4 Authors: Shellie J. Radford^{1,2}, Prof Stuart A Taylor³, Dr Gordon W. Moran^{1,2} 5 6 **Corresponding Author:** Miss Shellie J Radford 7 Corresponding Author address: The NIHR Nottingham Biomedical Research Centre, Liver and Gastrointestinal Research Theme. Nottingham University Hospitals NHS Trust. 8 Corresponding Author Email contact: shellie.radford1@nottingham.ac.uk 9 10 11 Affiliations: 12 13 1. The University of Nottingham, School of Medicine, Translation Medical Sciences. 14 2. The Nottingham NIHR Biomedical Research Centre, Liver and Gastrointestinal 15 Research Theme. Nottingham University Hospitals NHS Trust. 16 3. Centre for medical imaging, University College London. 17 18 **Conflict of interest:** 19 SJR,- nil to declare 20 • GWM is in receipt of research funding from Janssen, Arla foods and Astra Zeneca. Dr 21 Moran is a consultant for Alimentiv. 22 SAT is consultant to Alimentiv and has share options in Motilent 23 • There are no competing interests relating to any authors of this paper. 24 25 Keywords: Crohn's Disease, Ultrasound,. 26 27 Total word count of main text: 1827 28 29 Number of tables: 1 30 Table 1: Comparison of imaging modalities when assessing small bowel Crohn's Disease 31 32 Number of Figures: 2 33 Figure 1: Distribution of NHS centres in the UK who responded to the BSG survey on the use of 34 ultrasound. 35 Figure 2: Confidence in clinical decision making when using ultrasound and MRE 36 assessment. 37 38 Funding: This study is undertaken as part of Doctoral study funded by the National Institute 39 for Health Research (NIHR) Applied Research Collaboration East Midlands (ARC EM). The 40 views expressed are those of the author(s) and not necessarily those of the NIHR or the 41 Department of Health and Social Care. 42 43 Author Contributions: 44 SJR- Survey Design, Data analysis, whole study write up. 45 SAT- Survey Design, Whole manuscript review GWM- Survey Design, Survey distribution, Whole Manuscript review. 46 47 48 Acknowledgements: 49 We would like to thank the members of the BSG IBD group for their contributions. We would 50 like to thank Jacqueline Campbell, Head of Committee services at the BSG for her help in distributing and collating responses to this survey. 51 52

Ultrasound use to assess Crohn's Disease in the UK: a survey of British Society of

Gastroenterology Inflammatory bowel disease group members.

1

2

3

1

2 Abstract:

3

4 Background

5 Data from the METRIC trial (PMID:29914843) has shown that small bowel ultrasound has

- 6 very good diagnostic accuracy for disease extent, presence and activity in Crohn's Disease
- 7 (CD), is well tolerated by patients and is cheaper when compared to MRI. However, Uptake
- 8 of ultrasound in the UK is limited

9 Methods

- 10 We designed and conducted an online survey to assess the current usage of ultrasound
- 11 throughout the UK. The survey was undertaken by BSG IBD group members between 9
- 12 June 2021 25 June 2021. Responses were anonymous, respondents were able to skip
- 13 questions.

14 Results

- 15 103 responses were included in the data analysis Responses came from 14 different regions
- 16 of the UK, from 66 individual NHS trusts. 103 respondents reported that they currently have
- 17 an MRI service for Crohn's disease, where only 31 had an ultrasound service. Numbers of
- 18 MRIs per month was reported as an average of 15, with a range of 3-75. The average
- 19 number of ultrasounds undertaken was reported as 8 per month, with a range of 0-50.
- 20 Average time for results to be reported for MRI scans was reported as between 4-6 weeks,
- 21 with a range of 2 days to 28 weeks. The average time for an ultrasound to be reported was
- stated as 1-4 weeks, with a range of 0-8 weeks. 26 respondents were 'extremely confident'
- when using MRI data to make clinical decisions, 5 were 'very confident' were somewhat
- 24 confident and 3 were not so confident. Only 6 respondents stated they would be extremely
- confident in using ultrasound to make clinical decisions, 17 people stated they would be veryconfident, 20 were somewhat confident, 15 not so confident and 15 not at all confident. Of
- those respondents who did not have access to an ultrasound service, 72 stated that they
- 28 would be interested in developing an ultrasound service.

29 Conclusion

- 30 There is an appetite for the uptake of ultrasound in the UK for assessment of CD, however
- 31 there remains a significant number of UK centres with little or no access to an ultrasound
- 32 service. There is a difference in the levels of confidence that clinicians have in using
- 33 ultrasound as a diagnostic tool in the UK. Further research is necessary to understand why
- 34 this is the case. Results from this survey will go on to inform our future work in developing an
- 35 implementation package for ultrasound in the UK in the NHS

36

- 37 38
- 39
- 40 41

- 1 2
- 3 Summary:
- 4 1. What is already known about this subject?

5 ultrasound is used widely in central Europe and Canada. Despite ultrasound being a quicker,

6 cheaper and more preferable test for patients, the uptake of ultrasound use in the UK is still

7 limited. The METRIC study has shown that ultrasound has comparable sensitivity and

8 specificity to MRE when detecting presence and extent of small bowel CD.

9 2. What are the new findings?

10 Nationally there are longer waiting times for MRE and ultrasounds assessments.

11 Gastroenterologists report that they are more confident in using MRE reports to make clinical

decisions than ultrasound reports, its is not yet clear why this is the case. The survey has

shown that there are some centre in the UK that are using ultrasound as part of their IBD
assessment, however there still remains many UK NHS centres who do not use ultrasound

15 but have indicated that they would wish to in the future.

16 3. How might it impact on clinical practice in the foreseeable future?

17 This survey is part of a programme of work being led by the NIHR Nottingham Biomedical

18 Research Centre. This programme of work will investigate aspects of existing ultrasound

=use in the UK, training needs of the IBD team, confidence in clinical decision making, of the
IBD team using ultrasound, cost effectiveness of an ultrasound pathway in IBD care and

21 stakeholder perceptions of the implementation of ultrasound in the NHS. Mixed methods

22 data will be collected and used to create an implementation package to support the

23 implementation of ultrasound nationally for the care of patients living with IBD.

24 25

1 Introduction:

2

3 Inflammatory Bowel Disease (IBD) refers to two conditions; Crohn's Disease (CD) and

4 Ulcerative Colitis, typically characterised by chronic inflammation of the gastrointestinal tract.

5 Disease distribution in CD varies with up to 70% of patients having small bowel

6 involvement.¹7

8 The incidence and prevalence of CD in Europe ranges from 0.5 to 10.6 cases per 100,000

9 person-years, and from 1.522 to 21312 cases per 100,000 persons respetively.² In the

10 United Kingdom (UK) it is estimated that there are 300,000 people affected by IBD, one of

- 11 the highest world-wide.³
- 12

The mean cost per patient-year during follow-up has been reported as £2971 (median £602 [180–2948]) for patients with CD, with an overall annual cost to the National Health Service (NHS) of up to £470 million.⁴ During the first five years following IBD diagnosis 50-75% of the budget is attributed to the use of biologic therapy.⁴

17

18 To ensure optimal long term clinical outcomes, current recommendations based on the

19 Selecting Therapeutic targets in IBD (STRIDE-II⁵) suggest using objective measures as

20 treatment targets, rather than symptom resolution. A wide array of biological therapies are

21 employed in treating IBD and objectively assessing treatment response has significantly

22 increased the projected IBD healthcare burden for the next decade.⁶ To ensure cost-

23 effective IBD practice, complex and expensive pharmacological interventions should be

- 24 targeted at patients most likely to benefit.⁷
- 25

Cross sectional imaging is used to diagnose and monitor disease activity in small bowel CD (SBCD).⁸ Magnetic Resonance Enterography (MRE) is often employed as a first modality in the UK for assessment and monitoring of SBCD.⁸ Waiting times for an NHS MRE may be up to 4 weeks or in some instances longer, and have increased due to the impact of the Covid-19 pandemic. Radiological reporting is then undertaken at a later date and may also add to delays. There is still a clinical need to find quicker, more tolerable and cheaper alternatives for monitoring patients with IBD.

33

34 Small bowel (enteric) ultrasound is an alternative to MRE, and has the potential to

35 significantly reduce waiting times, speed up clinical decision making and improve patient

- 36 experience and outcomes.⁹ ultrasound is widely used for assessing and monitoring IBD
- 37 internationally, and the METRIC^{10,11} trial has demonstrated its relative diagnostic accuracy in
- 38 comparison to MRE.
- 39

The NIHR-funded METRIC trial is the largest comparative diagnostic accuracy trial of MRE
and ultrasound in CD.¹⁰ The study reported that sensitivity for detecting small bowel disease

42 was 97% and 92% for MRE and ultrasound respectively. Specificity was 96% for MRE and

43 84% for ultrasound.¹⁰ These findings were concordant in both new diagnosis and suspected

44 relapse.^{10,11}

45

46 NHS tariff reports from 2021/2022 detail the cost for a MRE procedure with intravenous

- 47 contrast to be £162, with a reporting cost of £22. In comparison the cost of ultrasound is £51,
- 48 inclusive of reporting, hence making it a less costly and potentially more cost-effective

- 1 alternative. There is a large clinical need to correctly identify responders and non-
- 2 responders to therapy in a timely, cost effective and efficient manner,.^{7,12} However
- 3 ultrasound is not commonly used in the NHS, unlike in Central Europe and Canada.^{13,14}.
- Many authors report this is likely down to lack of available training,^{9,15–17} although questions 4
- 5 over high interobserver variation and suboptimal accuracy have dogged ultrasound for many
- 6 vears. The actual barriers to adoption of ultrasound in the NHS UK are to date speculative.
- 7 and remain largely unknown.
- 8

9 Methods:

10

11 We designed and conducted an online survey to assess the current usage of ultrasound

- 12 throughout the UK (Table 1). The survey was undertaken by BSG IBD group members
- 13 between 9 June 2021 - 25 June 2021. The BSG IBD group consists of Consultant and
- 14 Trainee gastroenterologists with a special interest in IBD and IBD specialist Nurses. There 15
- are 1410 members of the BSG IBD group, The survey was sent to all members on the 9th
- 16 and 22nd of June 2021, the survey was sent twice as the deadline for responses was
- 17 extended by a week. Responses were anonymous, respondents were able to skip questions 18 if they were unsure of the answers or if the question was not relevant to them (i.e. they do
- 19 not currently use ultrasound). The survey was accessible via online link, no reminders were
- 20 sent.
- 21
- 22 The guestionnaire comprised of 14 guestions. Questions were focused on the respondents 23 experiences of MRE and ultrasound use in relation to the clinical IBD care they deliver. We
- 24 asked respondents to report only on plain ultrasound examinations. We did not collect data
- 25 regarding other forms of ultrasound examination such as elastography or doppler. We
- 26 collected data relating to the regions of the UK where respondents work clinically, and their
- 27 opinions about whether they would like to use ultrasound for monitoring of IBD in the future if
- 28 they did not already do so.
- 29

30 **Results:**

31

32 There were 106 respondents, this is a response rate of 7.5%. there were 2 incomplete 33 forms, these were removed, and one international respondent, was also removed given the

- 34 UK focus of the survey. 103 responses were included in the data analysis.
- 35

36 Responses came from 14 different regions of the UK, from 66 individual NHS trusts. Figure 37 1 shows the distribution of the responding centers, showing those that currently use

- 38 ultrasound, those that would like to in the future and those that do not.
- 39
- 40 103 respondents reported that they currently have an MRI service for Crohn's disease,
- 41 where only 31 had access to ultrasound service. Of those respondents who did not have
- 42 access to an ultrasound service, 72 stated that they would be interested in developing an ultrasound service.
- 43 44
- 45 55 of respondents reported that they always use MRI when clinically appropriate, 39
- 46 reported they 'usually' utilised MRI, 8 stated sometimes and 1 person stated that they never
- 47 use MRI. 46 respondents reported that they never use ultrasound, 12 rarely use it, 22
- 48 sometimes with only 5 respondents usually using it, and 6 always using ultrasound.

1 2

3 4 The number of MRIs performed per month was reported as an average of 15, with a range of 3-75. The average number of ultrasounds undertaken was reported as 8 per month, with a range of 0-50. Average time from referral for results to be reported for MRI scans was reported as between 4-6 weeks, with a range of 2 days to 28 weeks. The average time for an ultrasound to be reported was stated as 1-4 weeks, with a range of 0-8 weeks.

6 7

5

8 30 respondents reported that they had access to both MRE and ultrasound. Not all

9 respondents completed all sections of the survey questionnaire. 9 different sites were

10 reported to have access to both MRE and ultrasound, with five of those being University

11 hospitals Trusts, and four NHS Foundation trusts. 21 respondents did not complete which

12 NHS trust they were currently employed by. 25 of respondents with access to both

13 modalities submitted data relating to waiting times; in these centers the average waiting time 14 from referral to report was reported as 4.6 weeks for MRE and 3.4 weeks for ultrasound.

15

16 26 respondents were 'extremely confident' when using MRI data to make clinical decisions,

17 5 were 'very confident' were somewhat confident and 3 were not so confident. Only 6

18 respondents stated they would be extremely confident in using ultrasound to make clinical

decisions, 17 people stated they would be very confident, 20 were somewhat confident, 15

20 not so confident and 15 not at all confident (Figure 2)

21

22 Discussion:23

24 MRE is the first line imaging modality used to accurately stage small bowel disease location,

complexity and activity in newly diagnosed CD. ^{10,18} MRE is also most commonly used to

26 measure disease response to biological therapies. However, once disease location and

phenotype are established, in many patients, there is an equipoise between MRE and small
bowel ultrasound in subsequent disease follow up and monitoring. SBUS has been shown to

bower ultrasound in subsequent disease follow up and monitoring. SBOS has been shown to
be equally accurate for evaluating enteric disease ^{30–35}, cheaper, quicker, better tolerated

30 and, most importantly, preferred by patients.^{10,19–22} Despite this, US is not widely

31 implemented for CD in the UK, for reasons we do not fully understand.

32

33 The treat-to-target paradigm present in IBD management guidelines is similar in other

34 chronic diseases.^{23–26} Management strategies in CD reflect a step-up paradigm, where

35 patients clinical symptoms in conjunction with markers of inflammation tend to guide

³⁶ investigation or medical intervention.^{27,28} Mucosal healing, defined by the absence of

37 ulcerations, is recommended as the therapeutic goal in clinical practice.^{5,8,29}

38

39 The equipment required is readily available in most hospitals. ultrasound could be a robust 40 alternative to more invasive and expensive imaging techniques. Besides being quick, well 41 tolerated, relatively inexpensive and readily available, ultrasound is reported and interpreted at the time of scanning and allows for early clinical decision-making in routine IBD care.9,36 42 Importantly, the METRIC¹⁰ study found no major difference between MRE and ultrasound in 43 44 terms of therapeutic decision-making, indicating that the differences in accuracy between the 45 two tests do not translate to differences in patient management. Both tests had a similar 46 level of concordance compared to the reference standard in terms of therapeutic decisions 47 (77% for MRE and 78% for ultrasound). This sub-study on decision-making, although well1 designed, was a paper-based exercise with small numbers; further evidence is required to

- 2 ensure these results reflect real-world practice.
- 3

4 The results from the METRIC²¹ study were used to underpin a cost-effectiveness analysis

- 5 showing that ultrasound was more cost-effective than MRE in the management of suspected
- 6 relapse; it was estimated that ultrasound saves the NHS an average of £299 per patient,
- 7 with a negligible –0.0001 (–0.013 to 0.011) impact on QALYs. There is scarce empirical
- 8 evidence presenting comprehensive data relating to cost or cost effectiveness of
- 9 ultrasound.⁹ In the METRIC study ultrasound was considered highly acceptable by patients
- 10 when compared with MRE.²² ultrasound is often seen as having limited clinical utility due to
- 11 operator dependence.³⁶ However, every diagnostic technique, including endoscopy, has a
- degree of subjectivity and operator dependence and this criticism is perhaps more reflective
- 13 of a previous lack of identifiable international performance and training standards.³⁶ The 14 training needs for gastroenterologists are similar to those of radiologists as set out in the
- 14 training needs for gastroenterologists are similar to those of radiologists as set out in the 15 ECCO-ESGAR guidelines¹², this can be time consuming, even when supported by
- 16 abdominal radiology specialists and in partnership with radiology departments.^{9,16,36} There is
- 17 no current literature relating to any other IBD healthcare worker undertaking ultrasound
- 18 training.

19

20 Conclusions:

- 21
- 22 This survey was the first step in a project of further work to investigate patient or HCPs
- 23 preferences for service delivery for imaging for assessment and monitoring imaging in IBD.
- 24 ultrasound has been shown to be similar in accuracy to MRE in detecting presence SBCD.
- 25 ultrasound is reported as quicker, more acceptable to patients and potentially safer when
- compared to MRE. ultrasound is used widely in central Europe, Canada and some parts of
- 27 the USA, but has not been as widely embraced in the UK. It would seem prudent to
- 28 investigate broader stakeholder perceptions of the use of ultrasound to better understand
- 29 perceived or potential barriers and enablers to ultrasound implementation in the world-wide
- 30 healthcare systems and recognise and manage preferences for future service delivery.
- 31
- 32
- 33
- 34
- 35
- 36 37
- 37 38
- 39
- 40
- 41
- 42 43
- 44
- 45
- 46
- 46 47
- 48

Refe	rences:
1.	Jones, G. R. <i>et al.</i> IBD prevalence in Lothian, Scotland, derived by capture-recapture
-	methodology. <i>Gut</i> 68 , 1953–1960 (2019).
2.	Burisch, J., Jess, T., Martinato, M. & Lakatos, P. L. The burden of inflammatory bowel
	disease in Europe. Journal of Crohn's and Colitis (2013)
•	doi:10.1016/j.crohns.2013.01.010.
3.	Chu, I. P. C., Moran, G. W. & Card, I. R. The pattern of underlying cause of death in
	patients with Inflammatory Bowel Disease in England: A record linkage study. J.
	Cronn's Collitis 578–585 (2017) doi:10.1093/ecco-jcc/jjw192.
4.	Burisch, J. et al. Health-care costs of inflammatory bowel disease in a pan-European,
	study, Langet Castroonterol, Langetol, F. 454, 464 (2020)
E	Sludy. Luncel Gustroenleroi. Hepatol. 5 , 454–464 (2020).
5.	Inflammatory Rowel Disease (STRIDE) Initiative of the International Organization for
	the Study of IBD (IOIBD): Determining Therapeutic Goals for Treat-to-Target
	strategies in IBD. Gastroenterology 0 .
6.	Bouguen, G. <i>et al.</i> Treat to Target: A Proposed New Paradigm for the Management of
•	Crohn's Disease. <i>Clinical Gastroenterology and Hepatology</i> vol. 13 1042-1050.e2
	(2015).
7.	Kennedy, N. A. <i>et al.</i> Predictors of anti-TNF treatment failure in anti-TNF-naive
	patients with active luminal Crohn's disease: a prospective, multicentre, cohort study.
	Lancet Gastroenterol. Hepatol. 4 , 341–353 (2019).
8.	Peyrin-Biroulet, L. et al. Selecting Therapeutic Targets in Inflammatory Bowel Disease
	(STRIDE): Determining Therapeutic Goals for Treat-to-Target. Am. J. Gastroenterol.
	110 , 1324–1338 (2015).
9.	Allocca, M., Furfaro, F., Fiorino, G., Peyrin-Biroulet, L. & Danese, S. Point-of-Care
	Ultrasound in Inflammatory Bowel Disease. J. Crohn's Colitis 2020, 1–9 (2020).
10.	Taylor, S. A. et al. Diagnostic accuracy of magnetic resonance enterography and small
	bowel ultrasound for the extent and activity of newly diagnosed and relapsed Crohn's
	disease (METRIC): a multicentre trial. <i>Lancet Gastroenterol. Hepatol.</i> 3 , 548–558
11.	Bhatnagar, G. <i>et al.</i> Observer agreement for small bowel ultrasound in Crohn's
	disease: results from the IVIETRIC trial. Abdom. Rddiol. (2020) doi:10.1007/S00261-
17	UZU-UZ4UJ-W. Maasar (<i>at al</i> ECCO_ESGAR Guidaling for Diagnostic Assocsment in IPD Part 1:
12.	Initial diagnosis monitoring of known IRD, detection of complications 1. Crahn's
	Colitic 12 144–164 (2010)
13	Panes 1 et al Imaging techniques for assessment of inflammatory howel disease:
15.	Ioint ECCO and ESGAR evidence-based consensus guidelines / Crohn's Colitis 7 556–
	585 (2013).
14.	Panés, J. <i>et al.</i> Systematic review: The use of ultrasonography. computed tomography
	and magnetic resonance imaging for the diagnosis, assessment of activity and
	abdominal complications of Crohn's disease. Alimentary Pharmacology and
	Therapeutics vol. 34 125–145 (2011).
15.	Novak, K. L. <i>et al.</i> Point of Care Ultrasound Accurately Distinguishes Inflammatory

1 from Noninflammatory Disease in Patients Presenting with Abdominal Pain and 2 Diarrhea. Can. J. Gastroenterol. Hepatol. 2016, (2016). 3 16. Grunshaw, N. D. Initial experience of a rapid-access ultrasound imaging service for 4 inflammatory bowel disease. *Gastrointest. Nurs.* **17**, 42–48 (2019). 5 Wang, I. Kamm, M., Wong, D. et al. P260 Point of Care Ultrasound (POCUS) when 17. 6 performed by gastroenterologists with 200 supervised scans is accurate and clinically 7 useful for patients with Crohn's disease | Journal of Crohn's and Colitis | Oxford 8 Academic. Journal of Chron's and Colitis s232 https://academic.oup.com/ecco-9 jcc/article/12/supplement_1/S232/4807565 (2018). 10 18. Turner, D. et al. STRIDE-II: An Update on the Selecting Therapeutic Targets in 11 Inflammatory Bowel Disease (STRIDE) Initiative of the International Organization for 12 the Study of IBD (IOIBD): Determining Therapeutic Goals for Treat-to-Target 13 strategies in IBD. Gastroenterology 160, 1570–1583 (2021). 14 19. Miles, A. et al. Magnetic resonance enterography, small bowel ultrasound and 15 colonoscopy to diagnose and stage Crohn's disease: patient acceptability and 16 perceived burden. Eur. Radiol. (2019) doi:10.1007/s00330-018-5661-2. 17 20. Taylor, S. A. et al. Magnetic resonance enterography compared with ultrasonography 18 in newly diagnosed and relapsing crohn's disease patients: The METRIC diagnostic 19 accuracy study. Health Technol. Assess. (Rockv). 23, vii–161 (2019). 20 Allocca, M. et al. Comparative Accuracy of Bowel Ultrasound Versus Magnetic 21. 21 Resonance Enterography in Combination With Colonoscopy in Assessing Crohn's 22 Disease and Guiding Clinical Decision-making. J. Crohn's Colitis 1280–1287 (2018) 23 doi:10.1093/ecco-jcc/jjy093. 24 22. Evans, R. et al. Patient experience and perceived acceptability of whole-body 25 magnetic resonance imaging for staging colorectal and lung cancer compared with 26 current staging scans: A qualitative study. BMJ Open 7, e016391 (2017). 27 23. H.S., Y. et al. Update on the management and treatment of hepatitis C virus infection: 28 Recommendations from the Department of Veterans Affairs Hepatitis C Resource 29 Center Program and the National Hepatitis C Program Office. Am. J. Gastroenterol. 30 **107**, 669–689 (2012). 31 L.H., K. How and when to use biologics in psoriasis. J. Drugs Dermatology 9, s106-24. 32 s117 (2010). 33 25. P., H. The pathology induced by highly active antiretroviral therapy against human 34 immunodeficiency cirus: An update. Curr. Med. Chem. 13, 3121-3132 (2006). 35 26. Molodecky, N. A. et al. Increasing incidence and prevalence of the inflammatory 36 bowel diseases with time, based on systematic review. Gastroenterology 142, 46-54 37 e42; quiz e30 (2012). 38 27. Novak, K. et al. Clinic-based Point of Care Transabdominal Ultrasound for Monitoring 39 Crohn's Disease: Impact on Clinical Decision Making. J. Crohns. Colitis (2015) 40 doi:10.1093/ecco-jcc/jjv105. 41 Brazilian, S. G. of I. B. D. Consensus guidelines for the management of inflammatory 28. 42 bowel disease. Arq. Gastroenterol. 47, 313–25 (2010). 43 Lamb, C. A. et al. British Society of Gastroenterology consensus guidelines on the 29. 44 management of inflammatory bowel disease in adults. Gut (2019) doi:10.1136/gutjnl-45 2019-318484. 46 30. Horsthuis, K., Bipat, S., Bennink, R. J. & Stoker, J. Inflammatory bowel disease 47 diagnosed with US, MR, scintigraphy, and CT: Meta-analysis of prospective studies.

1		Radiology 247 , 64–79 (2008).
2	31.	Dong, J. et al. Ultrasound as a diagnostic tool in detecting active Crohn's disease: A
3		meta-analysis of prospective studies. <i>Eur. Radiol.</i> 24, 26–33 (2014).
4	32.	Puylaert, C. A. J., Tielbeek, J. A. W., Bipat, S. & Stoker, J. Grading of Crohn's disease
5		activity using CT, MRI, US and scintigraphy: a meta-analysis. Eur. Radiol. (2015)
6		doi:10.1007/s00330-015-3737-9.
7	33.	Ahmed, O., Rodrigues, D. M. & Nguyen, G. C. Magnetic Resonance Imaging of the
8		Small Bowel in Crohn's Disease: A Systematic Review and Meta-Analysis. Canadian
9		Journal of Gastroenterology and Hepatology vol. 2016 (2016).
10	34.	Greenup, A. J., Bressler, B. & Rosenfeld, G. Medical imaging in small bowel Crohn's
11		disease - Computer tomography enterography, magnetic resonance enterography,
12		and ultrasound: 'Which one is the best for what''". <i>Inflammatory Bowel Diseases</i> vol.
13		22 1246–1261 (2016).
14	35.	Alshammari, M. T. et al. Diagnostic Accuracy of Non-Invasive Imaging for Detection of
15		Colonic Inflammation in Patients with Inflammatory Bowel Disease: A Systematic
16		Review and Meta-Analysis. <i>Diagnostics 2021, Vol. 11, Page 1926</i> 11 , 1926 (2021).
17	36.	Bryant, R. V. et al. Gastrointestinal ultrasound in inflammatory bowel disease: An
18		underused resource with potential paradigm-changing application. Gut vol. 67 973–
19		985 (2018).
20		
21		
22		