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| 1 | Quantitative Assessment of Colorectal Morphology: Implications |
|-----------------|--|
| 2 | for Robotic Colonoscopy |
| 3 | |
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| 26 | |
| 27 | Abstract |

28 This paper presents a method of characterizing the distribution of colorectal morphometrics. It uses 29 three-dimensional region growing and topological thinning algorithms to determine and visualize the 30 luminal volume and centerline of the colon, respectively. Total and segmental lengths, diameters, 31 volumes, and tortuosity angles were then quantified. The effects of body orientations on these 32 parameters were also examined. Variations in total length were predominately due to differences in 33 the transverse colon and sigmoid segments, and did not significantly differ between body orientations. 34 The diameter of the proximal colon was significantly larger than the distal colon, with the largest 35 value at the ascending and caecum segments. The volume of the transverse colon was significantly the 36 largest, while those of the descending colon and rectum were the smallest. The prone position showed 37 a higher frequency of high angles and consequently found to be more torturous than the supine position. This study yielded a method for complete segmental measurements of healthy colorectal 38 39 anatomy and its tortuosity. The transverse and sigmoid colons were the major determinant in 40 tortuosity and morphometrics between body orientations. Quantitative understanding of these 41 parameters may potentially help to facilitate colonoscopy techniques, accuracy of polyp spatial 42 distribution detection, and design of novel endoscopic devices.

43

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45 Keywords: Robotic Colonoscopy, Colon, Large Intestine, Colorectal Cancer

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47 **1** Introduction

48 Colorectal cancer (CRC) is the third most common cancer worldwide, estimated to cause 10.2% 49 and 12.7% of the total cancer death in the UK and USA respectively [1, 2]. Currently, video-50 colonoscopy is the preferred screening method for the diagnosis of CRC [3, 4], however, it is attended 51 by certain clinical drawbacks such as patient discomfort, need for sedation, absence of 52 maneuverability of the scope, and a long learning curve. In order to improve compliance to the screening programs, an imaging modality - computed tomographic colonography (CTC) - was 53 54 developed for early detection of colorectal polyps and cancer [5]. Other new technologies are being 55 evaluated and have started to emerge, e.g. computer assisted colonoscopy [6-10] and active capsule 56 colonoscopy [11-13]. Similar to conventional colonoscopy, endoluminal navigation in these new 57 technologies is often challenging due to the convoluted nature of the colon.

58 Although the colon anatomy is well described, less is known about the variation of colorectal 59 morphometry in quantitative terms across the general population. Studies have shown that colonic anatomical features have a significant association with failure to achieve complete colonoscopy [14]. 60 61 An understanding of the length, diameter and tortuosity of the colon is important for the performance 62 of conventional colonoscopy; and specially for the future development of robotic colonoscopy 63 platforms that may have sections of fixed diameter and length. Consequently, using this information, 64 important features such as colonic elongation and distension can be extracted, and statistical 65 assessments such as polyp spatial distribution can be understood.

66 Colonic length has been investigated previously using barium enema, where accurate length 67 assessment is difficult due to the three-dimensional (3-D) intraluminal centreline. Only one study was found from the barium enema literature where the colonic diameter was estimated, in this case for 68 69 Japanese men and women [15]. It is of note that in these studies the colon contains a significant 70 volume of barium which may also influence the accuracy of measurements. Several barium enema 71 and intraoperative laparotomy studies have investigated how various colonic parameters and gender 72 might predict the degree of difficulty during colonoscopic examinations [16-18], but did not aim to 73 determine the morphology and tortuosity of the colon. One study described the intestinal length as a 74 whole in cadavers did not provide the length of the colonic segments used clinically [19]. Fig. 1 75 shows the 3-D geometry of the lumen and its anatomical segments. Several CTC studies reported the 76 length measurements for the colorectal anatomy [20, 21] and its correlation with difficult or 77 incomplete examinations [22, 23]. The high resolution of CTC technology, combined with regular CO2 insufflation and automated digital analyses, make this technology a far more accurate tool for 78 79 assessing the anatomy of the colon when compared to other methods such as barium enema. However, 80 little is known about the complete colon length, intra-colon segmental lengths, diameters, volume, and 81 tortuosity angle of colorectal anatomy in a healthy population.



82

Fig. 1. Schematic representation of the colonic segments and landmarks in anterior and lateral views; the dashed line represents the anatomical landmark position. All measurements were recorded for the six segments (rectum, sigmoid, descending, transverse, ascending, and cecum) using the automated centreline along the axis of the colonic lumen, solid arrow lines.

The aim of our investigation was to describe a method to quantitatively measure the luminal length, diameter, volume, and shape of the colon within an asymptomatic population undergoing primary CTC examinations. We also determined whether a correlation exists between these factors, and between scans taken in supine and prone orientations. These descriptors may have implications for conventional colonoscopy and CT colonography training and performance. Furthermore, they provide a quantitative description of the colonic environment that can be used for development of new colonoscopy devices, from incremental advances to current colonoscope technology to nextgeneration robotic systems [24]. An understanding of the diameter and tortuosity of the colon is particularly relevant for the future of these systems.

96 2 Materials and Methods

97 2.1 Patient Preparation and Data Acquisition

A single medically qualified researcher, under supervision of a consultant radiologist, searched the publically available TCIA (http://cancerimagingarchive.net/, sponsored by the Cancer Imaging Program, DCTD/NCI/NIH) retrospectively and selected clinical studies that demonstrated reportedly healthy colons in both prone and supine positions (no patients were excluded from this population). In total, 24 patient studies were selected at random: 12 men and 12 women. The average age of the sample was 54.8 ± 4.7 years ranging from 50 to 65.

103 The complete imaging methodology has been described previously [25]. Briefly, all patients had undergone 104 standard 24-hour colonic preparation with stool tagging following by the oral administration of 90 mL of 105 sodium phosphate (Phospho-soda, Fleet Pharmaceuticals) and bisacodyl tablets (10 mg) to reduce the presence 106 of any residual stool or fluid. The final step was to ingest a 6-oz (177 mL) glass of liquid containing at least 5 107 mL and up to 60 mL of water-soluble iodinated oral contrast material (diatrizoate meglumine and sodium 108 diatrizoate, Gastroview, Mallinckrodt Imaging) the night before the examination to label any residual colonic 109 fluid. All examinations were performed using at least a 16-channel helical CT scanner with 0.8 mm collimation, 110 1 mm reconstruction interval, matrix 512×512, 50 effective mAs, peak voltage of 120 kV, and B30f convolution 111 kernel. Data were obtained in the supine and prone positions; 1 mg of glucagon was administered 112 subcutaneously 7-15 minutes before the CT examination unless contraindicated or refused by the patient. 113 Colonic distention was achieved with an automated carbon dioxide insufflator (PROTOCO2L, E-Z-EM).

114 **2.2 Image Segmentation and Centerline**

115 We developed a multistage algorithm to process the CT data and generate a 3-D volumetric polygon mesh, 116 with an associated centerline, to be used in a quantitative description of the colon morphology. The first stage 117 employs clinical visualization software (Amira, FRI Visualization Sciences Group, USA) to identify low 118 attenuation voxels that represent gas in the colon lumen using a 3-D region growing algorithm [26]. An intensity 119 based threshold of less than -800HU was experimentally determined for each dataset (grey-level histogram of 120 the abdominal CT dataset) to segment the lumen's details. Increasing this threshold allows more details of the 121 lumen interior to be visualized, however, too high threshold may result in erroneous inclusion of the 122 surrounding tissue. Because the colon is not the only gas-filled organ, user-defined seed points were placed in 123 the lumen to spatially isolate the colon. If the value of the connected voxels were less than the intensity 124 threshold, they were included in the region. This algorithm stopped when no more voxels remain in the collected 125 neighbors. Two datasets required additional user interaction to guide the algorithm due to lumen occlusion by 126 peristalsis, spasm and/or residual faeces. This modeling was then checked against a visual review of the images 127 to ensure accuracy.

128 The segmented data was resampled to produce regular pixel spacing. The visualization software was then 129 used to implement a topological thinning algorithm to obtain a skeleton representation of the segmented colon 130 lumen. Using the anorectum as the starting point and most distal part of the cecum as the end point, the 131 algorithm automatically performs a skeleton tracing of the colon. The two endpoints required for centerline 132 calculation can also be automatically identified during the segmentation process based on distance fields [27]. 133 Due to the complexity of the segmented colonic lumen (e.g., haustral folds), the resulting skeleton may contain 134 many paths between two endpoints when only one corresponds to the centerline of the colon. We used a graph 135 theoretic algorithm, described by Ge et al., to remove extra branches [28] which are typically anomalies close to 136 the surface of the colon lumen rather than features associated with the actual centerline. The skeleton then 137 becomes a single continuous line from rectum to cecum, positioned centrally in the colon lumen and termed 138 herein the 'centreline'.

139 **2.3** Anatomical Colorectal Landmarks

140 As depicted in Fig. 1, the 3-D geometry of the lumen was classified into 6 anatomical segments using

141 landmarks identified on the centerline; the rectum, sigmoid colon, descending colon, transverse colon, ascending 142 colon, and cecum [29]. The spatial coordinates of these landmarks were evaluated by the same expert 143 researcher. The rectosigmoid junction was located between the sacral promontory and the S3 vertebral body 144 levels and included in the sigmoid colon. The pelvic brim was chosen as the location for sigmoid-descending 145 junction which the distal descending colon angles forwards. The leftmost and rightmost, most cranial inflexion 146 points of the colon were designated as the descending-transverse and transverse-ascending junctions, 147 respectively (the splenic and hepatic flexures). The ileocaecal valve was selected as a boundary between the 148 ascending colon and the cecum, where the valve was included in the proximal segment. The colon was also 149 divided into two functional parts: the proximal colon from the most proximal part of the cecum to the splenic 150 flexure; and the distal colon, from the splenic flexure to the anorectum.

151 **2.4 Data Processing**

The final stage of processing was performed using a custom program (implemented in Matlab, Mathworks Inc., USA) to analyze the 3-D volume and associated centerline of the colon for each dataset and from these compute the morphological metrics. To minimize local fluctuations in the centerline, the path was linearly resampled at regular 1 mm intervals along its length and then smoothed with a 40 point moving average filter.

157 Colorectal length and volume were calculated from the 3-D volume of the colon lumen and its 158 centerline between the successive colonic landmarks shown in Fig. 1. The effective colonic cross-159 section was described using a circle-fitting approach. This was done as the cross-section of the colonic lumen can have functionally redundant areas. Firstly, a series of planes were defined normal 160 161 to each component of the centerline (between successive 1 mm points) and then the intersection of 162 this plane with the colonic 3-D volume was found. This yielded a set of planar points for each cross-163 sectional plane. An iterative circle-fitting algorithm was then implemented to determine the radius of the largest circle which fits inside this point set, using the centerline point as an initial estimate for the 164 circle's center since it will always lie within the lumen. Collating these data then provides the 165 166 effective radius at 1 mm intervals along the colon's length.

Although the tortuosity does not have a formal clinical definition, there are clearly some intuitive properties, which a reasonable index must satisfy in order for it to correlate with the gross qualitative assessment of an expert observer. In order to obtain a meaningful tortuosity measure for the engineering community, we have calculated the in-plane angle of the centerline using a span size equal to the total mean diameter of the lumen (~3.5 cm), as shown in Fig. 2.

172 **3 Results**

We analyzed the colonic morphology in 24 patients in both supine and prone positions using the method described above. The variations in colonic elongation and tortuosity are visualized in Fig. 3. All the measurements were compared using the Wilcoxon signed-rank test and then underwent further post-hoc analysis using Exact testing to allow for the small number of subjects used with the significant level set at 0.05 (p<0.05). Table 1-3 summarize the length, diameter and volume of each

colonic segment and proximal, distal, and the total colons. The z-value is the test statistic for Wilcoxon signed-rank testing.



180 181 Fig. 2. Methodology for determining the angle (θ) of each point on the colonic centreline (e.g., at point n, this 182 angle was calculated by measuring the angle formed by two vectors between (n) \rightarrow (n-span) and (n) \rightarrow 183 (n+span).



184 185

Fig. 3. 3-D visualisation of the segmented colonic volumes from CTC, images are from patients TCIA 26, 87, 186 179, and 62. These cases demonstrate how colonic elongation and tortuosity (specifically for transverse and 187 sigmoid segments) differ from a relatively short textbook colon (a) to an elongated and tortuous colon (d). The automated centreline is overlaid on the rendered colons that allow morphometric measurements.

- 188
- 189

190 There was no significant difference in total colonic length between the supine (185.0 \pm 18.3 cm) 191 and prone (183.0 \pm 16.9 cm), nor was there any significant differences when mean lengths between 192 men and women groups were compared (187.7 \pm 19.0 cm in men and 182.2 \pm 18.1 cm in women). 193 However, when the colonic segments were considered, significant differences were found during positional. The descending colon was found to be significantly shorter in the supine position (z=-194 2.171, p < 0.03), while the ascending colon and cecum significantly shorter in the prone orientation. 195 196 The length of the transverse colon and sigmoid colon were significantly the largest in both 197 orientations, followed by the descending colon, rectum, ascending colon, and caecum. The transverse 198 and sigmoid segments also demonstrated the widest length variation in both positions. When 199 considering the proximal and distal lengths of the colon, the distal colon was found to be longer in 200 both orientations. The sigmoid colon was significantly longer (supine: p < 0.038 and prone: p < 0.04) in 201 women (supine: 54.4 ± 10.8 cm and prone: 53.5 ± 11.2 cm) than in men (supine: 46.8 ± 15.9 cm and 202 prone: 46.4 ± 11.6 cm). The descending and ascending colons were significantly longer in both 203 directions in men (p < 0.001). The cecum length was longer in male, but this finding attained significance only in the prone position (p < 0.001). The length of the other segments did not 204 205 significantly differ between the two gender groups.

206 The cross-sectional properties of the colon were considered across the entire colon length and per-207 segment. Fig. 4 shows a frequency distribution of colorectal radius across the group. Supine and prone 208 orientations exhibit similar trends, although the supine data are more positively skewed towards 209 higher radii. Fig. 5, Fig. 6 and Table 2 provide further detail of the radius within each segment of the 210 colon. The mean diameter of the ascending colon (supine: 4.5 cm and prone: 4.3 cm) was the largest followed by that of the cecum (supine: 4.4 cm and prone: 3.8 cm) in both orientations. The diameter 211 212 of the sigmoid colon was significantly the smallest (2.6 cm in both orientations) being the longest and narrowest segment of the colon. The diameter of the proximal colon was significantly larger than the 213 214 distal colon in both supine and prone orientations.



215

216 Fig. 4. Radius distribution for 24 patients in supine (white columns) and prone (pink columns) 217 orientations

218

Table 1. Comparison of the colonic segmental Length in supine and prone orientations in 24 patients (mean ± SD (median))

| | Length, cm | | | |
|-------------|--------------------|-----------------|--------------------|--|
| | Supine | z-value (p) | Prone | |
| Rectum | 23.4±6.7 (21.7) | -0.400 (0.705) | 23.1±3.9 (22.7) | |
| Sigmoid | 50.6±13.9 (51.6) | -0.971 (0.345) | 49.9±11.7 (48.7) | |
| Descending | 24.2±7.8 (23.1) | -2.171 (0.029*) | 26.0±7.8 (25.9) | |
| Transverse | 57.2±9.3 (56.6) | -0.086 (0.944) | 57.3±10.9 (56.9) | |
| Ascending | 21.7±4.2 (20.7) | -2.114 (0.034*) | 19.7±4.0 (20.3) | |
| Cecum | 7.8±2.9 (6.9) | -2.029 (0.042*) | 6.9±2.3 (6.8) | |
| Proximal | 86.6±9.7 (85.5) | -2.086 (0.037*) | 84.0±10.2 (82.9) | |
| Distal | 98.3±14.7 (97.7) | -0.057 (0.966) | 99.0±11.8 (98.8) | |
| Total Colon | 185.0±18.3 (187.5) | -1.086 (0.290) | 183.0±16.9 (185.0) | |

221 * Statistically Significant

222

Table 2. Comparison of the colonic segmental diameter in supine and prone orientations in 24 patients (mean \pm SD (median))

| | Diameter, cm | | | |
|-------------|---------------|-----------------|---------------|--|
| - | Supine | z-value (p) | Prone | |
| Rectum | 3.6±0.8 (3.4) | -1.400 (0.166) | 3.7±0.7 (3.6) | |
| Sigmoid | 2.6±0.4 (2.5) | -1.263 (0.214) | 2.6±0.3 (2.6) | |
| Descending | 3.3±0.6 (3.3) | -1.072 (0.294) | 3.2±0.5 (3.1) | |
| Transverse | 3.7±0.4 (3.7) | -2.030 (0.042*) | 3.6±0.5 (3.5) | |
| Ascending | 4.5±0.7 (4.7) | -2.359 (0.017*) | 4.3±0.7 (4.6) | |
| Cecum | 4.4±0.7 (4.5) | -3.187 (0.001*) | 3.8±0.6 (3.9) | |
| Proximal | 4.2±0.4 (4.3) | -3.514 (0.000*) | 3.9±0.5 (4.0) | |
| Distal | 3.1±0.5 (3.0) | -4.286 (0.000*) | 3.1±0.4 (3.1) | |
| Total Colon | 4.7±0.5 (4.7) | -4.286 (0.000*) | 3.5±0.4 (3.5) | |

225 * Statistically Significant

226

- Table 3. Comparison of the colonic segmental volume in supine and prone orientations in 24 patients
- 228 $(\text{mean} \pm \text{SD} (\text{median}))$

| | | Volume, cm ³ | |
|-------------|-----------------------|-------------------------|-----------------------|
| | Supine | z-value (p) | Prone |
| Rectum | 154.4±49.3 (156.7) | -1.686 (0.095*) | 170.1±44.0 (166.2) |
| Sigmoid | 195.6±104.9 (165.5) | -0.686 (0.509) | 190.2±92.2 (156.7) |
| Descending | 159.2±78.1 (154.8) | -1.057 (0.303) | 168.7±79.7 (165.3) |
| Transverse | 477.9±147.9 (466.9) | -1.429 (0.160) | 454.7±150.1 (433.2) |
| Ascending | 233.6±92.0 (237.0) | -2.914 (0.003*) | 198.0±94.9 (201.4) |
| Cecum | 156.1±370.3 (64.2) | -2.229 (0.025*) | 57.8±28.6 (57.2) |
| Proximal | 867.6±483.7 (823.0) | -2.943 (0.002*) | 710.5±212.0 (706.6) |
| Distal | 509.2±181.7 (448.2) | -0.857 (0.406) | 529.0±166.7 (481.7) |
| Total Colon | 1376.8±582.4 (1267.9) | -1.829 (0.069) | 1239.5±344.0 (1164.5) |

229 * Statistically Significant

230

Table 4. Comparison of the colonic segmental angle (mean \pm SD (median)), skewness (s), and kurtosis (k) in supine and prone orientations

| | Supine | | Prone | |
|------------|------------------|--------------|------------------|---------------|
| | Angle, deg | Skew (k) | Angle, deg | Skew |
| Rectum | 42.2±27.3 (42.2) | 0.03 (-0.84) | 45.2±28.1 (47.1) | -0.17 (-0.98) |
| Sigmoid | 46.9±24.2 (43.8) | 0.44 (-0.58) | 47.9±24.1 (45.1) | 0.34 (-0.71) |
| Descending | 33.6±21.1 (29.1) | 0.84 (0.07) | 36.1±21.7 (31.0) | 0.79 (-0.12) |
| Transverse | 38.7±23.3 (35.6) | 0.42 (-0.80) | 37.2±22.9 (33.0) | 0.51 (-0.69) |
| Ascending | 30.2±17.7 (27.1) | 0.69 (-0.09) | 31.8±18.0 (29.7) | 0.56 (-0.34) |
| Cecum | 27.1±15.5 (22.4) | 0.54 (-0.49) | 32.4±15.9 (31.9) | 0.42 (0.02) |
| Splenic * | 31.5±18.5 (27.4) | 0.84 (0.25) | 35.1±20.4 (30.3) | 0.74 (-0.17) |
| Hepatic * | 51.8±22.4 (52.5) | 0.00 (-0.82) | 51.6±21.8 (50.8) | 0.06 (-0.75) |

- ^{*} Artificially created segments defined around the splenic and hepatic flexures with a total length of
- 10 cm per segment





Fig. 5. The 5th (blue), 50th (green), and 95th (red) percentiles of the colorectal radius and the angle frequency distributions for individual colonic segments in supine and prone orientations; Dashed-lines represent prone orientation.

239

240 Across the colon as a whole, the total mean diameter was significantly larger in the supine position 241 (z=-4.29, p<0.01), demonstrated mainly in the significant difference between distal colons in two 242 different orientations (z=-4.29, p<0.01). There was no significant difference observed between the diameter of individual segments in the distal colon, however, all individual proximal segments 243 244 showed significant decrease in diameter when body orientation was changed to prone (cecum: z=-3.19, p < 0.01; ascending: z = -2.36, p < 0.02; and transverse: z = -2.03, p < 0.04). The diameter of the other 245 246 segments (distal portion) did not significantly differ between the two groups. The diameter of the proximal colon was larger in men than in women, whilst that of the distal colon was larger in women. 247 All other comparisons between the two gender groups yielded non-significant changes for the 248 249 diameter variations.

250 The total luminal volume of the colon was 1376.8 ± 582.4 cm³ in the supine position. This 251 measure was significantly larger in the proximal colon, with its biggest portion in the transverse colon 252 (477.9 cm3). Changing the body orientation to prone did not vary the total volume significantly 253 $(1239.5 \pm 344.0 \text{ cm}^3)$. However, when individual segmental volume was compared between the two 254 groups, the two most proximal segments were significantly smaller (cecum: z=-2.23, p<0.02 and 255 ascending: z=-2.91, p<0.01) in the prone orientation. When individual segmental volume was 256 compared between men and women, the cecum volume was significantly larger in men in both 257 orientations (supine: p < 0.013 and prone: p < 0.028), and the descending colon was significantly shorter in women (supine: p < 0.038 and prone: p < 0.005). All other segmental comparisons showed non-258 259 significant changes.

Assessment of colonic segmental tortuosity is achieved using a kernel density estimation to represent the segmental angle distribution, shown in Fig. 5 with the associated skewness (s) and kurtosis (k) parameters in Table 4. Prone data showed to be less skewed (more torturous) than the supine orientation in all segments except the transverse colon. In both orientations, the descending colon has the most skewed distribution (supine: s=0.84 and prone: s=0.79) closely followed by the ascending colon. Interestingly, the ascending and descending segments are held in place along their length by the colonic mesentery and considered to be less mobile than the transverse segment (which is much less constrained). The hepatic flexure exhibited a more normally distributed behavior (supine: s=0 and prone: s=0.06) when compared with the splenic flexure (supine: s=0.84 and prone: s=0.74). Data from the cecum were considered to be both less reliable and less relevant due to the segment's short length.



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Fig. 6. Two-dimensional visualisation of a segmented colon after total and segmental length normalisation; the colon centreline is laid flat and the diameter is shown as a shaded profile (for 5^{th} , 50^{th} , and 95^{th} percentiles) with anatomical landmarks.

275 4 Discussion

276 Colonoscopy is considered the gold standard for endoscopic examination of the whole colon. 277 However, it is still difficult to perform or complete, mainly due to the complex colonic anatomy. In 278 this study, we have reported a method, based purely on CTC data, to quantitatively describe the 279 complex morphology of the colon in terms of length, diameter, volume, and tortuosity. We have 280 examined these data across the total colon and in more detail within individual anatomical segments. 281 These data are intended to increase our understanding of this organ and thereby help the design of 282 future colonoscopy technology to accommodate the wide inter and intra-colon variations shown in 283 this study.

The mean total colonic length in this study averaged 185.0 cm in the supine orientation and 183.0 cm in the prone orientation. These are in close agreement with those found in the previous CTC study [20], however, they are significantly greater than the average reported in barium enama (up to 157 cm, for patient age range of 14-92 years) [15, 17] and in intraoperative studies (114 cm, for patient age range of 19-85 years) [18]. This discrepancy is likely to be a consequence of the population's age range and also the two-dimensional nature of these studies. Furthermore, it is possible that insufflation of the colon during CTC increases the length when compared to these studies.

This study demonstrates that the colonic segmental diameter changes from supine to prone orientation are influenced predominantly by intra-abdominal compression and pelvic motion. In a distended colon, the morphology of the majority of the colon is dictated primarily by changes in compressive effects, rather than by the effect of gravity. Our findings suggest that the prone position showed a higher frequency of high angles and consequently found to be more torturous than the supine position. We concluded that from these findings it would be beneficial to start device intubation in a prone position (for better rectal extension) and successively move the patient to the supine position for maximum extension of the ascending colon and cecum during the colonoscopyprocedure.

300 The descriptive metrics generated here agree with clinical reports of colonoscopy. The sigmoid 301 colon is typically considered to be a challenging segment and our metrics reveal this segment has the 302 smallest diameter but the second longest length. The angle distribution of this segment was also 303 negatively skewed (i.e. higher tortuosity), helping to explain the reported occurrences of loop formation, difficulty in passing the colonoscope, and patient discomfort during intubation found in 304 this segment [16, 30]. Normal adult human colons show considerable variations in length, radius, and 305 306 volume which would naturally modify the tortuosity of the colonic lumen. These factors along with 307 the pushing action exerted by the endoscopist during intubation and also the rigidity in the pushing 308 direction of the traditional colonoscope may contribute to excessive stress and stretch of bowel and mesenteries, and ultimately pain felt by the patient during an uncomfortable procedure. 309

310 The segmental diameters and lengths are very precious data for designers of robotic colonoscopies. 311 This study also defines a standardised metric that can be used to compare the tortuosity of colons between subjects and body orientations. Although the angle or tortuosity of the colon can be changed 312 313 under the pressure exerted by the scope or even body orientation, but it remains an important metric when designing a contactless robotic platforms or even atraumatic scopes. An example of this is 314 315 swimming or magnetic capsules, which are under development, and don't rely on contact with colonic 316 wall for their locomotion strategies. Our findings will help the community to optimise the length, diameter, and articulation parameters of new robotic platforms for an improved navigation and 317 318 locomotion strategies.

319 The geometry of colonoscopy devices (notably their diameter, segmental length and articulation range) are necessarily constrained and defined by the luminal environment of the colon. It is essential 320 that they are appropriately designed such that they can move within the torturous lumen without 321 322 imposing excessive force on the tissue (risking discomfort or colon damage) and support functionality 323 to image or take biopsies. Considering the conceptual design of a robotic system for colonoscopy requires guidance on factors such as the device diameter and ability to articulate. Referring to Fig. 5 324 325 and 6, it is evident that the rigid diameter of the robot should not be greater than 17 mm which is the 326 5th percentile of the sigmoid colon (the Olympus CF-140L adult video colonoscope features a 12.9 327 mm outer diameter tube). Furthermore, the body of the robot must be flexible enough, or articulate, to conform to the acute bends shown in Fig. 2. a significant number of which are $>90^{\circ}$. We used a span 328 329 size of 35 mm (which is the equivalent of total colonic average diameter in prone position) to analyze 330 the in-plane angle of the lumen centerline but this could readily be adapted for devices of different 331 lengths.

The results from this study give the research community a dataset defining engineering requirements for the design of colonoscopy devices. Furthermore, it provides a robust methodology with which to obtain these metrics. Thus the preliminary dataset presented here can be readily extended with complementary information and extended to cover different populations, age groups and disease types, helping to ensure that future colonoscopy systems are best designed to meet their clinical need.

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342 **6 References**

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