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**A small step in the right direction for reducing
postoperative pulmonary complications**

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6 A small step in the right direction for reducing postoperative pulmonary complications
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6 Only 12 years after the first successful public demonstration of general anaesthesia (GA) in
7 1846, one of the first adverse effects to be described was reported by John Snow: '*If the*
8 *inhalation is continued the breathing is rendered difficult, feeble, or irregular, and is*
9 *sometimes performed only by the diaphragm, whilst the intercostal muscles are paralysed. If*
10 *the dose of chloroform is gradually increased after these effects are produced, the breathing*
11 *entirely ceases*'.¹ Ever since ~~then~~ the fact that GA-anaesthesia can cause postoperative
12 chest complications has been widely accepted by anaesthetists, so it is surprising that only
13 relatively recently have serious efforts been made to understand the aetiology of this
14 common problem, with a view to being able to ~~their prediction~~ and prevention ~~chest~~
15 ~~complications~~. The term post-operative pulmonary complication (PPC) has become widely
16 adopted and describes a single outcome measure representing a disparate collection of
17 complications, linked only by a common organ of origin. PPCs are more- common than
18 cardiovascular- complications, have a significant impact on outcome measures such as
19 length of hospital stay, increase healthcare costs, and when severe are associated with
20 increased mortality.
21
22 Current knowledge of PPCs suggests they are multifactorial in origin.^{2,3} The well known
23 physiological changes associated with GA-anaesthesia such as reduced functional residual
24 capacity, disturbed ventilation-perfusion relationships and development of atelectasis ~~may~~
25 ~~can~~ continue into the post-operative period when surgical trauma, pain and the stress
26 response to surgery exacerbate these physiological abnormalities leading to pathological
27 conditions. Artificial ventilation during GA-anaesthesia remains ~~an~~ significantly
28 unphysiological process, applying pressures to delicate lung tissue that are several times
29 higher than physiological breathing, potentially causing physical lung damage or
30 inflammatory changes. There are over 50 factors that have been shown to be predictors for
31 developing a PPC,³ which can be categorised ~~into-as~~ those relating to the patient, the
32 procedure, or laboratory investigations. Approximately half of these are potentially
33 modifiable.³
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6 Many strategies to prevent PPCs are described, but good quality evidence of outcome
7 benefits for most of them remains elusive. Take preoperative smoking cessation as an
8 example of a non-invasive intervention that really should work. Studies in the 1980s of
9 patients having cardiac surgery paradoxically suggested that stopping smoking for less than
10 8 weeks pre-operatively was associated with a ~~worse-greater~~ respiratory complication risk
11 than keeping smoking until the day before surgery.⁴ Subsequent work has convincingly
12 challenged this finding such that current evidence suggests that stopping smoking pre-
13 operatively does indeed reduce ~~the~~ PPC risk, and that the longer the period of effective
14 smoking cessation the better the outcome.⁵ However there remains varied opinion on how
15 many weeks are required for this benefit to become clinically significant, or whether the
16 number of pack-years smoked before quitting affects the benefit. In the most recent and
17 widely used score for predicting PPC risk smoking did not prove to be an independent
18 predictor, while recent upper respiratory tract infection or a preoperative oxygen saturation
19 <96% on air did.⁶ This suggests that in the absence of these two clinical findings a smoker's
20 risk is the same as for a non-smoker.

21
22 A similar uncertainty exists around artificial ventilation during anaesthesia. Use of a
23 'protective ventilation' strategy of low tidal volume, moderate positive end-expiratory
24 pressure (PEEP), pressure limited ventilation and recruitment manoeuvres, as used for
25 many years in lung-injured patients in intensive care, is now generally agreed to reduce
26 PPCs.⁷ But there is less agreement on the ideal settings for these ventilation components, or
27 how they should be modified in specific patient groups such as those with pre-existing lung
28 disease, morbid obesity or during one-lung ventilation (~~OLV~~). For example pressure-
29 controlled ventilation as part of protective ventilation has shown some benefits in some of
30 these challenging patient groups such as one-lung ventilation OLV⁸ and obesity.⁹
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32 Conversely, a recent observational study involving more than 100 000 patients having varied
33 surgical procedures found a significant benefit for volume-controlled ventilation in reducing
34 PPC occurrence.¹⁰ Despite these uncertainties on the finer details of how best to prevent
35 PPCs there is general agreement that some form of intraoperative protective ventilation is

beneficial,^{11,12} but surveys of actual practice show adoption of this approach to be slow.^{13,14}

There is also controversy on the optimal inspired oxygen concentration to use in the perioperative period, with most anaesthetists advocating use of the lowest inspired concentration to avoid atelectasis and higher mortality,¹⁵ in contrast to a World Health Organisation (WHO) recommendation of 80% inspired oxygen to prevent surgical site infection.¹⁶ a recommendation that has been questioned.¹⁷

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A clearer understanding of potential ways of preventing PPCs is therefore required, but two problems are impeding progress in this area. First is variation in the components of ventilation 'bundles' used by studies in this area. A clear example is the lack of agreement on what constitutes an effective lung recruitment manoeuvre, with three quite different techniques used in different studies.^{18,7} Second, and in our opinion more fundamentally, is the lack of agreement on what constitutes a PPC. Considering the large number of individual patient events that may be counted as a PPC, use of composite measures is unavoidable to obtain large enough event rates to allow interventional studies. But the lack of agreement on which composite to use is currently a problem, both in terms of facilitating research and on the implications of those studies for patients. For example, consider two studies looking at PPC incidences with and without muscle relaxant usage. In the first a PPC was recorded if oxygen saturation ~~to~~ fell to <90% for ~~one-1 minute~~ in the 20 ~~minutes~~ after extubation,^{18,19} while in the second a diagnosis of one or more of respiratory failure, pulmonary oedema, tracheal reintubation, or pneumonia was required to be counted as having a PPC.^{19,20} The implications for patients from these two studies are quite different.

An international initiative in peri-operative medicine is beginning to address this second problem. In 2015 a joint taskforce of the European Society of Anaesthesiology and European Society of Intensive Care Medicine published definitions for 22 individual adverse respiratory events and some composite measures for PPCs.^{20,21} ~~On page @@@ of In~~ this month's *British Journal of Anaesthesia* ~~et al and colleagues,~~²² on behalf of the Standardized Endpoints for Perioperative Medicine (StEP) collaboration, have continued this work and reached a consensus for a definition of PPCs in an attempt to further unify research in this

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6 area.²⁴ Agreed definitions were reached by a systematic literature review followed by the
7 now widely-used Delphi process including three rounds with 75 expert participants in round
8 two. The group accepted existing definitions for pneumonia and acute respiratory distress
9 syndrome, the latter being combined with an adapted definition of re-institution of
10 mechanical or non-invasive ventilation to define postoperative respiratory failure. Perhaps
11 indicative of the challenges of existing definitions, the taskforce was unable to agree on a
12 ~~acceptable~~ previous definition of PPCs and instead proposed a new definition despite the
13 abundance of published studies that have examined post-operative pulmonary
14 complications.

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16 The paper is to be applauded for attempting to end the plethora of definitions used for a
17 PPC, and hopefully this will be accepted by the PPC research community for future studies.
18 For the first time, the PPC definition described introduces the concept of severity (classified
19 as mild, moderate or severe) rather than the all-or-none mostly used previously. This will
20 hopefully avoid future examples like that above where patients with a brief desaturation in
21 PACU-recovery or a prolonged stay in intensive care are both classed as simply having a
22 PPC for research purposes. Unfortunately, in order to achieve a clearly defined set of criteria
23 for a PPC, Abbot ~~et al. and colleagues~~ excluded many complications affecting the respiratory
24 system ~~which that~~ are seen on a daily basis in most hospitals. These include pulmonary
25 thromboembolism, bronchospasm, cardiogenic pulmonary oedema, pleural effusion and
26 pneumothorax, many of which were included in various previous definitions. The justification
27 for these exclusions was to avoid including complications that can only be identified by
28 specific screening of all study participants, such as pulmonary thromboembolism, and a
29 desire to keep the results applicable to perioperative care of a mixed surgical population
30 rather than focussing on specific technical complications of surgery. This is fine, but having
31 excluded so many common complications previously classed as PPCs, the incidence of
32 PPCs using these new criteria is likely to be much lower than previously cited. Furthermore,
33 can they really be called PPCs if all these critical complications involving the respiratory
34 system are excluded?

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6 This work also provides a welcome example of perioperative medicine on an international
7 scale: the group involved in agreeing the new definitions included an impressive
8 collaboration of experts from ten countries and three continents. This not only gives the work
9 international credibility but also makes wider acceptance of the definitions more likely, so
10 hopefully improving the performance and interpretation of future PPC research. As with all
11 new definitions, the validity, reliability and responsiveness of the proposed standardised
12 endpoints for pulmonary complications will have to be tested in the real world. The
13 systematic and transparent consensus process will lend it strength and credibility to the
14 clinical community.
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22 This example of international collaboration in peri-operative medicine is an overdue attempt
23 to tackle a common and hazardous complication of general anaesthesia. Since 2015, the
24 StEP Collaboration has been working towards standardised endpoints for perioperative
25 patient outcomes.²² An agreed definition has the potential to advance research in PPCs, but
26 there is still a long way to go before we fully understand PPCs and can finally start to change
27 practice to prevent them.
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34 **Authors' contributions**

35 AL and JY worked together to draft, produce and approve the manuscript.
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38 **Declarations of interest**

39 None declared
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