



Knight, S. R., Pathak, S., Christie, A., Jones, L., Rees, J., Davies, H., Wilson, M. S. J., Vaughan-Shaw, P., Roberts, K., Toogood, G., Harrison, E. M., & Taylor, M. A. (2019). Use of a modified Delphi approach to develop research priorities in HPB surgery across the United Kingdom. *HPB*, *21*(11). Advance online publication. https://doi.org/10.1016/j.hpb.2019.03.352

Peer reviewed version

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ORIGINAL ARTICLE

Use of a modified Delphi approach to develop research priorities in HPB surgery across the United Kingdom

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Abstract

Background: Research prioritisation can help identify clinically relevant questions and encourage highquality, patient-centred research. Delphi methodology aims to develop consensus opinion within a group of experts, with recent Delphi projects helping to define the research agenda and funding within several medical and surgical specialties.

Methods: All members of the Association of Upper Gastrointestinal Surgeons (AUGIS) were asked to submit clinical research questions using an online survey (Phase 1). Two consecutive rounds of Delphi prioritisation by multidisciplinary HPB healthcare professionals (Phase 2) were undertaken to establish a final list of the most highly prioritised research questions. A multidisciplinary steering committee analysed the results of each phase.

Results: Ninety-three HPB-focussed questions were identified in Phase 1, with thirty-seven questions of sufficient priority to enter a further prioritisation round. A final group of 11 questions considered highest priority were identified. The most highly ranked research questions related to treatment pathways, operative strategies and the impact of HPB procedures on quality of life, particularly for malignant disease.

Conclusion: Expert consensus has identified research priorities within the UK HPB surgical community over the coming years. Funding applications, to establish well-designed, high quality collaborative research are now required to address these questions.

Received 15 January 2019; accepted 11 March 2019

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Introduction

Methodological and practical difficulties present unique challenges to the surgical research community,¹ with research quality historically variable and often poor.^{1,2} Integration of research

This study was presented at the 21st Annual AUGIS Conference, September 2018, Edinburgh, UK.

and practice through a collaborative approach can often overcome such challenges³ and enhance research impact, such as the Dutch Pancreatic Cancer Group in hepato-pancreato-biliary (HPB) surgery.^{4–6}

Identifying a national consensus on research priorities provides the opportunity to increase value and reduce waste in HPB research.⁷ Furthermore, the engagement of stakeholders in

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research encourages the development of clinically relevant research priorities and the potential for national collaboration in HPB research. Using structured prioritisation methods to achieve consensus can maximise relevance, guide funding bodies and channel resources. The aim of this work is to identify a list of highly prioritised clinical research questions in HPB surgery using a modified Delphi approach to guide future research and funding bodies.

Methods

The modified Delphi methodology can be utilised to produce consensus within a group of experts by interaction to reduce individual bias. These methods have previously identified research questions in orthopaedic⁸ and colorectal surgery,⁹ with outputs frequently receiving pump-priming or significant funding following identification.^{10–12} However, to the authors knowledge, clinical priority setting in HPB surgery is yet to be performed.

A modified Delphi approach was performed with two distinct phases, including two rounds of prioritisation (Fig. 1) using methodology previously described across a broad range of medical topics.^{9,13,14} Throughout the process, stakeholders were encouraged to consider prioritisation responses based on disease



Figure 1 The modified Delphi process used to prioritise research questions in hepato-pancreato-biliary surgery

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burden, clinical relevance and answerability – factors identified as facilitators of high-impact research.¹⁵ During each round, any incomplete questionnaires were excluded prior to analysis.

Phase one

Healthcare professionals within the AUGIS membership were asked to submit potential research questions across upper gastrointestinal and HPB surgery via an online survey. This was open to both medical professionals and multidisciplinary team members, with no limit on the number of research questions an individual could submit. The survey was open for at least six weeks, with at least one reminder email was sent during this period and through social media platforms to increase engagement with the process.

All submitted questions were then grouped into four categories independently (hepato-pancreato-biliary, malignant upper gastrointestinal surgery, benign upper gastrointestinal surgery or bariatric surgery), with any disagreements resolved by consensus. HPB-focussed questions were then screened, with duplicate questions removed and similar questions streamlined following discussion by the steering group. Where questions related to an identical clinical research problem, questions were combined by group consensus. Where wording of questions was altered, agreement was reached within the steering group and care was taken not to alter the meaning of original questions posed.

Due to the separate sub-speciality interest within clinical practice, questions relating to the other three categories (malignant upper gastrointestinal surgery, benign upper gastrointestinal surgery and bariatric surgery) were each taken forward by individually selected and speciality-orientated steering groups.

Steering committee

The steering committee for the Delphi process contained two surgical registrars (SK, SP), a cancer nurse specialist (LJ), patient and public representation (HD), a medical oncologist with specialist interest in HPB disease (AC) and three consultant HPB surgeons (JR, EMH, MT). Representation from within the Association of Upper Gastrointestinal Surgeons (AUGIS) committee (Education, Training and Research Lead; MT) was also present.

Phase two

The collated list of research questions was then prioritised by attendees at the Great Britain and Ireland Hepato-Pancreato-Biliary Association (GBIHPBA) conference, March 2018, using printed questionnaires (*Prioritisation Round 1*). Healthcare professionals with a speciality interest in HPB surgery were asked to prioritise each individual research question based on their own opinion using a Likert scale (1 – low priority, 3 – neutral, 5 – high priority), in accordance with similar Delphi processes.^{8,9,13,14}

Scores were reviewed with a 'cut-off' point determined based on mean priority score and steering group consensus to select those questions to move through to the final round. Heterogeneity of responses was also examined by the steering committee to ensure 'block' voting was not performed. To ensure higher-level evidence did not already exist, a literature search of each remaining question was conducted prior to inclusion.

A second round of prioritisation was then performed by the GBIHPBA membership via a further online survey, with healthcare professionals asked to prioritise each question using the same Likert scale as the previous round. Again, respondents were given a minimum of six weeks with reminders sent through email and social media during this period. Results were reviewed by the steering committee to identify a final group of highly prioritised questions, with consensus reached by analysing the mean priority score and proportion of high priority responses each question received.

Results

A total of 427 research questions were produced by 140 respondents, representing 48% of the membership (Fig. 2). Subspecialisation in HPB surgery was declared by 47 (34%). Overall 153 (36%) research questions were HPB-focused. A summary of question topic in each round of the prioritisation exercise is shown in Table 1.

After duplicated and similar questions had been combined, 93 questions were moved forward into round 2. These research questions were prioritised by 44 respondents, representing 44% of the conference attendees, with a further four responses excluded due to incompleteness. Assessment of ranking heterogeneity excluded one further response due the possible presence of 'block' ranking (only priority ranks of 1 or 5 present). Following steering committee review, a consensus was reached regarding a cut-off value (mean priority score of \geq 3.0) and a total of 37 questions were brought forward into the last round (Supplementary Table 1).

In the final round 42 respondents prioritised these 37 questions, with three excluded due to incompleteness. Following steering committee review of responses, a consensus was again reached regarding cut-off value (questions receiving >50% high priority ranking (grade 4 or 5) in combination with a mean priority score of \geq 3.5; and <20% low priority ranking (1 or 2); Supplementary Table 2) to produce 11 final highly prioritised questions (Box 1). Across the three rounds consultant representation remained above 50% and group demographic remained similar (Fig. 3).

Discussion

A list of 11 high priority research topics in HPB surgery have been derived through a modified Delphi process.

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Figure 2 Overview of responses during modified Delphi process

A high proportion of these topics related to malignant disease, particularly regarding improving outcomes in pancreatic cancer and the operative management of colorectal cancer liver metastases. This is expected, particularly as they represent a high burden of disease,^{16,17} significant reduction in patient's quality of life and potential for improvement in survival outcomes.^{16,18} Furthermore there is considerable debate regarding these diseases if one considers the potential roles of neoadjuvant therapy or pathways to surgery among patients with pancreatic cancer and of two stage liver resection (Associating Liver Partition and Portal vein Ligation for Staged hepatectomy (ALPPS) in particular). In the case of pancreatic cancer, early stage disease detection and the development of new treatment strategies have previously been identified as priorities eight years ago.¹⁹ Progress within these domains has since been made, however they may remain high priorities due to difficulties in translating basic research to bedside²⁰ and persistent complexities within surgical research.21

A national quality improvement programme for iatrogenic common bile duct injury was sufficiently prioritised for

consideration in the final group of questions (Supplementary Table 2), however nearly a quarter of respondents (24%) ranked this question as low priority (grade 1 or 2). Due to priority rank heterogeneity and unanimous concern within the steering committee regarding difficulties in feasibility of delivering change at a national level, this question was not included within the final list. Thus, this Delphi process highlights areas where further review or data is required before those areas are the focus of study/improvement. All final round questions have been published here for transparency and to acknowledge that a proportion of questions are linked by topic or clinical management algorithms. In addition, differences in patient prioritisation may exist and as academic advances are achieved they may develop higher priority in the coming years. The authors also acknowledge that current ongoing studies, of which current progress and results are unknown, may answer some of the highranked questions. It is hoped this process will support any conclusions regarding the clinical significance of their findings.

Greater representation of benign and/or complex biliary disease may have been anticipated, particularly as it represents a

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Table 1 Question topics included in each round

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Торіс	Submitted in Phase 1	Collated questions in Round 2 voting	Highest scores for Round 3 voting	Final list of prioritisation research questions
Total	153 ^a	93	37	11
Cancer	49	31	17	5
Benign	21	13	3	-
Surgical technique	19	13	3	1
Complications	13	8	1	-
Prehabilitation	9	5	2	1
Screening	8	4	2	1
Liver transplant	3	3	2	-
Perioperative	5	3	1	-
Chronic pancreatitis	4	2	2	1
Palliative care	4	2	-	1
Education	1	1	_	_
Miscellaneous	17	8	4	1

^a Overall 427 questions were produced in Phase 1, of which 153 were hepato-pancreato-biliary (HPB) focused.

significant disease burden and associated clinical workload.²² Biliary sub-speciality interest was well represented across each round (data not shown) and therefore this is unlikely to explain their absence in the final set of topics for suggested review. It is possible that questions relating to biliary disease will be prioritised in the upper gastrointestinal surgery Delphi exercise run in parallel to this study, or that bias exists within the UK system towards cancer-based questions due to the focus on development of cancer care centres.

Throughout the prioritisation process, clinicians have been integral to identifying critical priorities in the surgical management of HPB disease, avoiding potential mismatches between those questions clinicians wish to be addressed in research and those questions that researchers choose to investigate.^{7,23,24}

The use of a large cohort of healthcare professionals through the AUGIS and GBIHPBA membership has reduced reporter bias and set priorities using a large multi-disciplinary cohort containing healthcare professionals representing both clinical and academic foci. While a variety of methods have been described to determine consensus, there is no agreement as to the best approach.²⁵ Therefore the 'certain level of agreement' method was adopted as this is the most commonly used methodology in consensus-based methodology.

Low member response rates could be highlighted as a limitation to this study, particularly as responses were received from less than half the membership in each round (47.6, 33.6 and 32% respectively). However, similar studies using Delphi methodology have demonstrated lower response rates, with many quoting

Pancreas

- 1. How do you identify patients with resectable pancreatic cancer who would benefit from neoadjuvant chemotherapy and/or radiotherapy? Is there a role for surgical resection in patients with pancreatic cancer down staged by neoadjuvant chemotherapy and/or radiotherapy? In those patients with locally advanced disease, in whom should resection be offered?
- 2. Which blood and urine markers can identify early stage pancreatic cancer?
- 3. Does fast track surgery for pancreatic carcinoma improve overall outcome in terms of reduced morbidity and improved survival? Which patient cohort is suitable for fast-track surgery and in whom should biliary drainage be performed preoperatively?
- 4. What is the natural history of intraductal papillary mucinous neoplasms? How should patients with detected lesions be managed?
- 5. In patients at high risk of pancreatic cancer, either hereditary or chronic pancreatitis, what is the optimal radiological surveillance (modality and frequency)?

Liver

- 6. In patients with colorectal cancer and synchronous liver metastases, which patients benefit from combined primary site surgery and metastasectomy?
- 7. Is there a role for genetic or molecular prognostic/ predictive markers in selecting patients for resection of colorectal liver metastases? What is the role of biological agents in colorectal liver metastases and what prognostic markers improve outcomes?
- 8. For patients with bilobar colorectal liver metastases in whom a single stage resection is not feasible, which technique is superior with regard to oncological outcomes and patient-reported outcomes – ALPPS or conventional 2-stage hepatectomy with portal venous embolization?

Cross-speciality

- 9. Does a prehabilitation programme, including the optimisation of nutrition, improve outcomes of HPB surgery?
- 10. What is the impact of the cancer nurse specialist on HPB patients, their clinical and patient-reported outcomes, and their carers?
- 11. How do we maximise quality of life and survival for patients with unresectable HPB cancer? What is the role of nutrition in this patient group?

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Figure 3 Distribution of responder demographics through three rounds of prioritisation by clinical grade

around 10-18%.^{9,14} This suggests adequate engagement in the process and represents the largest collation of expert opinion on HPB research priorities to date.

The use of paper questionnaires could have introduced the potential for respondents to readily identify questions of particular interest in block voting during Phase 2, particularly as questions were presented in identical order. However the assessment of rank heterogeneity identified only one response where this may have occurred, suggesting the introduction of bias remained limited.

This study also highlights the process of developing a national consensus statement for HPB research priorities. The final priority list may differ from other geographical and cultural settings, however this process could be utilised by other national and international groups to perform further Delphi prioritisation exercises specific to unanswered research questions prominent within their surgical community.

Within the steering group a patient representative reviewed questions during phase two to ensure patient relevance and interpretation of question structure. However, exploration of patient perceptions and validation of the final question group with a larger cohort is required to ensure patient-centred research is achieved.^{7,25} The development of clinical vignettes through patient and public involvement may also help further guide funding bodies, similar to the work performed by McNair *et al* following the similar process performed for colorectal surgery.²⁶

The questions will be disseminated to funding bodies and encourage championing of individual research questions by AUGIS members. Meanwhile, the newly appointed AUGIS pancreatic cancer research lead, a joint initiative with Pancreatic Cancer UK and Pancreatic Cancer Research Fund, will provide a platform on which such questions can be answered. It is anticipated the output from the current study will set the research agenda in HPB surgery and hope it will further encourage contribution to planned research and collaboration between centres over the coming years. In summary this Delphi process has set high research priorities in HPB surgery, which it is hoped will align research agendas and principle users. Malignant disease was highly prioritised, relating to treatment pathways and specific operative strategies, together with the impact of HPB procedures on quality of life.

Acknowledgements

The HPB Delphi research steering committee would like to thank AUGIS, GBIHPBA and, in particular, Nichola Bartlett for their support and administrative assistance throughout the Delphi process.

Funding sources

None.

Conflicts of interest

None to declare.

References

- Ergina PL, Cook JA, Blazeby JM, Boutron I, Clavien P-A, Reeves BC et al. (2009 Sep 26) Challenges in evaluating surgical innovation. *Lancet* 374:1097–1104.
- Solomon MJ, McLeod RS. (1993 Jan) Clinical studies in surgical journals – have we improved? Dis Colon Rectum 36:43–48.
- Barkun JS, Aronson JK, Feldman LS, Maddern GJ, Strasberg SM, Balliol Collaboration *et al.* (2009 Sep 26) Evaluation and stages of surgical innovations. *Lancet* 374:1089–1096.
- Rijssen LB van, Koerkamp BG, Zwart MJ, Bonsing BA, Bosscha K, Dam RM van *et al.* (2017 Oct 1) Nationwide prospective audit of pancreatic surgery: design, accuracy, and outcomes of the Dutch Pancreatic Cancer Audit. *HPB* 19:919–926.
- van der Geest LGM, van Eijck CHJ, Groot Koerkamp B, Lemmens VEPP, Busch OR, Vissers PAJ et al. (2018 Sep 6) Trends in treatment and survival of patients with nonresected, nonmetastatic pancreatic cancer: a population-based study. *Cancer Med* 7:4943–4951.
- de Rooij T, van Hilst J, van Santvoort H, Boerma D, van den Boezem P, Daams F et al. (2019 Jan) Minimally invasive versus open distal pancreatectomy (LEOPARD): a multicenter patient-blinded randomized controlled trial. *Ann Surg* 269:2–9.
- Chalmers I, Bracken MB, Djulbegovic B, Garattini S, Grant J, Gülmezoglu AM *et al.* (2014 Jan 11) How to increase value and reduce waste when research priorities are set. *Lancet* 383:156–165.
- Eubank BH, Mohtadi NG, Lafave MR, Wiley JP, Bois AJ, Boorman RS et al. (2016 20) Using the modified Delphi method to establish clinical consensus for the diagnosis and treatment of patients with rotator cuff pathology. *BMC Med Res Methodol* 16:56.
- Tiernan J, Cook A, Geh I, George B, Magill L, Northover J et al. (2014 Dec) Use of a modified delphi approach to develop research priorities for the association of coloproctology of Great Britain and Ireland. *Colorectal Dis* 16:965–970.
- Trial Detail UK Clinical Trial Gateway [Internet].(2019) [cited 2019 Mar 2]. Available from: https://ukctg.nihr.ac.uk/trials/trial-details/trialdetails?trialNumber=ISRCTN17573805.
- A Trial Looking at Surgery or Different Types of Radiotherapy for Rectal Cancer (STAR-TREC) [Internet]. (2017). Cancer Research UK [cited 2019 Mar 2]. Available from: https://www.cancerresearchuk.org/aboutcancer/find-a-clinical-trial/a-trial-looking-at-surgery-or-different-typesof-radiotherapy-for-rectal-cancer-star-trec.

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- ISRCTN ISRCTN52352431. (2019) Does Intravenous Lidocaine Speed up Gut Recovery after Large Bowel Surgery? [Internet]. [cited 2019 Mar 2]. Available from: http://www.isrctn.com/ISRCTN52352431? q=&filters=conditionCategory:Digestive%20System&sort=&offset=9& totalResults=602&page=1&pageSize=10&searchType=basic-search.
- **13.** Stewart RJ, Caird J, Oliver K, Oliver S. (2011 Dec) Patients' and clinicians' research priorities. *Health Expect* 14:439–448.
- Burt CG, Cima RR, Koltun WA, Littlejohn CE, Ricciardi R, Temple LK et al. (2009 May) Developing a research agenda for the American Society of Colon and Rectal Surgeons: results of a delphi approach. *Dis Colon Rectum* 52:898–905.
- Burden of Disease and Research Funding [Internet]. Articles, Reports and Web Pages.(2018) [cited 2018 Sep 14]. Available from: http://rm21. typepad.com/blog/2012/07/burden-of-disease.html.
- Mohammed A, Janakiram NB, Madka V, Li M, Asch AS, Rao CV. (2018) Current challenges and opportunities for chemoprevention of pancreatic cancer. *Curr Med Chem* 25:2535–2544.
- 17. Engstrand J, Nilsson H, Strömberg C, Jonas E, Freedman J. (2018 Jan 15) Colorectal cancer liver metastases – a population-based study on incidence, management and survival. *BMC Cancer* [cited 2018 Sep 14]. Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/ PMC5769309/.
- Hackl C, Neumann P, Gerken M, Loss M, Klinkhammer-Schalke M, Schlitt HJ. (2014 Nov 4) Treatment of colorectal liver metastases in Germany: a ten-year population-based analysis of 5772 cases of primary colorectal adenocarcinoma. *BMC Cancer* 14:810.
- Robotin MC, Jones SC, Biankin AV, Waters L, Iverson D, Gooden H et al. (2010 May) Defining research priorities for pancreatic cancer in

Australia: results of a consensus development process. *Cancer Causes Control* 21:729–736.

- 20. Ma Y, Wu Q, Li X, Gu X, Xu J, Yang J. (2016 Dec) Pancreatic cancer: from bench to bedside. *Ann Transl Med* [cited 2018 Sep 16];4(23). Available from: https://www.ncbi.nlm.nih.gov/pmc/articles/ PMC5220038/.
- **21.** McCulloch P. (2011 Jul 8) How to improve surgical research. *BMJ* 343: d4121.
- Stinton LM, Shaffer EA. (2012 Apr) Epidemiology of gallbladder disease: cholelithiasis and cancer. *Gut Liver* 6:172–187.
- Liberati A. (2011 Nov 19) Need to realign patient-oriented and commercial and academic research. *Lancet* 378:1777–1778.
- Tallon D, Chard J, Dieppe P. (2000 Jun 10) Relation between agendas of the research community and the research consumer. *Lancet* 355: 2037–2040.
- 25. Stewart D, Gibson-Smith K, MacLure K, Mair A, Alonso A, Codina C et al. (2017) A modified Delphi study to determine the level of consensus across the European Union on the structures, processes and desired outcomes of the management of polypharmacy in older people. *PLoS One* 12:e0188348.
- 26. McNair AGK, Heywood N, Tiernan J, Verjee A, Bach SP, Fearnhead NS et al. (2017 Jan) A national patient and public colorectal research agenda: integration of consumer perspectives in bowel disease through early consultation. *Colorectal Dis* 19:075–085.

Appendix A. Supplementary data

Supplementary data to this article can be found online at https://doi.org/10. 1016/j.hpb.2019.03.352.

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