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# across Human Cell Types **Nuclear DNA Content Varies with Cell Size**

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volume in other species, and interspecific relationships between diploid genome size and cell volume. Thus, we speculate that the quantity of nuclear DNA content in somatic cells of humans is perhaps best viewed as a distribution of values that reflects cell size distributions, quantity of nuclear DNA across 19 different human cell types increases with cell volume among cells have been frequently observed. Using published data, here we describe how the size is known to vary by several orders of magnitude, and differences in nuclear DNA content organisms that affects countless aspects of their structure and function. Within humans, cell rather than as a single, immutable quantity. This observed increase is similar to intraspecific relationships between DNA content and cell Variation in the size of cells, and the DNA they contain, is a basic feature of multicellular

or the rate at which different cells grow, divide, number of different cell types in an organism, tute multicellular organisms. For example, the the most basic features of the cells that constilecular level to the whole organism level. butes of multicellular organisms, from the mofects virtually all structural and functional attriand Skotheim 2015). Cell size, in particular, afcells that constitute an organism (see Amodeo understanding of the size and abundance of 2015). But perhaps most important, we lack an and die, remain poorly understood (see Niklas Yet, we still lack an understanding of some of ically increased over the past several decades. cells of which they are comprised, has dramatur understanding of the complexity of multicellular organisms, and the diverse

One key feature of organisms that may vary with cell size is the amount of nuclear DNA

Across species, genome size has long been known to correlate positively with cell and nuclear volume (Price et al. 1973; Szarski 1976; Olmo 1983). But within species, too, the nuclear DNA content of somatic cells has been shown in a few instances to increase with cell size in species such as *Daphnia* (Beaton and Hebert 1989) and *Arabidopsis* (Jovtchev et al. 2006). Such increases in nuclear DNA content can have important consequences for cell function, in general, and gene expression, in particular (Hancock et al. 2008; Lee et al. 2009; De Veylder et al. 2011; Marguerat and Bähler 2012).

In the case of humans, substantial differences in DNA content have been observed in many human cell types. Indeed, since Watson and Crick described the structure of DNA, studies of healthy human tissues have reported the presence of polyploid cells (Winkelmann et al.

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thology. A more synthetic view of differences in tions from the diploid quantity of DNA in hu-DNA constancy (Mirsky and Ris 1949). Deviapairs) based on the long-standing principle of characteristic quantity of DNA (  $\sim$  7 billion base somatic cells in the human body hold the same tle to change the traditional view that all healthy al. 1987). Still, these observations have done litstable, fully differentiated cells (Winkelmann et little in common, except that they are generally may provide some clarity on these and other nuclear DNA content across human cell types exceptional, tissue-specific, or indicative of pamans, like other animals, are still often viewed as which this has been observed appear to have 1987; Biesterfeld et al. 1994). The cell types in

ciated. However, as we later discuss, the mechain several broad taxonomic groups. These analsults with previously reported relationships benisms underlying these patterns remain in non in human cells than was previously appre-DNA content is a more ubiquitous phenomeyses suggest that systematic variation in nuclear genome size and cell size observed across species results with the relationships between diploid four other species. Finally, we compare these tween nuclear DNA content and cell size within lated with cell size. We then compare these recell types, and whether such variation is correclear DNA content varies across diverse human lished data to examine the extent to which nu-In this review, we compile and analyze pub-

# THE RELATIONSHIP OF NUCLEAR DNA CONTENT TO CELL SIZE IN HUMANS

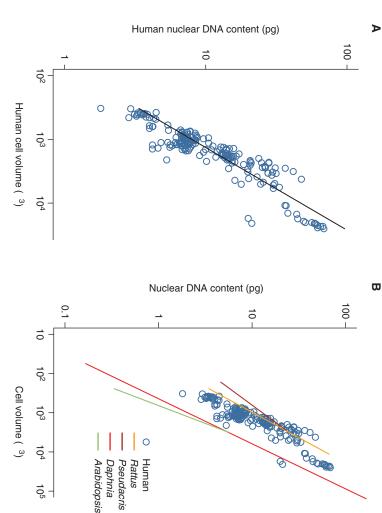
#### Methodology

Our analysis for this work used published data from healthy human cell populations representing 19 different cell types, as designated in the original studies (data provided in Table 1). In the original studies, DNA content was estimated using the Feulgen staining method, and the size of cells or cell nuclei were directly measured. Feulgen staining (Feulgen and Rosenbeck 1942) has been the most widely used method

terial is then measured to quantify the relative are converted from deoxyribose in DNA after by staining DNA owing to the reaction of Schiff among the cells we consider. The method works orders of magnitude variation in DNA content which for our analyses is negligible given the ically <5% using this method (Gregory 2005), terfeld et al. 2011). Measurement errors are typod for making quantitative measurements of DNA content of cells. The light absorbance of the stained genetic maor pseudo-Schiff reagents with aldehydes, which DNA content (Chieco and Derenzini 1999; Biesand is still generally considered a reliable methfor estimating DNA content for several decades, HCl hydrolysis (Chieco and Derenzini 1999).

In cases in which the relative content of DNA was originally expressed in "arbitrary units" in the original studies, we converted these measures to pg DNA given that the Feulgen staining method results in a linear relationship between light absorbance and DNA content (Biesterfeld et al. 2011). In each case, a specific conversion factor was used for each cell type based on the observed relationship between light absorbance and DNA content in the particular study (see Table 1). This was done to avoid any bias associated with differences in DNA compaction level or uptake of stain across cell types. The conversions used for each cell type are given in Table 1.

cell volumes were not directly reported, so we cell populations. In Table 1, we give all values for sent means of DNA content and cell size from values and used these in the analyses. So, all only the population means of cell size and of methodology and data) (Swanson et al. 1991) human cell types (see Table 1 for further details tween nuclear volume and cell volume across clear size based on a published relationship beestimated these volumes from measures of nuindividual cell values. In the majority of cases, inal sources, both population mean values and cell size and DNA content reported in the origpoints shown in the figures for humans reprevidual cell measurements, we calculated mean consistent, for those studies that reported indi-DNA content were reported. Thus, to remain Finally, in many studies considered here



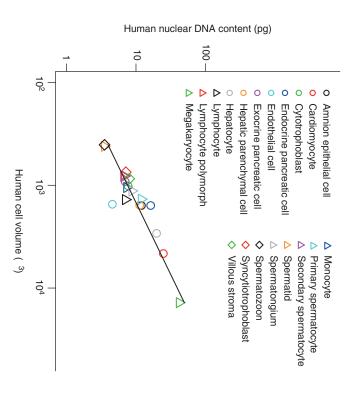
cell size in humans (blue) in comparison to previously reported relationships in the rat (Rattus norvegicus,  $\log_{10}$ human cell populations from 19 cell types ( $\log_{10} [DNA]$ ) =  $-1.3 + 0.74 \log_{10} [cell volume]$ ,  $r^2 = 0.81$ ; data in Table 1). The characteristic diploid human cell contains 7 pg DNA. (B) Relationship between DNA content and Figure 1. Mean DNA content for cell populations, (A) Mean nuclear DNA content versus cell volume for healthy regression. The human data shown here are the same as those shown in  $\it A$  $0.57\log_{10}$  [cell volume] [Bachmann et al. 1966]), crustacean ( $\it Daphnia~pulex, \log_{10}$  [DNA] -1.3 + 0.77 log<sub>10</sub> [cell volume] [Heizer 1955]), frog (Pseudacris obscura, log<sub>10</sub> [DNA] [Jovtchev et al. 2006]). Lines were fitted to data for nonhumans [Beaton and Hebert 1989]), and plant (Arabidopsis thalania, log10

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dom slope and intercept, respectively), and E nuclear DNA content (pg) and cell volume (µ<sup>3</sup> normality of residuals using quantile-quantile lated with cell volume within cell types, we used To determine whether DNA content was correcells from multiple cell types in a single analysis. in-group autocorrelation, allowing us to include 2000). This statistical model accounts for withrepresents residual errors (Pinheiro and Bates and D represent random effects of cell type (ranusing a linear mixed model of the form log<sub>10</sub> + E, where a and b are regression coefficients, C We evaluated the relationship between mean  $a + b \log_{10}$  (cell volume) + +D

#### Results

sure of nucleus size. Without converting to cell sperm to polyploid megakaryocytes. For 17 of (Fig. 1A) (mean  $\log_{10}$   $\log_{10}$  [cell volume],  $r^2$ atically with cell size among cell types, such that to 68.6 pg across cell types ranging from haploid DNA content for cell populations varies from 1.8 amount of DNA in human cells varies system-Our analyses of these data indicate that the  $1 \times 10^{-9}$ ). Figure 1A shows that the mean cells contain substantially  $= 0.82, F_{1, 175} = 36.6,$ more DNA



populations of a given cell type. Figure 2. Relationship between mean nuclear DNA content and mean cell size in diploid and polyploid human cells by cell type for data shown in Figure 1. Each point represents mean cell size and DNA content from all cell

relationship between nuclear volume and DNA content among these cell types ( $\log_{10}$  [DNA, pg] = -0.25 + 0.54  $\log_{10}$  [nuclear volume,  $\mu^3$ ],  $P = 3.8 \times 10^{-10}$ ). For the two remaining cell types in which we had direct measures of cell diameter (cardiomyocytes and megakaryocytes), we also observed a significant positive relationship between DNA content and cell volume ( $\log_{10}$  [DNA] = -0.71 + 0.52  $\log_{10}$  [cell volume],  $P = 1.0 \times 10^{-10}$ ).

When DNA content and cell volume were averaged over all cells for each cell type, we observed a similar relationship between mean DNA content and cell volume (Fig. 2) (mean  $\log_{10}$  [DNA] =  $-1.3 + 0.72 \log_{10}$  [cell volume],  $r^2 = 0.82$ ,  $F_{1,17} = 86.8$ ,  $P = 4.3 \times 10^{-8}$ ). Note, however, that the range in DNA content of individual cells from these populations was even greater, as one would expect. At one extreme, the largest megakaryocytes contained 448 pg of DNA (see Royere et al. 1988 in Table 1). This indicates that these cells have likely undergone six genome duplication events from their diploid state.

Moreover, within cell types, the same correlations between nuclear DNA content and cell volume were observed in five of the six cases in which mean cell volume varied by a factor of 5 or more (megakaryocytes, hepatocytes, lymphocytes, hepatic parenchymal cells, and amnion epithelial cells). In these five cases, DNA content was positively correlated with cell volume (ordinary least-squares [OLS] regression, all  $P \leq 1.6 \times 10^{-9}$ ). In the case of cardiomyocytes, however, we did not observe a statistically significant correlation (P = 0.1).

# RELATIONSHIP OF NUCLEAR DNA CONTENT TO CELL SIZE: HUMANS COMPARED WITH OTHER SPECIES

#### Methodology

To determine whether the relationships between cell size and DNA content were similar in humans and in other species, we used data from the few studies available on intraspecific variation in cell size and nuclear DNA content

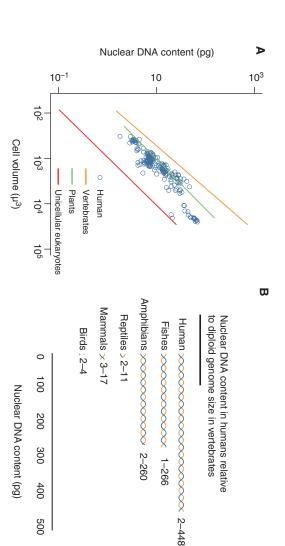
(Heizer 1955; Bachmann et al. 1966; Beaton and Hebert 1989; Jovtchev et al. 2006). Studies of nonhuman cells typically reported nuclear size instead of cell size. To convert nuclear volume to cell volume, we used the formula  $\log_{10}$  (cell volume) =  $0.88 + \log_{10}$  (nuclear volume), which we obtained by fitting a linear function to  $\log_{10}$ -transformed cell and nuclear size measurements from vertebrate (Olmo 1983) and plant (Price et al. 1973) cells from a diverse assortment of species using ordinary least squares.

#### results

The relationship between nuclear DNA content and cell volume in humans is similar to that previously reported for other species. DNA content is similarly correlated with cell volume in the rat *Rattus norvegicus*, the frog *Pseudacris obscura*, the crustacean *Daphnia pulex*, and the plant *Arabidopsis thalania* (Fig. 1B). These results indicate that concomitant changes in

cell volume and nuclear DNA content may occur across very different cell types and species, although this has only rarely been investigated.

given possible differences in methodology, posof species from these groups (e.g., unicellular and, in some cases, a fairly limited sampling nuclear volume and cell volume among taxa sible differences in the relationship between cautious in interpreting similarities and differdiploid genome size and cell volume in plants tent and cell volume in humans also appears eukaryotes). ences among these interspecific relationships all vertebrates (Fig. 3B). However, one must be rable to the range of diploid genome sizes across nuclear DNA content in human cells is compa-(Price et al. 1973), vertebrate animals (Olmo similar to the relationships observed between (Shuter et al. 1983). Interestingly, the range of 1983), and unicellular eukaryotes (Fig. 3A) The relationship between nuclear DNA con-



using ordinary least-squares regression. The range of unicellular eukaryote cell and genome sizes is truncated to previously reported relationships for diploid cells of vertebrates ( $log_{10}$  [DNA] = genome sizes within vertebrate groups. DNA content was rounded to the nearest whole number Figure 1A. (B) Range of nuclear DNA content in individual human cells in comparison to ranges of diploid clearly show data for other groups. Note that the human data shown in this figure are the same as those shown in tween nuclear DNA content and cell size in diploid and polyploid human cells (blue) in comparison to Figure 3. Relationship between nuclear DNA content and cell volume in humans. (A) Relationship beunicellular eukaryotes ( $log_{10}$  [DNA] =  $-1.75 + 0.912 \log_{10}$ [cell volume]). Lines were fitted to previously reported relationships  $2.81 + 0.945 \log_{10}$  [cell volume]), and angiosperms  $-1.91 + 1.07 \log_{10}$  [cell



#### CONCLUSIONS

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size, nuclear DNA content, and gene expression in other cell types not considered here (e.g., greater levels of protein synthesis for maintewithin and across cell types is only beginning to and functional. The relationship between cell muscle cells), multinucleated cells are common cell types (e.g., megakaryocytes), polyploid nuextra gene copies as an insurance policy against gued that perhaps this process serves to provide has long been viewed in plants. Some have armay be a common and/or functional trait in handful of model organisms. Endoreplication cell types of different sizes is restricted to only a have measured the amount of DNA in different stancy across somatic cells in humans and other to cell size. It remains to be seen whether the questions regarding the prevalence of endo-Our analysis of previously published data raises was previously thought (Parfrey et al. 2008). clearly genomes are much more dynamic than be explored (Marguerat and Bähler 2012), but clei are considered normal and functional. And Veylder et al. 2011). Certainly, in some human nance and production (Lee et al. 2009; De DNA damage, or that larger cells simply require Vinogradov 2011; De Veylder et al. 2011), as it (Lee et al. 2009; Ullah et al. 2009; Anatskaya and the somatic cells of humans and other animals The number of animal species for which we species has not been adequately investigated body. Our common assumption of DNA conhundreds of different cell types in the human relationship presented here extends across the replication in human cells and its relationship

We have moved beyond the view that genomes are immutable blueprints. Most of the cells shown in Figure 1 are presumably postmitotic, but many show quantities of nuclear DNA well beyond the expected value of about 7 pg.

The notion that human cells may have quite different genetic constitutions depending on their size would have profound implications for our understanding of cell structure and function.

The data shown here suggest that perhaps the quantity of nuclear DNA content in human cells is best viewed as a distribution of values that reflects cell size distributions, rather than

as a single value. That is, most cells may be of a similar size and contain something close to the characteristic amount of nuclear DNA, but a relatively small number of cells may be large with high amounts of DNA.

duplication in these two cell types. combination of partial and complete genome simply based on error. Thus, the data suggest around values expected from endopolyploidcloser look at the data provides some clues data on the individual cells of a cell population most data are averages of cell populations. Howhere, it is simply not possible to generalize the typical diploid quantity. Of the cell types shown cells may occur by endoreplication of individual DNA content has increased as a result of some erably more variation than one would expect ization (e.g., 7, 14, 28, etc.), but there is consid-DNA content with cell size appear to cluster (see Table 1). In these data, the increases in (amnion epithelial cells and hepatocytes), a ever, for the two cell types for which we have extent to which this may be the case because increases in nuclear DNA content beyond the cation via endoreplication is responsible for any tion). Often, it is assumed that genome dupligenes or the entire genome (i.e., polyploidiza-Changes in the DNA content of somatic

and functional features (gene expression and and how such increases affect gene expression the two. the qualitatively similar relationships between ships across species may help point to a more within species to genome size-cell size relationhas yet to be resolved. It is our hope that our dress the question of whether genome size conpolyploidization) to the life span, replication structural features (genome size and cell size) suggest that it may be useful to relate relevant proaches may be insightful. In particular, we 2012). Again, here, broadly comparative apthat lead to increases in cellular DNA content much to learn about the various mechanisms general understanding. We were surprised by comparisons of DNA-cell size relationships This has long been debated in the literature, and trols cell size, or cell size controls genome size rate, and metabolic rate of cells. (Therman et al. 1983; Marguerat and Bähler However, the results shown here do not ad-They highlight the fact that we have

Table 1. Human nuclear DNA content and cell volume data

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Amnion epithelial cell (s)	3.0	40.7	c, a	1072.1	5.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.6	41.6	c, a	958.0	6.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.9	41.7	c, a	1048.5	6.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.0	42.4	c, a	1437.0	6.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.4	42.7	c, a	1227.5	6.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.5	43.0	c, a	895.3	6.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.5	43.0	c, a	915.8	6.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.7	44.0	c, a	968.9	6.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.8	44.6	c, a	1002.2	6.5	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.9	44.6	c, a	1036.5	6.5	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.2	44.7	c, a	1147.2	6.5	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.0	45.4	c, a	1437.0	6.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.6	45.5	c, a	936.7	6.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.9	45.6	c, a	1036.5	6.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.6	46.8	c, a	1298.5	6.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.0	47.2	c, a	1072.1	6.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.4	47.6	c, a	865.5	6.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.8	47.7	c, a	1002.2	6.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.8	47.7	c, a	1013.5	6.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.9	48.0	c, a	1405.1	7.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.7	48.7	c, a	1682.4	7.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.7	48.8	c, a	968.9	7.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.9	48.8	c, a	1036.5	7.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.1	48.9	c, a	1109.2	7.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.8	49.1	c, a	1373.6	7.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.6	49.3	c, a	936.7	7.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.8	49.4	c, a	1013.5	7.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.7	49.9	c, a	968.9	7.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.2	50.3	c, a	1503.2	7.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.9	50.5	c, a	1036.5	7.3	Klinger and Schwarzacher 1960

Table 1. Continued

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Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Amnion epithelial cell (s)	3.1	50.6	c, a	1134.2	7.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.4	50.7	c, a	1213.8	7.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.5	51.0	c, a	905.6	7.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.0	52.0	c, a	1437.0	7.5	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	2.7	52.2	c, a	968.9	7.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.2	52.4	c, a	1173.3	7.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.5	53.0	c, a	1255.3	7.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	1.8	53.0	c, a	660.8	7.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.4	53.2	c, a	1572.4	7.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.0	53.5	c, a	1072.1	7.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.9	53.8	c, a	1405.1	7.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.9	54.4	c, a	1389.4	7.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.5	54.9	c, a	1255.3	8.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.2	57.4	c, a	1173.3	8.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.9	58.2	c, a	1405.1	8.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.6	62.2	c, a	1313.2	9.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.8	64.4	c, a	1373.6	9.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.9	71.4	c, a	1389.4	10.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.8	80.8	c, a	1358.1	11.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.6	82.6	c, a	1313.2	12.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.3	85.7	c, a	1537.5	12.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	3.8	87.5	c, a	1358.1	12.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.6	89.1	c, a	1992.2	12.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.0	89.6	c, a	1420.9	13.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.3	90.1	c, a	1904.3	13.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.5	91.9	c, a	1608.4	13.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.0	92.7	c, a	1420.9	13.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.3	93.9	c, a	1537.5	13.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.6	94.0	c, a	1663.6	13.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.2	97.1	c, a	1520.3	14.1	Klinger and Schwarzacher 1960



Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Amnion epithelial cell (s)	5.8	97.7	c, a	2060.6	14.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.9	98.5	c, a	1760.2	14.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.5	98.7	c, a	1970.0	14.3	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.3	100.5	c, a	1537.5	14.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.8	101.0	c, a	2060.6	14.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.1	102.0	c, a	1820.4	14.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	6.0	102.2	c, a	2131.4	14.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.6	102.9	c, a	1645.1	14.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.7	104.5	c, a	2014.8	15.1	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.1	106.7	c, a	1820.4	15.5	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.3	106.8	c, a	1883.0	15.5	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.0	107.9	c, a	1780.0	15.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	4.9	109.1	c, a	1740.3	15.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	6.9	114.8	c, a	2439.8	16.6	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.5	115.7	c, a	1970.0	16.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	6.7	118.7	c, a	2358.9	17.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	7.2	120.2	c, a	2552.4	17.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.9	124.0	c, a	2107.4	18.0	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.7	126.7	c, a	2014.8	18.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	5.9	132.7	c, a	2107.4	19.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	7.0	142.5	c, a	2467.4	20.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	7.8	149.4	c, a	2761.6	21.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	7.6	156.3	c, a	2700.3	22.7	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	8.7	171.5	c, a	3056.5	24.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	7.8	185.4	c, a	2761.6	26.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	8.3	185.5	c, a	2921.7	26.9	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	8.9	187.9	c, a	3126.2	27.2	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	8.3	192.0	c, a	2921.7	27.8	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	7.3	209.8	c, a	2581.4	30.4	Klinger and Schwarzacher 1960
Amnion epithelial cell (s)	6.8	224.3	c, a	2412.5	32.5	Klinger and Schwarzacher 1960

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Table 1. Continued

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	Cell/nuclear size (original	DNA content (original	Size, content (original	Cell volume	DNA content	
Cell type	values)	values)	units)	$(\mu^3)$	(pg)	Sources
Cardiomyocyte (m)	17401.0	20.0	v, pg	5646.7	20.0	Vliegen et al. 1990
Cardiomyocyte (m)	20794.0	21.2	v, pg	6339.9	21.2	Vliegen et al. 1990
Cardiomyocyte (m)	43.8	3.6	cc, p	1463.2	12.6	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	60.7	4.1	cc, p	2013.5	14.4	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	55.5	4.5	cc, p	1844.6	15.8	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	135.5	4.6	cc, p	4403.8	16.1	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	156.4	5.1	cc, p	5064.6	17.9	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	127.0	5.8	cc, p	4132.1	20.2	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	65.1	5.9	cc, p	2154.9	20.7	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	68.1	5.9	cc, p	2252.3	20.7	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	77.3	5.9	cc, p	2546.1	20.7	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	71.1	6.0	cc, p	2346.7	21.0	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	119.2	6.6	cc, p	3885.0	23.1	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	60.3	6.8	cc, p	2000.2	23.8	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	84.1	7.2	cc, p	2765.2	25.2	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	103.7	7.9	cc, p	3392.5	27.6	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	104.0	7.9	cc, p	3402.4	27.7	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	112.1	8.4	cc, p	3658.9	29.4	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	99.8	8.5	cc, p	3266.2	29.8	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	161.5	8.6	cc, p	5224.0	30.1	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	152.5	8.8	cc, p	4939.6	30.8	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	105.1	9.0	cc, p	3436.8	31.5	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	175.4	9.5	cc, p	5660.7	33.3	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	97.6	11.9	cc, p	3198.5	41.7	Wohlschlaeger et al. 2010
Cardiomyocyte (m)	129.9	13.6	cc, p	4225.6	47.6	Wohlschlaeger et al. 2010
Cytotrophoblast (m)	133.0	1.9	cc, p	1061.5	6.7	Galton 1962
Cytotrophoblast (m)	109.0	2.1	cc, p	932.7	7.4	Galton 1962
Cytotrophoblast (m)	125.0	2.4	cc, p	1019.6	8.3	Galton 1962
Cytotrophoblast (m)	128.0	2.4	cc, p	1035.4	8.5	Galton 1962
Endocrine pancreatic cell (m)	98.4	2.0	cc, p	872.7	7.0	Ehrie and Swartz 1974

 $\stackrel{\rightharpoonup}{\rightarrow}$ 



Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Endocrine pancreatic cell (m)	116.5	2.0	cc, p	973.9	7.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	119.0	2.0	cc, p	987.5	7.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	205.4	4.0	cc, p	1408.0	14.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	217.8	4.0	cc, p	1462.7	14.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	221.8	4.0	cc, p	1480.1	14.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	435.5	8.0	cc, p	2294.9	28.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	447.3	8.0	cc, p	2335.1	28.0	Ehrie and Swartz 1974
Endocrine pancreatic cell (m)	453.1	8.0	cc, p	2354.8	28.0	Ehrie and Swartz 1974
Endothelial cell (m)	36.5	4.2	cc, pg	1225.4	4.2	Dorman et al. 1990
Endothelial cell (m)	42.4	4.2	cc, pg	1418.2	4.2	Dorman et al. 1990
Endothelial cell (m)	41.1	4.6	cc, pg	1375.8	4.6	Dorman et al. 1990
Endothelial cell (m)	63.5	5.3	cc, pg	2102.6	5.3	Dorman et al. 1990
Exocrine pancreatic cell (m)	100.1	2.0	cc, p	882.5	7.0	Ehrie and Swartz 1974
Exocrine pancreatic cell (m)	107.7	2.0	cc, p	925.5	7.0	Ehrie and Swartz 1974
Exocrine pancreatic cell (m)	112.8	2.0	cc, p	953.7	7.0	Ehrie and Swartz 1974
Hepatic parenchymal cell (m)	34.7	1.8	cc, a	1124.6	4.8	Swartz 1956
Hepatic parenchymal cell (m)	34.4	2.0	cc, a	1118.2	5.4	Swartz 1956
Hepatic parenchymal cell (m)	45.1	2.1	cc, a	1333.5	5.6	Swartz 1956
Hepatic parenchymal cell (m)	34.8	2.3	cc, a	1126.7	6.1	Swartz 1956
Hepatic parenchymal cell (m)	26.1	2.3	cc, a	934.6	6.2	Swartz 1956
Hepatic parenchymal cell (m)	38.9	2.3	cc, a	1211.3	6.2	Swartz 1956
Hepatic parenchymal cell (m)	37.9	2.3	cc, a	1191.0	6.3	Swartz 1956
Hepatic parenchymal cell (m)	36.6	2.4	cc, a	1164.2	6.5	Swartz 1956
Hepatic parenchymal cell (m)	33.9	2.5	cc, a	1107.7	6.6	Swartz 1956
Hepatic parenchymal cell (m)	33.2	2.5	cc, a	1092.8	6.6	Swartz 1956
Hepatic parenchymal cell (m)	32.9	2.5	cc, a	1086.3	6.7	Swartz 1956
Hepatic parenchymal cell (m)	41.7	2.5	cc, a	1267.3	6.7	Swartz 1956
Hepatic parenchymal cell (m)	27.6	2.5	cc, a	969.1	6.8	Swartz 1956
Hepatic parenchymal cell (m)	51.2	2.5	cc, a	1448.1	6.8	Swartz 1956
Hepatic parenchymal cell (m)	35.2	2.6	cc, a	1135.1	6.8	Swartz 1956

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Table 1. Continued

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Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Hepatic parenchymal cell (m)	41.8	2.6	cc, a	1269.2	6.9	Swartz 1956
Hepatic parenchymal cell (m)	29.5	2.6	cc, a	1012.0	7.0	Swartz 1956
Hepatic parenchymal cell (m)	29.4	2.6	cc, a	1009.7	7.1	Swartz 1956
Hepatic parenchymal cell (m)	40.4	2.6	cc, a	1241.5	7.1	Swartz 1956
Hepatic parenchymal cell (m)	47.4	2.7	cc, a	1377.3	7.2	Swartz 1956
Hepatic parenchymal cell (m)	39.9	2.7	cc, a	1231.4	7.2	Swartz 1956
Hepatic parenchymal cell (m)	32.5	2.7	cc, a	1077.7	7.2	Swartz 1956
Hepatic parenchymal cell (m)	55.3	2.7	cc, a	1522.5	7.2	Swartz 1956
Hepatic parenchymal cell (m)	30.6	2.7	cc, a	1036.3	7.4	Swartz 1956
Hepatic parenchymal cell (m)	36.9	2.8	cc, a	1170.5	7.4	Swartz 1956
Hepatic parenchymal cell (m)	22.3	2.8	cc, a	843.7	7.5	Swartz 1956
Hepatic parenchymal cell (m)	36.4	2.8	cc, a	1160.1	7.5	Swartz 1956
Hepatic parenchymal cell (m)	28.1	2.8	cc, a	980.5	7.5	Swartz 1956
Hepatic parenchymal cell (m)	33.9	2.8	cc, a	1107.7	7.5	Swartz 1956
Hepatic parenchymal cell (m)	54.5	2.8	cc, a	1508.1	7.5	Swartz 1956
Hepatic parenchymal cell (m)	38.6	2.9	cc, a	1205.2	7.7	Swartz 1956
Hepatic parenchymal cell (m)	49.3	2.9	cc, a	1413.0	7.7	Swartz 1956
Hepatic parenchymal cell (m)	36.1	2.9	cc, a	1153.9	7.7	Swartz 1956
Hepatic parenchymal cell (m)	44.6	2.9	cc, a	1323.9	7.8	Swartz 1956
Hepatic parenchymal cell (m)	33.3	2.9	cc, a	1094.9	7.9	Swartz 1956
Hepatic parenchymal cell (m)	31.4	3.1	cc, a	1053.9	8.4	Swartz 1956
Hepatic parenchymal cell (m)	36.0	3.2	cc, a	1151.8	8.5	Swartz 1956
Hepatic parenchymal cell (m)	71.4	3.4	cc, a	1797.6	9.1	Swartz 1956
Hepatic parenchymal cell (m)	98.7	3.8	cc, a	2218.6	10.1	Swartz 1956
Hepatic parenchymal cell (m)	78.1	4.4	cc, a	1905.5	11.8	Swartz 1956
Hepatic parenchymal cell (m)	44.5	4.8	cc, a	1321.9	12.8	Swartz 1956
Hepatic parenchymal cell (m)	65.8	4.8	cc, a	1704.6	12.9	Swartz 1956
Hepatic parenchymal cell (m)	71.6	4.9	cc, a	1800.8	13.1	Swartz 1956
Hepatic parenchymal cell (m)	65.0	5.0	cc, a	1691.1	13.3	Swartz 1956
Hepatic parenchymal cell (m)	69.0	5.1	cc, a	1758.1	13.7	Swartz 1956

Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Hepatic parenchymal cell (m)	64.1	5.2	cc, a	1675.9	13.9	Swartz 1956
Hepatic parenchymal cell (m)	72.9	5.3	cc, a	1822.0	14.1	Swartz 1956
Hepatic parenchymal cell (m)	70.7	5.3	cc, a	1786.1	14.1	Swartz 1956
Hepatic parenchymal cell (m)	91.7	5.3	cc, a	2115.0	14.2	Swartz 1956
Hepatic parenchymal cell (m)	87.0	5.4	cc, a	2043.9	14.5	Swartz 1956
Hepatic parenchymal cell (m)	78.4	5.4	cc, a	1910.2	14.5	Swartz 1956
Hepatic parenchymal cell (m)	147.2	5.4	cc, a	2876.9	14.5	Swartz 1956
Hepatic parenchymal cell (m)	70.3	5.5	cc, a	1779.5	14.7	Swartz 1956
Hepatic parenchymal cell (m)	81.6	5.6	cc, a	1960.6	15.0	Swartz 1956
Hepatic parenchymal cell (m)	72.9	5.7	cc, a	1822.0	15.2	Swartz 1956
Hepatic parenchymal cell (m)	73.6	6.1	cc, a	1833.4	16.4	Swartz 1956
Hepatic parenchymal cell (m)	65.3	6.4	cc, a	1696.2	17.2	Swartz 1956
Hepatic parenchymal cell (m)	177.5	7.4	cc, a	3249.1	19.9	Swartz 1956
Hepatic parenchymal cell (m)	106.7	9.3	cc, a	2333.9	24.9	Swartz 1956
Hepatic parenchymal cell (m)	129.4	9.8	cc, a	2645.7	26.2	Swartz 1956
Hepatic parenchymal cell (m)	141.9	9.8	cc, a	2809.1	26.4	Swartz 1956
Hepatic parenchymal cell (m)	136.8	9.9	cc, a	2743.1	26.5	Swartz 1956
Hepatic parenchymal cell (m)	142.1	11.6	cc, a	2811.7	31.1	Swartz 1956
Hepatic parenchymal cell (m)	144.7	13.6	cc, a	2845.0	36.6	Swartz 1956
Hepatocyte (s)	35.7	18.3	cc, a	1200.9	8.7	Meek and Harbison 1967
Hepatocyte (s)	46.1	20.7	cc, a	1537.3	9.9	Meek and Harbison 1967
Hepatocyte (s)	41.9	21.0	cc, a	1400.8	10.0	Meek and Harbison 1967
Hepatocyte (s)	59.3	22.4	cc, a	1965.6	10.7	Meek and Harbison 1967
Hepatocyte (s)	39.0	22.6	cc, a	1306.2	10.8	Meek and Harbison 1967
Hepatocyte (s)	42.2	23.6	cc, a	1411.4	11.3	Meek and Harbison 1967
Hepatocyte (s)	73.4	23.8	cc, a	2422.6	11.4	Meek and Harbison 1967
Hepatocyte (s)	64.7	24.4	cc, a	2142.5	11.6	Meek and Harbison 1967
Hepatocyte (s)	46.7	25.6	cc, a	1558.2	12.2	Meek and Harbison 1967
Hepatocyte (s)	68.0	25.8	cc, a	2246.3	12.3	Meek and Harbison 1967
Hepatocyte (s)	64.1	26.7	cc, a	2121.7	12.7	Meek and Harbison 1967

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Table 1. Continued

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Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Hepatocyte (s)	89.5	29.5	cc, a	2939.3	14.1	Meek and Harbison 1967
Hepatocyte (s)	54.1	30.6	cc, a	1798.8	14.6	Meek and Harbison 1967
Hepatocyte (s)	73.1	32.1	cc, a	2412.3	15.3	Meek and Harbison 1967
Hepatocyte (s)	89.5	32.5	cc, a	2939.3	15.5	Meek and Harbison 1967
Hepatocyte (s)	64.1	33.3	cc, a	2121.7	15.9	Meek and Harbison 1967
Hepatocyte (s)	73.1	34.4	cc, a	2412.3	16.4	Meek and Harbison 1967
Hepatocyte (s)	96.3	35.8	cc, a	3155.6	17.1	Meek and Harbison 1967
Hepatocyte (s)	53.8	35.9	cc, a	1788.3	17.1	Meek and Harbison 1967
Hepatocyte (s)	68.0	38.3	cc, a	2246.3	18.2	Meek and Harbison 1967
Hepatocyte (s)	84.1	39.1	cc, a	2763.9	18.6	Meek and Harbison 196
Hepatocyte (s)	78.9	39.7	cc, a	2598.6	18.9	Meek and Harbison 196
Hepatocyte (s)	63.8	39.9	cc, a	2111.3	19.0	Meek and Harbison 196
Hepatocyte (s)	68.3	40.3	cc, a	2256.7	19.2	Meek and Harbison 196
Hepatocyte (s)	72.8	40.9	cc, a	2401.9	19.5	Meek and Harbison 196
Hepatocyte (s)	113.4	41.9	cc, a	3699.8	19.9	Meek and Harbison 196
Hepatocyte (s)	126.2	43.3	cc, a	4109.2	20.6	Meek and Harbison 196
Hepatocyte (s)	90.2	43.7	cc, a	2959.9	20.8	Meek and Harbison 1967
Hepatocyte (s)	106.9	44.1	cc, a	3494.7	21.0	Meek and Harbison 1967
Hepatocyte (s)	101.1	45.7	cc, a	3309.8	21.8	Meek and Harbison 1967
Hepatocyte (s)	96.0	47.0	cc, a	3145.3	22.4	Meek and Harbison 1967
Hepatocyte (s)	90.2	47.6	cc, a	2959.9	22.7	Meek and Harbison 1967
Hepatocyte (s)	140.1	48.0	cc, a	4548.1	22.9	Meek and Harbison 1967
Hepatocyte (s)	95.3	48.7	cc, a	3124.7	23.2	Meek and Harbison 1967
Hepatocyte (s)	107.2	49.1	cc, a	3504.9	23.4	Meek and Harbison 196
Hepatocyte (s)	120.8	49.8	cc, a	3935.3	23.7	Meek and Harbison 1967
Hepatocyte (s)	90.2	50.3	cc, a	2959.9	23.9	Meek and Harbison 1967
Hepatocyte (s)	140.1	50.9	cc, a	4548.1	24.3	Meek and Harbison 1967
Hepatocyte (s)	101.4	52.7	cc, a	3320.1	25.1	Meek and Harbison 1967
Hepatocyte (s)	113.7	53.4	cc, a	3710.1	25.4	Meek and Harbison 1967
Hepatocyte (s)	156.2	56.6	cc, a	5057.1	27.0	Meek and Harbison 1967

Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Hepatocyte (s)	113.0	58.7	cc, a	3689.5	27.9	Meek and Harbison 1967
Hepatocyte (s)	120.8	63.0	cc, a	3935.3	30.0	Meek and Harbison 1967
Hepatocyte (s)	140.7	79.6	cc, a	4568.4	37.9	Meek and Harbison 1967
Hepatocyte (s)	170.4	91.6	cc, a	5503.9	43.6	Meek and Harbison 1967
Hepatocyte (s)	177.5	97.9	cc, a	5726.9	46.6	Meek and Harbison 1967
Lymphocyte (m)	17.9	4.0	cc, pg	611.8	4.0	Dorman et al. 1990
Lymphocyte (m)	19.8	4.0	cc, pg	675.0	4.0	Dorman et al. 1990
Lymphocyte (m)	18.5	4.2	cc, pg	631.8	4.2	Dorman et al. 1990
Lymphocyte (m)	37.6	5.8	cc, pg	1261.4	5.8	Dorman et al. 1990
Lymphocyte (s)	18.1	17.2	cc, a	618.7	6.3	Bedi and Goldstein 1976
Lymphocyte (s)	19.9	17.4	cc, a	679.4	6.3	Bedi and Goldstein 1976
Lymphocyte (s)	18.6	17.7	cc, a	633.9	6.4	Bedi and Goldstein 1976
Lymphocyte (s)	23.5	18.5	cc, a	796.5	6.7	Bedi and Goldstein 1976
Lymphocyte (s)	21.2	18.6	cc, a	721.0	6.7	Bedi and Goldstein 1976
Lymphocyte (s)	22.8	18.8	cc, a	773.8	6.8	Bedi and Goldstein 1976
Lymphocyte (s)	21.4	18.9	cc, a	728.5	6.9	Bedi and Goldstein 1976
Lymphocyte (s)	22.9	18.9	cc, a	777.6	6.9	Bedi and Goldstein 1976
Lymphocyte (s)	24.9	19.0	cc, a	845.4	6.9	Bedi and Goldstein 1976
Lymphocyte (s)	22.4	19.0	cc, a	762.5	6.9	Bedi and Goldstein 1976
Lymphocyte (s)	24.3	19.0	cc, a	822.9	6.9	Bedi and Goldstein 1976
Lymphocyte (s)	23.0	19.1	cc, a	781.4	6.9	Bedi and Goldstein 1976
Lymphocyte (s)	22.8	19.3	cc, a	773.8	7.0	Bedi and Goldstein 1976
Lymphocyte (s)	23.8	19.7	cc, a	807.8	7.1	Bedi and Goldstein 1976
Lymphocyte (s)	28.0	19.9	cc, a	946.9	7.2	Bedi and Goldstein 1976
Lymphocyte (s)	7.4	2.7	n, a	1437.3	4.1	Petrakis 1953
Lymphocyte (s)	5.5	3.0	n, a	793.2	4.5	Petrakis 1953
Lymphocyte (s)	7.0	3.0	n, a	1293.6	4.6	Petrakis 1953
Lymphocyte (s)	5.2	3.0	n, a	712.0	4.6	Petrakis 1953
Lymphocyte (s)	7.7	3.0	n, a	1562.6	4.6	Petrakis 1953

DNA—Cell Size Relationship in Humans

#### Cold Spring Harbor Perspectives in Biology www.cshperspectives.org

Table 1. Continued

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	Cell/nuclear size (original	DNA content (original	Size, content (original	Cell volume	DNA content	
Cell type	values)	values)	units)	$(\mu^3)$	(pg)	Sources
Lymphocyte (s)	7.5	3.1	n, a	1462.0	4.7	Petrakis 1953
Lymphocyte (s)	7.0	3.1	n, a	1278.2	4.8	Petrakis 1953
Lymphocyte (s)	7.1	3.1	n, a	1340.8	4.8	Petrakis 1953
Lymphocyte (s)	5.6	3.2	n, a	824.1	4.8	Petrakis 1953
Lymphocyte (s)	7.0	3.3	n, a	1293.6	5.0	Petrakis 1953
Lymphocyte (s)	8.0	3.3	n, a	1666.4	5.0	Petrakis 1953
Lymphocyte (s)	7.5	3.3	n, a	1462.0	5.0	Petrakis 1953
Lymphocyte (s)	8.2	3.3	n, a	1746.7	5.0	Petrakis 1953
Lymphocyte (s)	7.8	3.4	n, a	1597.1	5.2	Petrakis 1953
Lymphocyte (s)	6.5	3.4	n, a	1118.1	5.2	Petrakis 1953
Lymphocyte (s)	6.9	3.5	n, a	1239.8	5.3	Petrakis 1953
Lymphocyte (s)	7.7	3.5	n, a	1562.6	5.3	Petrakis 1953
Lymphocyte (s)	8.0	3.5	n, a	1666.4	5.3	Petrakis 1953
Lymphocyte (s)	7.5	3.5	n, a	1495.0	5.3	Petrakis 1953
Lymphocyte (s)	5.0	3.5	n, a	680.8	5.4	Petrakis 1953
Lymphocyte (s)	6.0	3.5	n, a	953.5	5.4	Petrakis 1953
Lymphocyte (s)	7.4	3.5	n, a	1453.6	5.4	Petrakis 1953
Lymphocyte (s)	6.5	3.6	n, a	1118.1	5.4	Petrakis 1953
Lymphocyte (s)	5.0	3.6	n, a	671.6	5.5	Petrakis 1953
Lymphocyte (s)	8.2	3.6	n, a	1737.6	5.5	Petrakis 1953
Lymphocyte (s)	6.0	3.6	n, a	950.4	5.5	Petrakis 1953
Lymphocyte (s)	7.0	3.6	n, a	1293.6	5.5	Petrakis 1953
Lymphocyte (s)	7.5	3.7	n, a	1462.0	5.6	Petrakis 1953
Lymphocyte (s)	7.8	3.7	n, a	1579.6	5.6	Petrakis 1953
Lymphocyte (s)	7.0	3.7	n, a	1286.0	5.7	Petrakis 1953
Lymphocyte (s)	5.0	3.7	n, a	671.6	5.7	Petrakis 1953
Lymphocyte (s)	6.2	3.8	n, a	1028.4	5.8	Petrakis 1953
Lymphocyte (s)	6.0	3.8	n, a	956.9	5.8	Petrakis 1953
Lymphocyte (s)	6.0	3.8	n, a	953.5	5.8	Petrakis 1953

Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Lymphocyte (s)	6.6	3.8	n, a	1142.7	5.8	Petrakis 1953
Lymphocyte (s)	7.1	3.8	n, a	1317.1	5.8	Petrakis 1953
Lymphocyte (s)	6.5	3.8	n, a	1113.7	5.8	Petrakis 1953
Lymphocyte (s)	6.0	3.9	n, a	953.5	5.9	Petrakis 1953
Lymphocyte (s)	10.1	3.9	n, a	2627.9	5.9	Petrakis 1953
Lymphocyte (s)	6.8	3.9	n, a	1217.0	5.9	Petrakis 1953
Lymphocyte (s)	6.1	3.9	n, a	980.3	5.9	Petrakis 1953
Lymphocyte (s)	6.3	3.9	n, a	1035.5	6.0	Petrakis 1953
Lymphocyte (s)	7.2	3.9	n, a	1364.7	6.0	Petrakis 1953
Lymphocyte (s)	7.5	3.9	n, a	1495.0	6.0	Petrakis 1953
Lymphocyte (s)	7.0	3.9	n, a	1299.0	6.0	Petrakis 1953
Lymphocyte (s)	7.2	4.0	n, a	1370.2	6.1	Petrakis 1953
Lymphocyte (s)	8.0	4.0	n, a	1657.9	6.1	Petrakis 1953
Lymphocyte (s)	8.1	4.0	n, a	1711.1	6.1	Petrakis 1953
Lymphocyte (s)	10.0	4.0	n, a	2594.9	6.1	Petrakis 1953
Lymphocyte (s)	6.5	4.0	n, a	1113.7	6.1	Petrakis 1953
Lymphocyte (s)	6.8	4.0	n, a	1209.7	6.1	Petrakis 1953
Lymphocyte (s)	5.6	4.0	n, a	836.6	6.2	Petrakis 1953
Lymphocyte (s)	9.0	4.1	n, a	2115.1	6.2	Petrakis 1953
Lymphocyte (s)	7.0	4.1	n, a	1278.2	6.2	Petrakis 1953
Lymphocyte (s)	6.8	4.1	n, a	1217.0	6.3	Petrakis 1953
Lymphocyte (s)	6.0	4.2	n, a	950.4	6.4	Petrakis 1953
Lymphocyte (s)	6.5	4.2	n, a	1113.7	6.4	Petrakis 1953
Lymphocyte (s)	7.0	4.2	n, a	1286.0	6.4	Petrakis 1953
Lymphocyte (s)	7.6	4.2	n, a	1512.1	6.4	Petrakis 1953
Lymphocyte (s)	6.6	4.2	n, a	1142.7	6.4	Petrakis 1953
Lymphocyte (s)	7.0	4.3	n, a	1299.0	6.5	Petrakis 1953
Lymphocyte (s)	7.6	4.3	n, a	1503.6	6.5	Petrakis 1953
Lymphocyte (s)	6.0	4.3	n, a	950.4	6.5	Petrakis 1953

DNA—Cell Size Relationship in Humans

#### Table 1. Continued

Cite this article as Cold Spring Harb Perspect Biol 2015;7:a019091

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Lymphocyte (s)	6.9	4.3	n, a	1252.5	6.5	Petrakis 1953
Lymphocyte (s)	8.3	4.3	n, a	1818.2	6.6	Petrakis 1953
Lymphocyte (s)	6.0	4.3	n, a	966.9	6.6	Petrakis 1953
Lymphocyte (s)	6.6	4.3	n, a	1150.1	6.6	Petrakis 1953
Lymphocyte (s)	7.0	4.3	n, a	1286.0	6.6	Petrakis 1953
Lymphocyte (s)	7.1	4.3	n, a	1325.1	6.6	Petrakis 1953
Lymphocyte (s)	7.2	4.3	n, a	1356.6	6.6	Petrakis 1953
Lymphocyte (s)	7.5	4.4	n, a	1485.0	6.6	Petrakis 1953
Lymphocyte (s)	6.5	4.4	n, a	1106.4	6.7	Petrakis 1953
Lymphocyte (s)	7.6	4.4	n, a	1503.6	6.7	Petrakis 1953
Lymphocyte (s)	7.4	4.4	n, a	1435.4	6.7	Petrakis 1953
Lymphocyte (s)	6.3	4.4	n, a	1056.6	6.7	Petrakis 1953
Lymphocyte (s)	6.6	4.4	n, a	1135.6	6.7	Petrakis 1953
Lymphocyte (s)	6.0	4.4	n, a	966.9	6.7	Petrakis 1953
Lymphocyte (s)	7.0	4.4	n, a	1299.0	6.8	Petrakis 1953
Lymphocyte (s)	8.0	4.5	n, a	1666.4	6.8	Petrakis 1953
Lymphocyte (s)	9.0	4.5	n, a	2095.0	6.8	Petrakis 1953
Lymphocyte (s)	5.5	4.5	n, a	811.7	6.8	Petrakis 1953
Lymphocyte (s)	6.2	4.5	n, a	1014.6	6.8	Petrakis 1953
Lymphocyte (s)	6.8	4.5	n, a	1221.9	6.8	Petrakis 1953
Lymphocyte (s)	7.6	4.5	n, a	1503.6	6.8	Petrakis 1953
Lymphocyte (s)	5.8	4.5	n, a	897.3	6.9	Petrakis 1953
Lymphocyte (s)	8.0	4.6	n, a	1666.4	6.9	Petrakis 1953
Lymphocyte (s)	7.0	4.6	n, a	1299.0	6.9	Petrakis 1953
Lymphocyte (s)	7.4	4.6	n, a	1453.6	7.0	Petrakis 1953
Lymphocyte (s)	7.8	4.6	n, a	1588.3	7.0	Petrakis 1953
Lymphocyte (s)	9.0	4.6	n, a	2105.0	7.0	Petrakis 1953
Lymphocyte (s)	8.9	4.6	n, a	2075.5	7.0	Petrakis 1953
Lymphocyte (s)	6.1	4.6	n, a	1000.9	7.0	Petrakis 1953

Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Lymphocyte (s)	7.0	4.6	n, a	1286.0	7.0	Petrakis 1953
Lymphocyte (s)	6.3	4.6	n, a	1035.5	7.1	Petrakis 1953
Lymphocyte (s)	6.6	4.6	n, a	1150.1	7.1	Petrakis 1953
Lymphocyte (s)	7.6	4.6	n, a	1503.6	7.1	Petrakis 1953
Lymphocyte (s)	6.9	4.7	n, a	1239.8	7.1	Petrakis 1953
Lymphocyte (s)	7.8	4.7	n, a	1605.5	7.1	Petrakis 1953
Lymphocyte (s)	9.0	4.7	n, a	2085.5	7.1	Petrakis 1953
Lymphocyte (s)	7.8	4.7	n, a	1595.5	7.1	Petrakis 1953
Lymphocyte (s)	6.2	4.7	n, a	1007.9	7.2	Petrakis 1953
Lymphocyte (s)	7.5	4.7	n, a	1478.5	7.2	Petrakis 1953
Lymphocyte (s)	8.0	4.7	n, a	1666.4	7.2	Petrakis 1953
Lymphocyte (s)	6.5	4.7	n, a	1132.6	7.2	Petrakis 1953
Lymphocyte (s)	6.8	4.7	n, a	1209.7	7.2	Petrakis 1953
Lymphocyte (s)	7.0	4.7	n, a	1286.0	7.2	Petrakis 1953
Lymphocyte (s)	7.4	4.7	n, a	1437.3	7.2	Petrakis 1953
Lymphocyte (s)	7.0	4.7	n, a	1307.0	7.2	Petrakis 1953
Lymphocyte (s)	7.4	4.8	n, a	1429.3	7.3	Petrakis 1953
Lymphocyte (s)	5.8	4.8	n, a	894.0	7.3	Petrakis 1953
Lymphocyte (s)	8.0	4.8	n, a	1675.4	7.3	Petrakis 1953
Lymphocyte (s)	7.6	4.8	n, a	1520.3	7.3	Petrakis 1953
Lymphocyte (s)	7.0	4.8	n, a	1275.9	7.4	Petrakis 1953
Lymphocyte (s)	7.0	4.9	n, a	1291.5	7.4	Petrakis 1953
Lymphocyte (s)	7.2	4.9	n, a	1346.3	7.4	Petrakis 1953
Lymphocyte (s)	7.4	4.9	n, a	1437.3	7.4	Petrakis 1953
Lymphocyte (s)	7.5	4.9	n, a	1470.4	7.4	Petrakis 1953
Lymphocyte (s)	6.7	4.9	n, a	1194.4	7.5	Petrakis 1953
Lymphocyte (s)	9.0	4.9	n, a	2095.0	7.5	Petrakis 1953
Lymphocyte (s)	6.0	4.9	n, a	953.5	7.5	Petrakis 1953
Lymphocyte (s)	8.0	4.9	n, a	1666.4	7.5	Petrakis 1953

Table 1. Continued

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Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Lymphocyte (s)	7.1	5.0	n, a	1314.9	7.6	Petrakis 1953
Lymphocyte (s)	8.1	5.0	n, a	1719.8	7.6	Petrakis 1953
Lymphocyte (s)	8.0	5.0	n, a	1666.4	7.7	Petrakis 1953
Lymphocyte (s)	7.9	5.1	n, a	1614.3	7.7	Petrakis 1953
Lymphocyte (s)	6.0	5.1	n, a	966.9	7.7	Petrakis 1953
Lymphocyte (s)	7.0	5.1	n, a	1283.6	7.7	Petrakis 1953
Lymphocyte (s)	7.0	5.1	n, a	1293.6	7.8	Petrakis 1953
Lymphocyte (s)	8.4	5.1	n, a	1846.4	7.8	Petrakis 1953
Lymphocyte (s)	7.7	5.1	n, a	1571.3	7.8	Petrakis 1953
Lymphocyte (s)	6.6	5.1	n, a	1150.1	7.8	Petrakis 1953
Lymphocyte (s)	8.0	5.1	n, a	1657.9	7.8	Petrakis 1953
Lymphocyte (s)	8.1	5.1	n, a	1711.1	7.8	Petrakis 1953
Lymphocyte (s)	6.8	5.2	n, a	1224.7	7.9	Petrakis 1953
Lymphocyte (s)	7.5	5.2	n, a	1495.0	7.9	Petrakis 1953
Lymphocyte (s)	8.0	5.2	n, a	1684.4	7.9	Petrakis 1953
Lymphocyte (s)	6.0	5.2	n, a	970.6	7.9	Petrakis 1953
Lymphocyte (s)	8.0	5.3	n, a	1691.8	8.0	Petrakis 1953
Lymphocyte (s)	6.0	5.3	n, a	940.2	8.1	Petrakis 1953
Lymphocyte (s)	8.0	5.3	n, a	1684.4	8.1	Petrakis 1953
Lymphocyte (s)	8.1	5.3	n, a	1719.8	8.1	Petrakis 1953
Lymphocyte (s)	5.6	5.4	n, a	843.0	8.1	Petrakis 1953
Lymphocyte (s)	7.0	5.4	n, a	1307.0	8.2	Petrakis 1953
Lymphocyte (s)	9.8	5.4	n, a	2486.4	8.2	Petrakis 1953
Lymphocyte (s)	9.0	5.4	n, a	2085.5	8.2	Petrakis 1953
Lymphocyte (s)	8.0	5.4	n, a	1674.1	8.2	Petrakis 1953
Lymphocyte (s)	8.0	5.5	n, a	1666.4	8.3	Petrakis 1953
Lymphocyte (s)	7.1	5.5	n, a	1314.9	8.4	Petrakis 1953
Lymphocyte (s)	8.0	5.5	n, a	1691.8	8.4	Petrakis 1953
Lymphocyte (s)	7.3	5.5	n, a	1388.8	8.4	Petrakis 1953



Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Lymphocyte (s)	8.6	5.6	n, a	1912.5	8.5	Petrakis 1953
Lymphocyte (s)	9.0	5.6	n, a	2095.0	8.5	Petrakis 1953
Lymphocyte (s)	7.5	5.6	n, a	1487.0	8.5	Petrakis 1953
Lymphocyte (s)	7.2	5.6	n, a	1356.6	8.6	Petrakis 1953
Lymphocyte (s)	6.1	5.7	n, a	977.2	8.7	Petrakis 1953
Lymphocyte (s)	6.8	5.7	n, a	1229.6	8.7	Petrakis 1953
Lymphocyte (s)	7.6	5.7	n, a	1512.1	8.7	Petrakis 1953
Lymphocyte (s)	8.4	5.7	n, a	1856.2	8.7	Petrakis 1953
Lymphocyte (s)	5.6	5.8	n, a	824.1	8.8	Petrakis 1953
Lymphocyte (s)	7.1	5.8	n, a	1309.1	8.8	Petrakis 1953
Lymphocyte (s)	9.2	5.9	n, a	2215.0	9.0	Petrakis 1953
Lymphocyte (s)	6.3	6.0	n, a	1049.4	9.1	Petrakis 1953
Lymphocyte (s)	7.8	6.0	n, a	1597.1	9.1	Petrakis 1953
Lymphocyte (s)	8.0	6.0	n, a	1675.4	9.1	Petrakis 1953
Lymphocyte (s)	8.0	6.1	n, a	1684.4	9.3	Petrakis 1953
Lymphocyte (s)	9.0	6.1	n, a	2095.0	9.4	Petrakis 1953
Lymphocyte (s)	7.0	6.2	n, a	1307.0	9.5	Petrakis 1953
Lymphocyte (s)	6.9	6.3	n, a	1267.8	9.5	Petrakis 1953
Lymphocyte (s)	9.2	6.3	n, a	2205.2	9.6	Petrakis 1953
Lymphocyte (s)	8.0	6.4	n, a	1684.4	9.7	Petrakis 1953
Lymphocyte (s)	9.2	6.4	n, a	2205.2	9.8	Petrakis 1953
Lymphocyte (s)	9.2	6.5	n, a	2215.0	9.9	Petrakis 1953
Lymphocyte (s)	9.1	6.6	n, a	2134.8	10.0	Petrakis 1953
Lymphocyte (s)	8.0	6.6	n, a	1666.4	10.1	Petrakis 1953
Lymphocyte (s)	8.0	6.7	n, a	1666.4	10.2	Petrakis 1953
Lymphocyte (s)	6.6	6.8	n, a	1162.1	10.4	Petrakis 1953
Lymphocyte (s)	9.0	6.9	n, a	2124.7	10.5	Petrakis 1953
Lymphocyte (s)	9.1	6.9	n, a	2164.8	10.5	Petrakis 1953
Lymphocyte (s)	9.1	7.0	n, a	2144.5	10.7	Petrakis 1953

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Table 1. Continued

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Cell/nuclear **DNA** content Size, content size (original (original (original Cell volume DNA content Cell type  $(\mu^3)$ values) values) units) (pg) Sources Lymphocyte (s) Petrakis 1953 11.1 7.4 n, a 3158.4 11.3 Lymphocyte (s) 12.1 8.5 3762.4 13.0 Petrakis 1953 n, a Lymphocyte polymorph (m) 17.9 849.2 Bedi and Goldstein 1976 25.1 cc, a 6.5 Lymphocyte polymorph (m) 22.8 18.0 773.8 6.5 Bedi and Goldstein 1976 cc, a Lymphocyte polymorph (m) 21.4 18.0 728.5 Bedi and Goldstein 1976 6.5 cc, a Lymphocyte polymorph (m) 20.0 18.1 683.1 6.6 Bedi and Goldstein 1976 cc, a Lymphocyte polymorph (m) 18.2 6.6 Bedi and Goldstein 1976 24.4 cc, a 826.6 Lymphocyte polymorph (m) 21.3 18.2 724.8 6.6 Bedi and Goldstein 1976 cc, a Lymphocyte polymorph (m) 18.2 Bedi and Goldstein 1976 27.1 cc, a 916.9 6.6 Lymphocyte polymorph (m) 22.9 18.5 777.6 6.7 Bedi and Goldstein 1976 cc, a 18.9 Lymphocyte polymorph (m) 25.7 cc, a 871.8 6.9 Bedi and Goldstein 1976 Lymphocyte polymorph (m) 28.0 19.0 946.9 6.9 Bedi and Goldstein 1976 cc, a Lymphocyte polymorph (m) 6.9 Bedi and Goldstein 1976 26.2 19.0 cc, a 886.8 Megakaryocyte (m) 15.5 3.9 d, p 1949.8 13.7 Ishibashi et al. 1986 Megakaryocyte (m) 15.3 4.4 d, p 1875.3 15.4 Ishibashi et al. 1986 Megakaryocyte (m) 16.1 4.4 d, p 2352.1 15.4 Ishibashi et al. 1986 Megakaryocyte (m) 16.5 4.4 2185.1 15.4 Ishibashi et al. 1986 d, p Megakaryocyte (m) 16.3 4.6 d, p 2267.6 16.1 Ishibashi et al. 1986 Megakaryocyte (m) 16.8 5.3 d, p 2482.7 18.6 Ishibashi et al. 1986 7.0 5059.8 Megakaryocyte (m) 21.3 d, p 24.5 Ishibashi et al. 1986 Megakaryocyte (m) 7.3 Ishibashi et al. 1986 20.8 d, p 4711.8 25.6 Megakaryocyte (m) 22.4 8.6 d, p 5884.9 30.1 Ishibashi et al. 1986 Megakaryocyte (m) 27.5 Ishibashi et al. 1986 10.4 d, p 10889.2 36.4 Megakaryocyte (m) 10.6 d, p 11008.4 37.1 Ishibashi et al. 1986 27.6 Ishibashi et al. 1986 Megakaryocyte (m) 30.2 10.9 d, p 14421.8 38.2 Megakaryocyte (m) 30.6 12.6 15002.5 Ishibashi et al. 1986 d, p 44.1 Megakaryocyte (m) 32.3 12.8 d, p 17644.4 44.8 Ishibashi et al. 1986 Megakaryocyte (m) 33.3 14.1 d, p 19334.4 49.4 Ishibashi et al. 1986 Megakaryocyte (m) 33.8 14.5 d, p 20218.5 50.8 Ishibashi et al. 1986 Megakaryocyte (m) 33.8 16.1 d, p 20218.5 56.4 Ishibashi et al. 1986

Table 1. Continued

Cell type	Cell/nuclear size (original values)	DNA content (original values)	Size, content (original units)	Cell volume $(\mu^3)$	DNA content (pg)	Sources
Megakaryocyte (m)	34.2	16.8	d, p	20944.8	58.8	Ishibashi et al. 1986
Megakaryocyte (m)	34.1	16.9	d, p	20761.6	59.2	Ishibashi et al. 1986
Megakaryocyte (m)	34.4	17.2	d, p	21314.4	60.2	Ishibashi et al. 1986
Megakaryocyte (m)	34.4	17.2	d, p	21314.4	60.2	Ishibashi et al. 1986
Megakaryocyte (m)	35.0	17.7	d, p	22449.3	62.0	Ishibashi et al. 1986
Megakaryocyte (m)	36.0	17.7	d, p	24429.0	62.0	Ishibashi et al. 1986
Megakaryocyte (m)	35.4	18.6	d, p	23227.8	65.1	Ishibashi et al. 1986
	35.4	19.3		23227.8	67.6	Ishibashi et al. 1986
Megakaryocyte (m)	35.4 36.4	19.5	d, p	25252.4		Ishibashi et al. 1986
Megakaryocyte (m)	21.4	19.6	d, p	728.5	68.6 7.0	Bedi and Goldstein 1976
Monocyte (s)			cc, a			Bedi and Goldstein 1976
Monocyte (s)	32.1	19.9	cc, a	1081.8	7.2	
Monocyte (s)	26.3	20.0	cc, a	890.6	7.3	Bedi and Goldstein 1976
Monocyte (s)	30.8	20.1	cc, a	1036.9	7.3	Bedi and Goldstein 1976
Monocyte (s)	30.6	20.2	cc, a	1033.1	7.3	Bedi and Goldstein 1976
Monocyte (s)	30.9	20.3	cc, a	1040.7	7.4	Bedi and Goldstein 1976
Monocyte (s)	46.2	21.5	cc, a	1543.3	7.8	Bedi and Goldstein 1976
Primary spermatocyte (m)	37.6	2.6	cc, a	1184.8	7.6	Leuchtenberger et al. 1956
Primary spermatocyte (m)	28.1	3.3	cc, a	980.5	9.7	Leuchtenberger et al. 1956
Primary spermatocyte (m)	59.3	3.3	cc, a	1593.2	9.7	Leuchtenberger et al. 1956
Primary spermatocyte (m)	42.1	3.9	cc, a	1275.2	11.3	Leuchtenberger et al. 1956
Primary spermatocyte (m)	69.4	4.6	cc, a	1764.7	13.3	Leuchtenberger et al. 1956
Primary spermatocyte (m)	45.1	4.6	cc, a	1333.5	13.4	Leuchtenberger et al. 1956
Primary spermatocyte (m)	48.6	5.1	cc, a	1399.9	14.7	Leuchtenberger et al. 1956
Primary spermatocyte (m)	48.2	5.4	cc, a	1392.4	15.8	Leuchtenberger et al. 1956
Secondary spermatocyte (m)	14.2	1.5	cc, a	629.2	4.2	Leuchtenberger et al. 1956
Secondary spermatocyte (m)	13.1	1.9	cc, a	597.0	5.4	Leuchtenberger et al. 1956
Secondary spermatocyte (m)	17.6	1.9	cc, a	723.4	5.5	Leuchtenberger et al. 1956
Secondary spermatocyte (m)	19.9	2.3	cc, a	783.5	6.8	Leuchtenberger et al. 1956
Secondary spermatocyte (m)	33.4	2.6	cc, a	1097.0	7.7	Leuchtenberger et al. 1956

Cell/nuclear

Size, content

**DNA** content

7.5

7.7

7.9

1.8

11.4

13.0

11.7

81.0

Spermatozoon (m)

Spermatozoon (m)

Spermatozoon (m)

Syncytiotrophoblast (m)

24

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cc, a

cc, a

cc, a

cc, p

394.0

447.9

405.1

769.0

3.7

3.8

3.9

6.4

Royere et al. 1988

Royere et al. 1988

Royere et al. 1988

Galton 1962



Table 1. Continued

Cell type	Cell/nuclear size (original	DNA content (original	Size, content (original	Cell volume	DNA content	
	values)	values)	units)	$(\mu^3)$	(pg)	Sources
Syncytiotrophoblast (m)	72.0	1.9	cc, p	712.3	6.7	Galton 1962
Syncytiotrophoblast (m)	79.0	2.2	cc, p	756.6	7.7	Galton 1962
Syncytiotrophoblast (m)	77.0	2.2	cc, p	744.1	7.8	Galton 1962
Villous stroma (m)	82.0	2.1	cc, p	775.2	7.4	Galton 1962
Villous stroma (m)	108.0	2.1	cc, p	927.1	7.5	Galton 1962
Villous stroma (m)	96.0	2.4	cc, p	858.8	8.5	Galton 1962
Villous stroma (m)	102.0	2.5	cc, p	893.3	8.7	Galton 1962

Data from 19 cell types used in analyses shown in Figures 1 and 2. The original (i.e., reported) values for cell/nuclear size and DNA content are listed in columns 2 and 3, and the original units in which these values were expressed are listed in column 4. For analyses, all of the measures of cell/nuclear size were converted to units of cell volume ( $\mu^3$ ) shown in column 5, and measures of DNA content were converted to picograms of DNA shown in column 6. Letters in parentheses following the listing of cell type in column 1 denote whether the value is a measurement of a single cell (s) or the mean of a population of cells (m). The first notation used in column 4 indicates whether cell/nuclear size measurements were originally reported as cell diameter (d), nuclear cross-sectional area in calibrated arbitrary units (c), nuclear cross-sectional area in  $\mu^2$  (cc), nuclear diameter in  $\mu$  (n), or cell volume in  $\mu^3$  (v). The second notation in column 4 indicates whether DNA content measurements were reported in units of ploidy level (p), arbitrary absorbance units (a), or picograms of DNA (pg). Cell types are reported based on cell-type designations given in original studies.

Computation of mean cell volumes: Studies reported cell or nuclear size using one of four measurements: cell volume, cell diameter/radius, nuclear cross-sectional area, or nuclear diameter/radius. Measurements of cell diameter/radius were converted to cell volume assuming cells were spherical. Measurements of nuclear cross-sectional area and diameter/radius were converted to nuclear volume assuming nuclei were spherical and that nuclear cross-sectional areas were circular. All estimates of nuclear volume were then converted to cell volume using a previously reported relationship between cell and nuclear volume of human cells (Swanson et al. 1991): cell volume = 44.2 (nuclear volume)<sup>0.65</sup> as indicated in the text. In one case (Klinger and Schwarzacher 1960), measurements of nuclear area were given in calibrated arbitrary units. In this case, we converted back from arbitrary units to units of squared micrometers before computing nuclear radius (in micrometers).

Computation of mean DNA content: DNA content measurements were originally reported in one of three ways: as picograms of DNA, as ploidy level, or as arbitrary absorbance units of DNA. To convert ploidy level to pg DNA, we multiplied by the typical haploid DNA content of human cells (3.5 pg [Gregory 2012]). To convert arbitrary units to pg DNA, we determined the mean arbitrary unit value of cells that the original investigators of each study designated as diploid, and then assumed this corresponded to  $2 \times 3.5 = 7$  pg of DNA to convert values from arbitrary units to pg DNA. Thus, each of the seven studies in which this conversion was performed had a unique conversion factor: Bedi and Goldstein 1976 (2.75 arbitrary units/pg DNA), Klinger and Schwarzacher 1960 (6.9 arbitrary units/pg DNA), Leuchtenberger et al. 1956 (0.34 arbitrary units/pg DNA), Meek and Harbison 1967 (2.1 arbitrary units/pg DNA), Petrakis 1953 (0.66 arbitrary units/pg DNA), Royere et al. 1988 (2.03 arbitrary units/pg DNA), and Swartz 1956 (0.37 pg). This conversion is possible because the Feulgen staining method results in a linear relationship between light absorbance and DNA content as discussed in the text.

Averaging individal cell measurements: For cases in which individual cell measurements were reported, we averaged all cell volume and DNA content measurements to yield a single population mean for each cell type within each study. Mean cell volumes of these cell types were 1574  $\mu^3$  (amnion epithelial cells [Klinger and Schwarzacher 1960]), 2936  $\mu^3$  (hepatocytes [Meek and Harbison 1967]), 1441  $\mu^3$  (lymphocytes [Petrakis 1953]), 764  $\mu^3$  (lymphocytes [Bedi and Goldstein 1976]), 817  $\mu^3$  (lymphocyte polymorphs [Bedi and Goldstein 1976]), and 1051  $\mu^3$  (monocytes [Bedi and Goldstein 1976]). Corresponding mean DNA contents were 12, 19.9, 7.1, 6.8, 6.7, and 7.3 pg, respectively. These means were used in the analyses presented in the text.

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#### REFERENCES

\*Reference is also in this collection

- \* Amodeo AA, Skothein JM. 2015. Cell size control. Cold Spring Harb Perspect Biol doi: 10.1101/cshper spect.a019083.
- Anatskaya OV, Vinogradov AE. 2011. Somatic polyploidy promotes cell function under stress and energy depletion: Evidence from tissue-specific mammal transcriptome. Funct Integr Genomic 10: 433–446.
- Bachmann K, Goin OB, Goid CJ. 1966. Hylid frogs: Polyploid classes of DNA in liver nuclei. Science 154: 650– 651
- Beaton MJ, Hebert PDN. 1989. Miniature genomes and endopolyploidy in cladoceran crustaceans. Genome 32: 1048-1053.
- Bedi KS, Goldstein DJ. 1976. Apparent anomalies in nuclear Feulgen-DNA contents. Role of systematic microdensitometric errors. *J Cell Biol* 71: 68–88.
- Biesterfeld S, Gerres K, Fischer-Wein G, Bocking A. 1994. Polyploidy in non-neoplastic tissues. *J Clin Pathol* 47: 38–42.

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- Biesterfeld S, Beckers S, Del Carmen Villa Cadenas M, Schramm M. 2011. Feulgen staining remains the gold standard for precise DNA image cytometry. *Anticancer Res* **31**: 53–58.
- Chieco P. Derenzini M. 1999. The Feulgen reaction 75 years on. *Histochem Cell Biol* 111: 345–358.
- De Veylder L, Larkin JC, Schnittger A. 2011. Molecular control and function of endoreplication in development and physiology. *Trends Plant Sci* **16:** 624–634.

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- Dorman A, Graham D, Curran B, Henry K, Leader M. 1990. Ploidy of smooth muscle tumours: Retrospective image analysis study of formalin fixed, paraffin wax embedded tissue. *J Clin Pathol* **43**: 465–468.
- Ehrie MG, Swartz FJ. 1974. Diploid, tetraploid, and octaploid β cells in the islets of Langerhans of the normal human pancreas. *Diabetes* 23: 583–588.
- Feulgen R, Rosenbeck HC. 1942. Manual of histological demonstration technique. Butterworth, London.
- Galton M. 1962. DNA content of placental nuclei. *J Cell Biol*13: 183–191.
- Gregory TR. 2005. *The evolution of the genome*, pp. 1–768 Academic, New York.
- Gregory TR. 2012. Animal genome size database, www...genomesize.com.
- Hancock V, Martin JF, Lelchuk R. 2008. The relationship between human megakaryocyte nuclear DNA content and gene expression. *Brit J Haematol* 85: 692–697.
- Heizer P. 1955. Desoxyribose nucleic acid (DNA) content and size of rat liver nuclei during thioac etamide intoxication and recovery. *Chromosoma* 7: 281–327.
- Ishibashi T, Ruggeri ZM, Harker LA, Burstein SA. 1986. Separation of human megakaryocytes by state of differ-

- entiation on continuous gradients of Percoll: Size and ploidy analysis of cells identified by monoclonal antibody to glycoprotein IIb/IIIa. *Blood* 67: 1286–1292.
- Jovtchev G, Schubert V, Barrow M, Schubert I. 2006. Nuclear DNA content and nuclear and cell volumes are positively correlated in angiosperms. Cytogenet Genome Res 114: 77–82.
- Klinger HP, Schwarzacher HG. 1960. The sex chromatin and heterochromatic bodies in human diploid and polyploid nuclei. J Biophys Biochem Cytol 8: 345–364.
- Lee HO, Davidson JM, Duronio RJ. 2009. Endoreplication: Polyploidy with purpose. Genes Dev 23: 2461–2477.
- Leuchtenberger C, Leuchtenberger R, Schrader F, Weir DR. 1956. Reduced amounts of desoxyribose nucleic acid in testicular germ cells of infertile men with active spermatogenesis. *Lab Invest* 5: 422–440.
- Marguerat S, Bähler J. 2012. Coordinating genome expression with cell size. *Trends Genet* **28:** 560–565.
- Meek ES, Harbison JFA. 1967. Nuclear area and deoxyribonucleic acid content in human liver cell nuclei. *J Anat* 101: 487–489.
- Mirsky AE, Ris H. 1949. Variable and constant components of chromosomes. *Nature* 163: 666–667.
- Niklas KJ. 2015. A phyletic perspective on cell growth. Cold Spring Harb Perspect Biol doi: 10.1101/cshper spect.a019158.
- Olmo E. 1983. Nucleotype and cell size in vertebrates: A review. *Basic Appl Histochem* **27**: 227–256.
- Parfrey LW, Lahr DJG, Katz LA. 2008. The dynamic nature of eukaryotic genomes. *Mol Biol Evol* **25:** 787–794.
- Petrakis NL. 1953. Microspectrophotometric estimation of the desoxyribonucleic acid (DNA) content of individual normal and leukemic human lymphocytes. *Blood* 8: 905-915
- Pinheiro J, Bates D. 2000. Mixed-effects models in S and S-PLUS. Springer, New York.
- Price HJ, Sparrow AH, Nauman AE 1973. Correlations between nuclear volume, cell volume and DNA content in meristematic cells of herbaceous angiosperms. Cell Mol Life Sci 29: 1028–1029.
- Royere D, Hamamah S, Nicolle JC, Barthelemy C, Lansac J. 1988. Freezing and thawing alter chromatin stability of ejaculated human spermatozoa: Fluorescence acridine orange staining and Feulgen-DNA cytophotometric studies. *Gamete Res* 21: 51–57.
- Shuter BJ, Thomas JE, Taylor WD, Zimmerman AM. 1983. Phenotypic correlates of genomic DNA content in unicellular eukaryotes and other cells. Am Nat 122: 26–44.
- Swanson JA, Lee M, Knapp PE. 1991. Cellular dimensions affecting the nucleocytoplasmic volume ratio. J Cell Biol 115: 941–948.
- Swartz FJ. 1956. The development in the human liver of multiple desoxyribose nucleic acid (DNA) classes and their relationship to the age of the individual. *Chromo-soma* 8: 53-72.
- Szarski H. 1976. Cell size and nuclear DNA content in vertebrates. *Int Rev Cytol* **44:** 93–111.

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Therman E, Sarto GE, Stubblefield PA. 1983. Endomitosis: A reappraisal. *Human Genet* **63**: 13–18.

Ullah Z, Lee CY, Lilly MA, DePamphilis ML. 2009. Developmentally programmed endoreduplication in animals. *Cell Cycle* **8**: 1501–1509.

Vliegen HW, Bruschke VG, Van der Laarse A. 1990. Different response of cellular DNA content to cardiac hypertrophy in human and rat heart myocytes. *Comp Biochem Physiol* **95A:** 109–114.

Winkelmann M, Pfitzer P, Schneider W. 1987. Significance of polyploidy in megakaryocytes and other cells in health and tumor disease. *Klin Wochenschr* **65**: 1115–1131.

Wohlschlaeger J, Levkau B, Brockhoff G, Schmitz KJ, von Winterfeld M, Takeda A, Takeda N, Stypmann J, Vahlhaus C, Schmid C, et al. 2010. Hemodynamic support by left ventricular assist devices reduces cardiomyocyte DNA content in the failing human heart. Circulation 121:



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