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Research prioritisation exercises related to the care of children and young people with life-limiting conditions, their parents, and all those who care for them: a systematic scoping review

ABSTRACT

Background: In planning high quality research in any aspect of care for children and young people with life-limiting conditions it is important to prioritise resources in the most appropriate areas.

Aim: To map research priorities identified from existing research prioritisation exercises relevant to infants, children, and young people with life-limiting conditions, in order to inform future research.

Design: We undertook a systematic scoping review to identify existing research prioritisation exercises; the protocol is publicly available on the project website.

Data sources: The bibliographic databases ASSIA, CINAHL, MEDLINE/MEDLINE In Process and Embase were searched from 2000. Relevant reference lists and websites were hand searched. Included were any consultations aimed at identifying research for the benefit of neonates, infants, children and/or young people (birth to age 25 years) with life-limiting, -threatening or -shortening conditions; their family, parents, carers; and/or the professional staff caring for them.

Results: Twenty four research prioritisation exercises met the inclusion criteria, from which 279 research questions or priority areas for health research were identified. The priorities were iteratively mapped onto an evolving framework, informed by WHO classifications. This resulted in identification of 16 topic areas, 55 sub-topics and 12 sub-sub-topics.

Conclusions: There are numerous similar and overlapping research prioritisation exercises related to children and young people with life-limiting conditions. By mapping existing research priorities in the context in which they were set, we highlight areas to focus research efforts on. Further priority setting is not required at this time unless devoted to ascertaining families’ perspectives.

Keywords: Delphi Technique; Consensus; infant; infant, newborn; child; Adolescent; Palliative Care; Humans.

What is already known about the topic?

- There is limited high quality research in many aspects of care for infants, children and young people with life-limiting conditions.
- It is important to minimise waste in research and maximise use of limited resources.
• A range of research prioritisation exercises for a variety of aspects of care or conditions exist within the broad scope of this population.

What this paper adds
• This paper provides a unique overview of where and by whom a wide range of research priorities for infants, children and young people with life-limiting conditions have been agreed.
• The research priorities identified are mapped in the context in which they were agreed, while common topics and themes are highlighted.

Implications for practice, theory or policy
• This paper presents an overview of consensus derived research priorities for infants, children and young people with life-limiting conditions, providing the opportunity for a coherent approach to improving the evidence base for this area of practice.
• This study highlights the need for broader consideration of stakeholder perspectives when undertaking research prioritisation exercises.
• However further identification of research priorities cannot be justified at this time unless ascertaining the perspectives of children and young people and their families.

BACKGROUND
The number of children with life-limiting or life-threatening conditions has been rising with latest figures estimating 49,000 children and young people with a life-limiting condition in the UK and approximately 21 million worldwide. These include conditions for which there is no reasonable hope of cure and from which children or young people will die, as well as conditions for which curative treatment may be feasible but can fail, such as cancer or heart failure. In children and young people, more than 300 diagnoses are life-limiting or life-threatening, including Duchene muscular dystrophy, severe cerebral palsy, neurodegenerative conditions, and severe congenital anomalies. Although many of the individual diagnoses are rare, as a group children and young people with a life-limiting condition are a larger patient population than many other long term conditions in children and young people, such as diabetes mellitus. Many of these children are living longer due to the use of medical technologies, e.g. ventilation and gastrostomy feeding, and more aggressive treatment of complications and they are often high users of healthcare services. The recent UK National Institute for Health and Care Excellence guidance on End-of-life care for children and young people, although focussed on end of life care, highlighted the lack of evidence base on which the care of these children and young people was based. The recent Lancet Commission on Palliative Care and Pain Relief states that globally, nearly 2.5 million children die in need of palliative care and pain relief, and over 90% of paediatric deaths associated with serious health-related suffering are avoidable. The American Academy of Pediatrics guidelines and recommendations on paediatric hospice and palliative care are based on available evidence and consensus expert opinion, while acknowledging the need for further clinical and health service research.
In terms of medical research involving children and young people with life-limiting conditions has in the past been seen as difficult, for example around issues of access, clinical considerations and obtaining informed consent. This has resulted in clinical decisions having to be based on converting the findings of research in adults to apply to children. For example, drugs are licenced for specific indications and patient groups based on the results of clinical trials, usually carried out in the adult population, age 18 to 65. It is common therefore in the absence of evidence for children to be prescribed medicines in ways that are not included in the license. Children and young people are different biochemically and physiologically from adults, so this is not a satisfactory solution. However, over the last decade or so initiatives such as the International Alliance for Better Medicines for Children in 2006 have been set up and it has become commonly agreed that children are not 'little adults'. Ethical considerations have been overcome, so children and young people are encouraged to be involved in investigations into the most effective diagnoses, treatments and delivery of care specifically for them.

Involving children, including those with life-limiting conditions, directly in research about their lives and the services they use is, however, more established. The evidence base generally for all aspects of care for children and young people and understanding of what is important to patients and their families has grown. But there are still some significant gaps in terms of evidence with respect to particular populations and/or the scope, or comprehensiveness of that evidence.

The identification of priority areas for research using consensus methods is recognised as a good way of ensuring that finite research resources are used to maximum effect. Prior knowledge and preparatory internet searches identified a significant number of published research prioritisation exercises relevant, in varying degrees, to neonates, infants, children and young people with life-limiting conditions and their parents and carers. We therefore planned to examine the focus, context and questions identified by existing prioritisation exercises in this area by undertaking a scoping review to:

- Systematically identify existing research prioritisation exercises relevant to infants, children, and young people with life-limiting conditions and their families and carers.
- Formulate the findings into a framework, mapping existing priorities.

**METHODS**

The exploratory nature of the review, the broad scope and anticipated volume of literature was suited to scoping review methods. In addition, the absence of any validated quality appraisal tools for consensus exercises precluded a systematic review. The scoping review was undertaken using systematic methods and is reported in line with the PRISMA statement. The review protocol was agreed and made publicly available on the Martin House Research Centre website prior to screening studies against inclusion criteria; scoping review protocols are not accepted for registration on PROSPERO. Amendments made to the protocol were highlighted and dated.

**Eligibility criteria**

Studies using any consensus consultation method were included. Studies had to include at least two rounds of consultation.
Excluded were research and development analyses and knowledge gap analyses as these are different concepts to agreeing research priorities.

There were no restrictions on the people or stakeholders undertaking the prioritisation exercise. Research prioritisation exercises aimed at identifying research for the benefit of: neonates (birth to <28 days), children and/or young people (28 days to age 25) with life-limiting, life-shortening and/or life-threatening conditions; their family, parents, carers; and/or the professional staff caring for them. We used the definitions for life-limiting, life-shortening and life-threatening conditions (hereafter encompassed in the term life-limiting) adopted by Together for Short Lives:

- **Life-limiting/life-shortening conditions** are those for which there is no reasonable hope of cure and from which children or young people will die. Some of these conditions cause progressive deterioration rendering the child increasingly dependent on parents and carers.
- **Life-threatening conditions** are those for which curative treatment may be feasible but can fail, such as cancer; are also included. Children in long-term remission or following successful curative treatment are not included in this review.

Exercises seeking to identify research priorities for mixed age groups (children and/or young people and adults) were included if details of the priorities were reported separately for children and young people. Likewise, exercises setting priorities for our target age groups but not exclusively those with life-limiting conditions, were included if priorities specifically related to life-limiting conditions were reported.

The outcomes of interest were the top ten priorities for future research, such as topic areas or specific research questions presented as the main result of the consultation. Secondary lists, for example priorities for a sub-group of participants, were excluded.

**Search sources and strategy**

To identify studies for inclusion we searched ASSIA, CINAHL, MEDLINE/MEDLINE In Process and Embase. The search strategy was developed by an experienced information specialist in collaboration with the rest of the review team. The strategy consisted of thesaurus and free text terms for "research prioritisation" combined (using AND) with terms for children and young people and their families. The search strategies are presented in Supplementary File 1.

The reference lists of included papers and relevant websites, such as the James Lind Alliance (JLA) Prioritisation Setting Partnership (PSP) website, were hand-searched for on-going and completed research prioritisation exercises.

There were no restrictions on the setting in which the exercise was undertaken or for the research priorities set. However, to aid generalisability to settings with similar healthcare provision, we restricted exercises to those in the English language and undertaken in Organisation for Economic Co-operation and Development countries. Healthcare, treatment and service delivery for children and young people with life-limiting conditions have changed considerably over the last few decades. In addition, the identification of priorities
for future research implies that subsequent research is likely to have been undertaken. For these reasons the searches were restricted to exercises published from 2000 to date.

**Study selection and data collection**

Study selection was performed independently by two researchers with discrepancies resolved through discussion or by recourse to a third researcher. Titles and abstracts were screened first, then at second screening full papers were assessed for inclusion.

A data extraction form was designed and piloted independently by two researchers. Once finalised, one researcher data extracted the included studies and a second researcher checked 30% of the records. Discrepancies were resolved by discussion or by recourse to a third researcher.

Items data extracted were: publication details; funding; aims and objectives; study country/ies; setting; health condition/s; age group; methods used to achieve consensus; degree of consensus; priorities identified; and planned use of priorities.

**Strategy for collating, summarising and reporting the data**

We charted the data, and collated, summarised and reported the results based on the enhancements of Levac et al. to the scoping review framework suggested by Arksey and O’Malley. This included the research team collectively developing the data extraction tables and discussing sequential iterations and piloting until an appropriate data set had been agreed. Simple narrative and descriptive statistics were used for reporting the included study characteristics and methods.

For the synthesis we used the WHO established classifications of functioning, disability and health as a starting point for developing a framework. Three researchers independently categorised the identified priorities into the framework. Many of the authors had grouped their questions or issues into topic areas which were also used to inform the process. Repeated discussions and iterations took place, including consideration of the factors that informed prioritisation decision-making.

**RESULTS**

The electronic searches, run in February 2017, identified a total of 7447 records which were loaded into EndNote v7 bibliographic software and de-duplicated leaving a total of 4970 records for screening. The study flow chart is presented in Figure 1.

Initial screening identified 189 records for full text assessment, from which 24 studies were found to meet the inclusion criteria. A table of characteristics of the included studies is provided in Table 1. A list of excluded studies and the full data extraction tables are available on request from the authors.

**Theoretical consensus models used**

The consensus method most frequently used in the included consultations was the Delphi Consensus method. The majority of consultations used questionnaires to obtain responses, mainly distributed electronically, but a few were distributed as hard copies.

**Settings**
Most of the consultations were undertaken in single countries. Three consultations were Europe wide, and a fourth attempted global reach. The majority of consultations were undertaken in, and for use by, centres providing specialist care such as paediatric palliative care (including hospices), neo-natal/paediatric intensive care or oncology units.

**Participants in the consultations**

Details of participants’ professions/roles were generally not clearly reported but the majority were health professionals, including academic health professionals. Other participants included academics, social workers, volunteers, carers, charity staff, teachers, policy makers, and priests.

Only eight studies reported including children and young people and/or parents and family members (Table 1). None of the included studies only consulted with these groups. There were some reports on the difficulties of recruiting children, young people and parents, but the majority of papers did not mention any attempts to include family members or explain why they had not.

The number of individuals taking part in a consultation exercise overall, and within individual rounds, varied considerably and not all study reports included details of numbers at every stage.

**Focus for research priorities**

The priority setting exercises either focussed on conditions, such as cancer or cystic fibrosis, an overarching ‘any life-limiting condition’, or a specified care need, such as requiring paediatric palliative/end of life care, or other therapies.

The majority of papers reported age descriptively, for example “children and young adults” or “paediatric”, rather than specifying an age range.

**Planned use of priorities**

The majority of research prioritisation exercises were undertaken with the intension of the results being used by the wider health professional and research communities. Others were undertaken to provide priorities for the benefit of funding and commissioning agencies; raising the profile of paediatric palliative care; ensuring the perspective of the family was considered in neonatal research. Eight studies were undertaken specifically to inform the research agenda of those undertaking the exercise for their particular setting.

Seventeen studies referred to searching for existing prioritisation exercises in the background to their papers, either not finding any relevant to their area of interest or justifying a repeat for differences in service delivery between countries. Fourteen studies discussed the priorities identified in the context of existing research; two of these reported modifying the priorities in light of finding relevant research.

**Research priorities**

From the 24 studies meeting the inclusion criteria, we identified 279 research questions or priority areas for research. The majority of studies presented up to 10 top priorities, 11 listed between 12 and 33 and one listed 82 items. The range was 4 to 82, the average 17
and the median 11. Eight of the studies reported more than one final list of priorities: these were differentiated by topics, categories, or by groups of participants in six cases, and in two cases, as well as ranking questions, the authors ranked research domains.

The priorities were variously called research priorities, items, themes, questions, statements, or issues, and were framed in different ways. We use the collective term ‘research priorities’ from here on. This ranged from detailed specific questions including population, intervention, comparator and outcome (PICO) to less explicit, more exploratory questions such as what is best practice for a specific stage of care. Some studies simply listed, or included, one or two word topic areas. This ranged from detailed specific questions including population, intervention, comparator and outcome (PICO) to less explicit, more exploratory questions such as what is best practice for a specific stage of care. Some studies simply listed, or included, one or two word topic areas.

Synthesis of research priorities

As the majority of studies sought to identify 10 or less research priorities, and given our objective to identify issues/topic areas considered most pressing or causing greatest concern, we limited inclusion in our synthesis of the data to a maximum of 10 of the overall top priorities reported.

This yielded a total of 279 research priorities from the included studies. Some studies presented lists of priorities under more than one category, and some had less than 10 priorities (hence the odd number). The complete list of synthesised research priorities identified by the included studies is very large and therefore presented by topic, in Supplementary File 2.

Some studies concerned specific types of service or setting (palliative, hospice and end-of life care; tertiary referral centres; oncology; and PICU and NICU). In terms of the scope of the studies with respect to diagnostic categories, they were grouped as either life limiting conditions or cancer.

An iterative analysis of the research priorities generated 16 topic areas across which were subsumed 55 sub-topics and 12 sub-sub-topics. These are presented in Table 2 and a high level summary of the key issues provided here.

- Epidemiology / population (including access to services): the need to and feasibility of collecting data; barriers and facilitators to accessing services; the needs of CYP and their parents; and strategies for health promotion.
- Measurement and assessment: understanding what quality of life means; assessment of symptoms; levels of sedation in babies; assessment of antibiotic levels; psychosocial issues for children with cancer.
- Service delivery and models of care: best models for organisation and delivery of care; differences in quality and cost; how to maintain good governance; use of tele-care; barriers and facilitators to shared care; inter- and multi-agency collaboration; effective transfer between services; palliative care outcomes of importance to CYP and families; effective ways to measure, prevent, record aspects of care.
- Health interventions: pharmacological and/or invasive: avoiding necrotising enterocolitis; preparing children for cancer treatment; best practice in ventilation; meeting nutritional needs; long term effects of cancer treatments.
Symptom management and control: effective management of symptoms e.g. pain control, in different settings and different stages; non-pharmacological interventions; drug trials.

Other interventions: physical health and functioning: optimum timing and delivery of interventions; long-term safety and effectiveness of techniques to manage spasticity; achieving, or improving, continence; effective communication with CYP with neurodisability; strategies to manage sleep disturbance in CYP with neurodisability; facilitating engagement in physical activity.

Intervention adherence: barriers and facilitators.

Emotional and psychological issues: factors to protect or risk factors when adjusting to living with a LLC; coping mechanisms; anxiety, breathlessness and sleeplessness as clusters; emotional and psychological challenges and experience for CYP; needs of parents, siblings and wider family; support systems and practices; promotion of well-being.

Participation and inclusion: promoting positive attitudes; supporting independence in terms of participation and mobility.

Communication and decision-making: recording care; communications within multi-disciplinary teams; communications between staff and CYP and their families; shared decision making.

Other family needs and support: impact on the family; support for family; support for CYP to self-manage.

Practices related to palliative and end-of-life care: understanding and supporting parent expectations; understanding what dying well means.

Bereavement: CYP experience of and coping with bereavement; care and support needs of families.

Ethics: parental role in ethical decision making; ethical dilemmas for staff.

Workforce: recruitment, retention and support for staff; training to equip staff to deliver high quality evidence based care; strategies to help staff communicate with CYP and their families.

Funding: who and how research questions should be set; where funding should be targeted.

DISCUSSION

Main findings/results of the study

In this scoping review we aimed to map research priorities identified from existing research prioritisation exercises relevant to infants, children, and young people with life-limiting conditions. Extensive searches of bibliographic databases and hand searching identified 24 research prioritisation exercises that met the inclusion criteria. From these 279 research questions or priority areas for health research were identified. The content of these research priorities was subject to a thematic analysis which generated 16 topic areas with numerous sub-topics across these areas. This demonstrates the wide range of research priorities identified by existing studies, with major themes including medical treatments, condition and symptom management; non-medical aspects of care and support; and topics of shared interest and expertise. No single research prioritisation exercise captured the total
range. This demonstrates the value of our prudent use of resources having taken the
approach of a scoping review.

Given the number and diversity of life limiting conditions and the range of services involved
in their care, the large number of published exercises and identified priorities was not
unexpected. Whilst these studies generated a large volume of research priorities, it was
possible to organise these into a relatively small number of overarching and meaningful
topic areas.

The research prioritisation exercises included in this review were wide-ranging in terms of
purpose, inclusivity of stakeholders and experts, topic areas and scope. However, a key
limitation of many studies was the lack of involvement of children and young people and
parents in the research prioritisation exercise or, where sought, only minimal involvement
was secured. It was not possible to map the priorities by condition or by age group as we thought might be possible based on the systematic approach to
undertaking research prioritisation taken by the JLA PSP. The rigorous methods used by
the JLA set a standard to aspire to, but which are challenging even for the JLA to meet. For
example we included the JLA PSP on neurodisability, where the authors report the problems
of including children and young people in the consensus process. An issue echoed in other
studies where attempts were made. While including children and young people, parents and
other family members is always likely to be challenging given the nature of the conditions
concerned, it is still disappointing that so few studies reported even considering
perspectives other than those of the health professionals. Even within the involvement of
health professionals, this almost exclusively included doctors and nurses, with very little
involvement of other relevant professions such as physiotherapists, speech and language
therapists, occupational therapists, clinical psychology, and dieticians. This is concerning
given the evidence from various fields that what is important to patients can be different to
what is important to clinicians. Also, other professional groups outside of the healthcare
sectors such as social workers, priests/chaplains, and teachers may bring new insights and
perspectives. A useful and efficient way forward to address this inequity and imbalance in
the stakeholder groups involved in identifying research priorities to date would be to use
the findings from this review as a basis for consultation with families and particular
professional groups. We recently carried out such an exercise to inform and guide the work
of our newly established research centre.

The number of exercises identified may illustrate a wide spread awareness of the lack of evidence in this area, and the interest in providing the evidence in a way that maximises
value. We limited the search period from post-2000 to February 2017. It may be that some progress has been made in producing research evidence. We are aware that the JLA PSP on
neurodisability has informed the commissioning of research by the UK’s National Institute
for Health Research. However, within the studies included in this review, only a few
reported that research priorities were limited to those where there was a known degree of
uncertainty. It was beyond the remit of this review to evaluate the current evidence
available against each of the topic and sub-topics identified by our synthesis.

Undertaking thorough literature searches to justify carrying out any study is good research
practice. So, while this review is a valuable resource, it does not negate the need for
evidence reviews to inform future research and decisions regarding the commissioning of research.

The drivers behind the production of the prioritisation exercises varied. For many the reason for doing the exercise was specific to a profession or service at a single site. This may have been in the belief that priorities would vary between sites and professions. However, again the overlap of priorities in this scoping review demonstrates the commonality of issues that need to be addressed. When more research has been carried out and it becomes appropriate to take a fresh look at priorities, we believe a more comprehensive approach would be justified.

**Strengths and weaknesses/limitations of the study**

A key strength of our review is in the systematic methods used to ensure only those consultations that met our pre-specified criteria were included. Our focus on research priorities for children and young people meant we had to exclude studies that included all ages but did not present the results in a way in which we could identify those relevant to children and young people. This meant included studies where the top ten priorities were clearly not relevant to children and young people but also some where they may have been relevant but not necessarily arrived at with this age group specifically in mind. For example, we had to exclude a well conducted JLA PSP which set priorities for clinical research in primary brain and spinal cord tumours related to any age and which included paediatric representatives on the panel.We also excluded studies that focussed on the Emergency Department setting as generally the population and conditions presenting are very mixed and priorities therefore not focussed on children and young people with life limiting conditions. This demonstrates the rigor of our selection process but also underlines further the extensive number of priority setting exercises closely related to the 24 studies that met our inclusion criteria.

As this is a scoping review we have not included an appraisal of the quality of the included studies. Given the potential for consensus priorities to influence the future direction of research bids and funding, consideration should be given to the way in which the consensus has been carried out. There is extensive literature on consensus methods and now on the reporting of Delphi studies in palliative care. In synthesising the included priorities, we used the existing recognised terminology related to children and young people with life-limiting conditions as a starting point for the framework. Although three researchers independently allocated priorities to topics in an iterative analytical process, we have to acknowledge a level of subjectivity in the synthesis. At all times we were conscious of the need to acknowledge the drivers behind individual studies and the factors that informed the decision-making processes. For example, it was important to know who set the questions, in what context and with what degree of consensus.

We placed no restrictions on the setting in which priorities were set. However, the epidemiology of life-limiting conditions in children, their management, the availability of services and treatments and outcomes are very different in the high resource and low/middle resource countries therefore the research priorities here will be very different. So, to aid generalisability to settings with similar healthcare provision, we restricted
inclusion to those in the English language and undertaken in Organisation for Economic Co-
operation and Development countries.

What this study adds
We believe this scoping review presents a unique overview of research priorities, arrived at
through consensus processes, with respect to infants, children and young people with life-
limiting, life-threatening and life-shortening conditions. By detailing the individual
consultations and presenting our synthesis of the priorities identified, we provide evidence
of the range and focus of areas for improving the evidence base for care of this population
and their families. As such it provides a useful resource for researchers, professionals,
funders and commissioners of research and other stakeholders involved in supporting
evidence informed practice. The findings provide an opportunity at this time to maximise
the use of limited research resources by focusing on filling priority evidence gaps.

CONCLUSION
By taking systematic scoping approach to identification of existing research prioritisation
exercises and providing transparency in our methods we believe we present a reliable
overview of the priorities already set in this area. Anyone wishing to develop a programme
of research or planning a specific project could use this review as a starting point, and
justification, for choosing topics or questions. Users will of course need first to check if
research has been done or is in progress, on priorities before proceeding.

There are a significant number of research prioritisation exercises related to children and
young people with life-limiting conditions. By mapping the priorities while being sensitive to
the context in which they were set, we have achieved our aim to provide an overview of
existing research priorities.

DECLARATIONS
Authorship. AB conceived the approach, all authors contributed to the design of the review
and approved the protocol. KW developed the search strategy and ran the searches. AB and
JM undertook the screening and data extraction with BB and LF providing arbitration; AB,
JM, BB and LF synthesised the priorities identified. AB drafted the paper and JM, KW, BB and
LF provided critical comments. The final version was approved by all authors.

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Declaration of conflicts of interest. The Authors declare that there is no conflict of interest.

Research ethics and patient consent. Research ethics and consent were not required for
this review of published studies.

Data management and sharing. The data extraction tables are available from the authors
on request.

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5. Feudtner C, DiGiuseppe DL and Neff JM. Hospital care for children and young adults in the last year of life: a population-based study. *BMC Medicine* 2003; 1: 3-3. DOI:


11. Bourns IM. Unlicensed medicines in the UK – legal frameworks, risks, and their management. @Medicine Access@Point of Care 2017; 1: 0. 10.5301/maapoc.0000006. DOI: 10.5301/maapoc.0000006.


