998) suggest that the following three vascular patterns can occur in thyroid lesions:
1. Absence of blood flow both within the nodule and
   around it
2. Blood flow around the nodule
3. Blood flow both within the nodule and around it

Some authors (Messina et al. 1996; Morozov 1997; Abdulhalimova et al. 1999) additionally describe an intranodular type of vascular pattern, where individual or multiple color signals are registered within the lesion.

Zubarev et al. (2000) suggest that the following three vascular patterns of thyroid nodules should be used in daily practice:

1. Perinodular: the blood flow is mainly in the periphery of the nodule
2. Mixed: vascularization occurs in the periphery of and within the nodule
3. Avascular: there is no sonographically discernible blood flow.

The thyroid nodules can also be divided into the following groups according to the blood flow intensity:

1. Hypervascular nodules show a peripheral rim and multiple arterial and venous vessels within (the sign of a “color crown”)
2. Nodules with a medium degree of vascularization have 5–6 color spots within the nodule
3. Hypovascular nodules show 2–3 color spots
4. Avascular nodules have no inner color spots and no peripheral rim

CDI has some disadvantages, such as table distortions of the Doppler spectrum (aliasing artifact), baseline noise, and dependence on the angle of the US beam.

**Power Doppler imaging** (PDI) permits images of smaller vessels with sharper contours to be obtained. This increases the diagnostic value of US (Fig. 2.8). PDI demonstrates a decreased dependence on the angle between the US beam and the blood flow, shows no aliasing artifact, and has a lower noise level. Therefore, PDI is three- to fivefold more sensitive than CDI (Lagalla et al. 1994, Adler et al. 1995; Spiezia et al. 1996). According to Zubarev (1997), PDI increases diagnostic sensitivity to thyroid pathology from 36 to 79% and specificity from 58 to 62% as compared with CDI.

PDI has some disadvantages, such as its high dependence on the motions of surrounding structures (leading to “motion artifacts”) and the staining of perivascular areas.

Fast computer processing of US images permits the **three-dimensional (3D) reconstruction** of the thyroid structure, lesions, the vascular tree, and surrounding tissues (Fig. 2.9). This option may be incorporated into ordinary US scanners as additional software. Data acquisition is achieved by a freehand scan with a usual 2D probe. Such cases demand subsequent computational processing of the data obtained. Some scanners can be equipped with special probes for mechanical 3D scanning. 3D imaging confers many advantages, such as the possibility of viewing planes that are usually inaccessible, and improved accuracy of volume estimation. It is useful for archiving US data in an objective form suitable for delayed analysis and digital transfer.

In comparison with 2D PDI, **3D reconstruction of vascular structures** (3D power Doppler imaging or 3DPD) enables more specific diagnoses of neoplasms based on the objective visual data for the structure.
Fig. 2.8 (a, b) Thyroid sonograms. PDI (a) hypervascular thyroid structure (b) avascular thyroid nodules

Fig. 2.9 Thyroid sonograms. 3D reconstruction
and the intensity of lesion vascularity, and the spatial relationships of different vascular structures of the neck (Fig. 2.10). According to Markova (2004), 3DPD is helpful when assessing the type of lesion vascularity, and it increases the sensitivity of US for the detection and categorization of thyroid lesions from 46 to 80%, and the specificity from 72 to 84%.

4D (real-time 3D) has reportedly been utilized to examine the thyroid gland. The 4D mode is a 3D scanning in real time using special US probes and high-class equipment. 3D image acquisition and reconstruction are performed quickly enough to allow real-time 3D visualization. 4D allows the spatial features of the thyroid to be defined more precisely, with a smaller dependence on noise artifacts. This is especially valuable for thyroid lesions.

In pulsed-wave (PW) Doppler, a curve resulting from the Doppler shift is produced via computer processing. This permits the analysis of the velocity and spectral parameters of the blood flow as well as the calculation of some indices (Fig. 2.11).

Markova (2001) suggests the following normal values for various blood flow parameters: the average peak systolic velocity (PSV) in the upper thyroid artery (UTA) is 16.8±0.94 cm/s; in the inferior thyroid artery (ITA) it is 15.8±0.77 cm/s; the end-diastolic velocity (EDV) in UTA is 7±1.2 cm/s; in the ITA it is 6.36±0.29 cm/s; the resistance index (RI) in UTA is 0.56±0.01; in the ITA it is 0.58±0.01. Struchkova (2003) also suggests nominal blood flow data for all thyroid arteries: PSV is 10.4–28.1 cm/s; EDV is 3.1–9.6 cm/s; RI is 0.5–0.75; and the pulsatility index (PI) is 0.7–1.2. RI and PI have been reported to be the most informative.

PW Doppler can confirm an increase in blood flow within the nodule as compared to that in the surrounding parenchyma (much more rarely, the vascularity is
Blood flow data within one nodule may vary substantially, which complicates the interpretation. The vascularity of a nodule was shown to be defined by both its morphological structure and its size. The bigger the nodule, the greater the observed increase in blood flow in one or several vessels.

Argalia et al. (1995) consider that PSV and RI are important for the differential diagnosis of thyroid nodules, and that they help to define the nodules that should be subjected to biopsy. According to Pinsky et al. (1999), blood flow data are undoubtedly higher in the vessels of the lobe that contains a tumor in comparison with the other lobe and the norm. Thus, PW Doppler allows thyroid tumors to be classified without separating them into benign or malignant. According to Kharchenko et al. (1994), malignant tumors are characterized by decreased PSV (39 ± 11 cm/s on the average) compared to those in adenomas.

Delorme et al. (1995) indicate that PW Doppler is subjective in assessments of blood flow changes. This may influence the examination and result in diagnostic errors. Our own research revealed no regularity in blood flow parameters. PW Doppler data in thyroid nodules show a wide dispersion and do not carry significant additional information. This precludes PW Doppler from being used for the differential diagnosis of thyroid nodules, although it may be used as an accessory feature.

Panoramic scan is an option that permits an extended field of view, thus simplifying the visualization and measurement of long structures. This helps when attempting to assess the precise dimensions of the thyroid and to calculate the volume of the lobes and the whole gland (Fig. 2.12).

The sensitivity of CDI and PDI can be significantly increased by intravenously introducing ultrasound contrast agents in a manner similar to contrast enhancement for CT and MRI. Lacocita et al. (2005) report that contrast-enhanced ultrasound (CEUS) is valuable for the diagnosis of thyroid diseases. They used SonoVue as a contrast medium for thyroid nodules. Nikolaeva et al. (2000) and Argalia et al. (2002) note the improvement in visualization of nodules of 0.5–1 cm in size with the use of Levovist. Fukunari et al. (2000) used Levovist to monitor the changes in thyroid nodules after PEI (858 observations).

Ultrasound elastography refers to a number of techniques that assist in the assessment of tissue softness (Fig. 2.13). The examined tissue is periodically exposed to pressure in order to create some form of displacement. The response is measured and processed to form an image. The tissue softness is color coded and observed superimposed on a grayscale image. Different colors correspond to different tissue elasticities. The best application of this modality is for the investigation of stiff tumors in soft tissues. It may be useful for both detection and categorization purposes. Moreover, it allows malignant tumor invasion to be defined more precisely, and small cancers to be...
2.1 Ultrasound Modalities

Fig. 2.12  Thyroid sonogram. Panoramic scan. (a) Transverse scan. (b) Longitudinal scan

Fig. 2.13  Thyroid sonogram. US elastography

In multislice viewing, a 3D US image is converted into a series of consecutive sections corresponding to intervals of 0.5–5 mm in any plane, similar to CT representations. This aids in the analysis of thyroid images, and makes it objective.

Enhancements to traditional procedures and advances in new technologies are leading to continual improvements in the accuracy and value of diagnostic ultrasound.

2.2 Technology Used in Ultrasound Examinations of the Thyroid Gland

Special preparation of the patient for thyroid US is not required. The patient is positioned supine, with the head thrown back and a bolster under the shoulders (Fig. 2.14). Seriously ill patients may sometimes be examined in a sitting position with the head thrown back.

Thyroid US is performed using a linear probe with a frequency of 5–17 MHz (most often 7.5–12 MHz). A 3.5–5 MHz convex probe is sometimes more convenient for measurements of large thyroids. A sector probe with a frequency of 2.5–5 MHz may be required for the substernal thyroid.

An outline of an US examination is provided below:

(a) The thyroid as a whole
- Location (typical, dystopia, ectopia)
- Dimensions and volume (also in comparison with the norm)
- Margins (regular/irregular, accurate/indistinct)
- Shape (typical; congenital anomalies: lobed constitution, aplasia, hypoplasia; goiter)
- Echodensity (normal, increase, decrease)
- Echostructure (homogeneous, heterogeneous)
- Blood vessels of the thyroid parenchyma (intensity, symmetry)

(b) Thyroid abnormalities
- Character of changes (diffuse, focal, mixed)
- Location (in lobes and segments)
- Number of lesions
- Contours (sharpness)
- Borders (smoothness)
- Dimensions (in three mutually perpendicular planes)
- Echodensity, echostructure of focal lesions
- Vascularity

(c) Mutual relations of the thyroid with the surrounding structures

(d) The status of regional lymph nodes

The US probe is positioned on the front surface of the neck and moved from the breastbone to the hyoid bone. The probe should produce minimal compression in order to avoid shape distortion of the thyroid gland.

The location and the parts of the thyroid are defined by measuring its dimensions and calculating its volume. At least five scanning planes should be evaluated to assess the dimensions of the thyroid: transverse, longitudinal, and oblique for the right and the left lobes (Fig. 2.15).

Thyroid size assessment is based on the linear dimensions and the volumes of the lobes. It is important to measure the linear dimensions only in the transverse or longitudinal sections of the thyroid lobes that show the

Fig. 2.14 (a, b) Thyroid US. The position of the patient (a) transverse thyroid scan, (b) longitudinal thyroid scan
maximum value (Fig. 2.16). When choosing the cross-section, it is necessary to follow the anatomical transverse plane and position the probe perpendicular to the skin with no angle. The longitudinal lobe dimension (the length or height of the lobes) is the largest size of the lobe. It is actually obtained in the plane that deviates from the anatomical longitudinal plane of the neck. The optimal position of the probe is close to parallel with the inner edge of the sternomastoid. Since the length of the lobe usually exceeds the length of a linear probe, it is preferable to measure it with a convex probe adjusted to the maximum possible frequency. The time expended and the reliability of this method of measurement are comparable with panoramic reconstruction of the image.
Fig. 2.16 (a1–a5, b1–b5) Thyroid US. Measurements of the widths, the depths, and the lengths of thyroid lobes, as well as the thickness of the isthmus (a1–b1) the depth and the width of the right lobe, (a2–b2) the depth and the width of the left lobe, (a3–b3) the isthmus (a4–b4) measurement of the lobe length in one view (a5–b5) combination of two measurements to calculate the lobe length.
The normal US dimensions of an adult thyroid can vary. A thyroid lobe is about 13–18 mm wide, 16–18 mm deep, and 45–60 mm long, while the isthmus is 2–6 mm deep; the thyroid has a volume of 7.7–22.6 cm³ in men and 4.55–19.32 cm³ in women (Ilyin 1995). The literature does not report any significant difference in US dimensions between the right and left thyroid lobes. Hence, separately defined linear parameters are of no value. It is important to note that the size of the organ is characterized only by the total volume of the glandular tissue.

The volume of a thyroid lobe is calculated by the formula \( A \times B \times C \times 0.479 \), where \( A \) is the length, \( B \) is the width, and \( C \) is the thickness (depth) of the lobe, while 0.479 (0.524) is the correction factor for the ellipsoidal shape of the lobe (Brunn et al. 1981).

The total thyroid volume encompasses the volumes of the right and left lobes. The volume of the isthmus (if thinner than 10 mm) is omitted.

The volume of a normal thyroid in both adults and children is still the source of debate (see Chap. 3). The World Health Organization suggests a normal volume in men of 7.7–25 cm³ and in women of 4.4–18 cm³. The calculated thyroid volume in adults can be compared with recommended standards that depend on age, height, weight, and body surface area (Parshin 1994; Ilyin 1995).

The optimal volume of the thyroid and criteria for its enlargement are currently being studied. No unified classification of thyroid enlargement based on sonographic data is being utilized yet. The classifications available are not accepted by the professional societies for general use.
They anchor the US data to the degree of enlargement of the thyroid gland based on imperfect palpation and visual assessment (for example the 1994 WHO scale).

At the same time, only one aspect is important in most cases: whether the patient’s thyroid volume differs from the norm. Many authors suggest that presenting the degree of deviation as a percentage may be of benefit for the dynamic assessment of changes in thyroid volume during treatment.

### 2.3 Basic Mistakes in Thyroid Ultrasound

Factors that result in inaccurate US assessment of the status of the thyroid gland can be divided into the following groups:

1. **Objective**
   - Anatomical, physiological, and constitutional features of the patient leading to a decrease in visualization
   - Limitations of the equipment (the quality of the scanner, the characteristics of the probes, etc.)

2. **Subjective**
   - Insufficient US specialist experience
   - Faulty thyroid US technique

High intra- and interobserver variations in thyroid sonography are largely due to the quality of the equipment and the skill level of the operator. According to Bataeva et al. (2006), an expert fails to reproduce the results of the 2D thyroid volume calculation in 8.7% of cases; assessments performed by different experts differ by 12.8%. Measurements taken in 3D methods of surface reconstruction and segmentation show differences of 4 and 4.8%, respectively. One disadvantage of using US in some cases is a low detection efficacy for thyroid dystopia and ectopia. Retrosternal location of the thyroid below the tracheal bifurcation significantly limits the possibility of ultrasonography (see Chap. 10).

High-resolution US equipment allows several thyroid pathologies to be detected, which used to be considered the norm. For example, medium- and large-sized cellular patterns (multiple fine hypo- and anechoic lesions 2–4 mm in size) that were interpreted as the norm years ago are now considered abnormal. This kind of sonogram is widely seen for people living in iodine-deficient regions, and precedes a diffuse endemic goiter. It corresponds to colloid cystic change with dilatation of the follicles due to extra colloid accumulation. The hyperchoic points within such follicles are a consequence of the dense consistency of the colloid. Another type of small change is inflammatory foci during the initial stages of autoimmune thyroid disease (AITD). These relate to a tendency for a decrease in echodensity and a slight heterogeneity of thyroid tissue resulting from small foci of lymphoplasmacytic infiltration and edema.

Alternatively, there are hyperdiagnostic cases when normal thyroid structures are interpreted as nodules. This especially concerns structures behind the left lobe or along the posterior margin of the inferior compartment of the right lobe. This type of error can be caused by the proximity of the esophagus, which may be reported as a nodule if imaged only with a transverse scan. The normal vascular pattern of the inferior thyroid artery (ITA) can result in hyperdiagnostics of thyroid nodules. In some cases, the ITA trunk does not split into fine branches when entering the inferior pole of the thyroid lobe. It may be traced within the lobe, where it borders a roundish region of healthy tissue that imitates a nodule.

Lymph nodes adjacent to the isthmus frequently complicate the diagnosis. The enlarged lymph nodes present in AITD are often located close to the upper or inferior part of the isthmus and can be interpreted as thyroid nodules. Differential diagnosis may benefit from the use of the highest probe frequency, as it permits a more detailed image. It is necessary to pay attention to specific features of lymph nodes, such as form, echosstructure with differentiation of the cortical and central parts, and type of blood flow. The opposite type of error, where nodules of the thyroid isthmus are interpreted as neck lymph nodes, are quite rare.

Differential diagnosis of thyroid nodules and inflammatory foci in AITD is rather difficult (see Chap. 5). In this case, it is necessary to consider the sharpness and regularity of the contours of the lesion, its form, and the blood flow pattern. Nevertheless, the correct diagnosis sometimes requires long observations or the use of other diagnostic modalities.

Rare neck pathologies may be misdiagnosed due to insufficient experience of the US operator. In many cases, adenomas and hyperplasia of the parathyroid
glands are interpreted as thyroid nodules. This pathology is relatively rare, so general practitioners or US specialists who do not practice at a specialized endocrinology center are not aware of it and have little experience in its diagnosis (see Chap. 11). The same can be said about the diverticulum of the esophagus (see Chap. 12). In most cases, sonographers have a mental image of “typical” thyroid nodules based on their own experience, so any lesion differing from this typical image should be interpreted with extra caution.

Special opportunities for differential diagnosis are provided by auxiliary methods, such as turning the patient’s head, compression of the neck tissues with fingers, swallowing, and others.

Note that modern equipment permits accurate US visualization of solid thyroid nodules larger than 2–3 mm in size. The presence of smaller lesions is preferably reported without the term “nodule.” Drawing a conclusion about the lesion is expedient only when it can be clearly visualized in at least two mutually perpendicular scans.

It is very important to adhere to the correct examination technique. Thyroid structure including small parts can be estimated only with linear probes at frequencies of 7.5 MHz and higher. Convex abdominal probes may be used to measure the lengths of thyroid lobes or for large thyroids. The use of a convex probe alone for thyroid examination results in multiple severe errors and discredits the field of sonography.

Rational timescales for thyroid US are as follows:

- Normal thyroid: once every two years for preventive purposes
- Benign diffuse or nodular thyroid pathology: 1–2 times a year to monitor the dynamics of the disease
- Inoperable thyroid malignancy: once every two months to define the stage.
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