## REVIEW

# Update of complications and functional outcome of the ileo-pouch anal anastomosis: overview of evidence and meta-analysis of 96 observational studies

Sharonne de Zeeuw • Usama Ahmed Ali • Rogier A. R. T. Donders • Willem E. Hueting • Frederik Keus • Cees J. H. M. van Laarhoven

Accepted: 22 December 2011 / Published online: 10 January 2012 © The Author(s) 2012. This article is published with open access at Springerlink.com

## Abstract

*Objective* The objective of this study is to provide a comprehensive update of the outcome of the ileo-pouch anal anastomosis (IPAA).

*Data sources* An extensive search in PubMed, EMBASE, and The Cochrane Library was conducted.

*Study selection and data extraction* All studies published after 2000 reporting on complications or functional outcome after a primary open IPAA procedure for UC or FAP were selected. Study characteristics, functional outcome, and complications were extracted.

*Data synthesis* A review with similar methodology conducted 10 years earlier was used to evaluate developments in outcome over time. Pooled estimates were compared using a random-effects logistic meta-analyzing technique. Analyses focusing on the effect of time of study conductance, centralization, and variation in surgical techniques were performed. *Results* Fifty-three studies including 14,966 patients were included. Pooled rates of pouch failure and pelvic sepsis were 4.3% (95% CI, 3.5–6.3) and 7.5% (95% CI 6.1–9.1),

S. de Zeeuw (⊠) · U. A. Ali · F. Keus · C. J. H. M. van Laarhoven Department of Surgery, (Division of Abdominal Surgery), Radboud University Nijmegen Medical Centre, PO Box 9101, 6500 HB Nijmegen, The Netherlands e-mail: s.dezeeuw@chir.umcn.nl

R. A. R. T. Donders Department of Epidemiology, Biostatistics and HTA, Radboud University Nijmegen Medical Centre, Nijmegen, The Netherlands

W. E. Hueting Department of Surgery, Diakonessenhuis, Leiden, The Netherlands respectively. Compared to studies published before 2000, a reduction of 2.5% was observed in the pouch failure rate (p=0.0038). Analysis on the effect of the time of study conductance confirmed a decline in pouch failure. Functional outcome remained stable over time, with a 24-h defecation frequency of 5.9 (95% CI, 5.0–6.9). Technical surgery aspects did not have an important effect on outcome.

*Conclusion* This review provides up to date outcome estimates of the IPAA procedure that can be useful as reference values for practice and research. It is also shows a reduction in pouch failure over time.

Keywords Functional outcome  $\cdot$  Ileo-pouch anal anastomosis  $\cdot$  Meta-analysis

### Introduction

Restorative proctocolectomy by means of an ileo-pouch anal anastomosis (IPAA) is the procedure of choice for the surgical treatment of ulcerative colitis (UC) and familial adenomatous polyposis (FAP). The IPAA procedure generally results in acceptable long-term functional outcomes and improvement of quality of life. It is, however, associated with substantial morbidity with complications up to 50% of patients [1]. Since its introduction in 1978, the IPAA procedure has continuously been subjected to attempts of improvements in technique [55]. Additionally, much has been achieved in improving the perioperative care. Centralization of complex surgery has been a recent development in many countries. All these developments may have resulted in improved outcomes. A large number of observational studies have reported outcomes of the IPAA procedure, most commonly from a selected group of patients in one hospital. This review aims to provide an overview of the available evidence and to evaluate the effect of the continuous developments on outcomes. Previously, a systematic review summarizing complications and functional results after an IPAA procedure in studies published until 2000 was conducted. This current systematic review provides an update regarding the outcomes of the IPAA procedure and uses the combined data set of both reviews to analyze changes in the outcomes of the IPAA procedure.

## Methods

This systematic review aims to provide an overview regarding the outcomes of the open IPAA procedure. Additionally, it evaluates the effect of development of practice on the most important outcomes over time.

## Search strategy

A systematic literature search with predefined search terms was carried out in four electronic databases: Medline (1-1-2000 to 1-1-2010), EMBASE (1-1-2000 to 1-1-2010), and the Cochrane Library (Issue 1, 2010) (Fig. 1). Two authors (SZ and UAA) independently performed the selection of the publications according to inclusion criteria. Disagreements were discussed with a third reviewer (FK). Additional relevant studies were looked for by cross-reference checking of all included studies. Language restrictions were not applied.

## Selection of studies

## Inclusion criteria

Title and abstract of all identified publications (prospective and retrospective and observational and comparison studies) were screened according to the following inclusion criteria: The study population consists of adult patients with established UC or FAP undergoing a primary IPAA procedure; the intervention is a clearly documented open IPAA procedure in an elective setting with or without a diverting ileostomy and irrespective of the number of stages of the operation; the study reports at least one of the primary outcomes reported below; and the study includes a consecutive series with a minimal sample size of 50 patients.

## Exclusion criteria

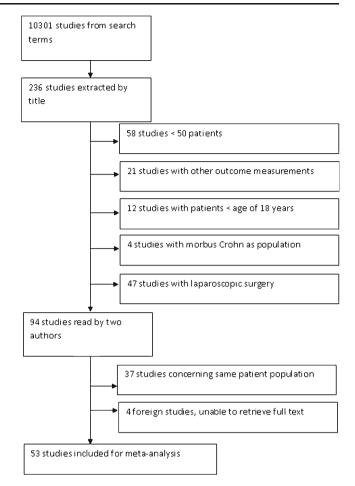


Fig. 1 Flowchart of study selection resulting in the 53 included studies

(>65 years). Studies were excluded as well when the IPAA was performed for other indications than UC or FAP (e.g., Morbus Crohn) or when a secondary IPAA was performed (e.g., after ileo-rectal anastomosis). Studies were also excluded when there was selective outcome reporting, i.e., studies focusing on one parameter (e.g., pouchitits) without reporting of any additional outcomes.

Outcomes of interest and definitions

Primary outcomes were pouch failure, pelvic sepsis, and severe day incontinence. Secondary outcomes included other complications (stricture, fistula, small bowel obstruction, pouchitis, and sexual dysfunction) and parameters of functional outcome (defecation frequency and incontinence). Table 1 shows the definitions of all outcomes.

## Data extraction

After assessment for eligibility, two authors (SZ and UAA) independently extracted the following data if available: numbers of patients, patient characteristics, dates of start and end of the inclusion period, duration of follow-up,

#### Table 1 Definition of outcomes

Pouch failure	Pouch excision or a nonfunctioning pouch at 12 months after IPAA
Pelvic sepsis	Pelvic abscess, anastomotic leakage or dehiscence, pelvic/perineal wound infection
Fistula	Any pouch-related fistula
Stricture	Anastomotic fibrosis necessitating dilatation
Mild fecal incontinence	Soiling, spotting in underwear
Severe fecal incontinence	Regularly severe leakage or fecal loss passive fecal incontinence

variations in surgical technique, numbers and type of complications, and parameters of pouch functional outcome. Authors were contacted when data were missing. One author provided updated unpublished information [66]. Double publications describing identical populations were considered as one study.

## Data of studies published until 2000

Previously, a systematic review was performed summarizing complications and functional results after an IPAA procedure in studies published until 2000 [39]. For time-frame analyses regarding the development of clinical practice and its effect on outcomes, data from both reviews were combined. Both reviews were conducted with similar methodology, and the full data set from the previous review was available.

## Statistical analysis

Relevant study characteristics and outcomes were extracted and presented for each study individually. Outcomes were subsequently pooled, and cumulative probabilities were calculated with 95% confidence intervals (95% CI). Data were pooled using a "random-effects model, based on the restricted maximum likelihood estimator in order to incorporate the heterogeneity between studies. Pooled results of this review (all studies published since 2000) were compared with pooled results of the previously published review (all studies published before 2000) by analyzing the complete set in one random-effects model and incorporating a dummy variable that coded for period. The Knapp and Hartung adjustment was used to obtain estimates and confidence intervals [33]. Additionally, analysis focusing on the effect of time of study conductance, centralization, and variation in techniques were performed using linear regression and one-way ANOVA as appropriate. Data management and statistical analyses were conducted using SPSS (version 15) and R Statistical Software (R Development Core Team, version 2.11.1) with the metafor package [52, 70].

#### Results

#### Selection process

The search resulted in a total of 10,301 hits. Initial selections based on titles identified 236 potentially relevant articles. Further selections based on abstracts excluded 142 studies. The full text of the remaining 94 studies was evaluated. Finally, a total of 53 studies with 14,966 patients were included (Fig. 1).

#### Description of identified studies

The most important characteristics of the included studies are presented in Table 2. Most studies were retrospective cohort studies (70%). The median sample size was 127 (range, 50–2,490) over a median inclusion periods of 12 years (range, 4–30). A diverting ileostomy was used in 79% (range, 5–100%) of all the IPAA procedures. A hand-sewn anastomosis was used in 40% of the procedures (range, 0–100%). Median duration of follow-up was 75 months (range, 6–180). Thirty-one studies mentioned postoperative mortality with a median of 0% (range, 0–2.9%).

#### Outcomes

#### **Complications**

Data on complications were available from all 53 studies. We found a pooled incidence of pouch failure of 4.3% (95% CI, 3.5–6.3; Fig. 2). A sensitivity analysis including only studies with minimal follow-up of 5 years showed a pouch failure rate of 4.7%. The pooled incidence of pelvic sepsis was 7.5% (95% CI, 6.1–9.1; Fig. 3). Pooled incidence rates for other important complications are presented in Table 3.

### Functional results

Data on functional results after IPAA were available from 26 studies including 5,321 patients. The pooled incidence of mild and severe fecal incontinence during the day was 14.3% (%CI, 7.3–25.9) and 6.1% (2.9–12.3), respectively (Table 3). Mean defecation frequency was 5.9 (4.9–6.7) per 24 h with a mean nighttime frequency of 1.5 (1.0–2.1).

Comparison with studies published before 2000

The previous review of studies published before 2000 identified 43 studies with a total of 9,317 patients [33]. Median duration of follow-up was 36.7 months (range, 12–99). Detailed discussion of the characteristics of these studies is published elsewhere [33].

Table 2 Characteristics of studies and selected outcomes following IPAA in the 53 included studies published since 2000

Author	Year of publication	Inclusion period	Number of patients	FU (months)	Age	Gender (% female)	Type of disease	Handsewn anastomosis (%)	With diverting ileostoma (%)
Fonkalsrud et al. [20]	2000	1993–1997	77		34.5	50	UC	57.1	100
Karlbom et al. [36]	2000	1983–1996	168	29	32	39.3	UC	58.9	70.8
Seidel et al. [60]	2000	1985–1996	55		31.2	45.5	Both		
Simchuk [61]	2000	1987–1996	114	38		43.9	Both		100
Gullberg and Liljeqvist [25]	2001	1990–1995	86	72	34.1	45.3	Both	2.3	10.5
Barton et al. [5]	2001	1983-2000	110	30	36.8		Both	100	100
Heuschen et al. [29]	2001	1991-2000	171	30	36.7	45.6	UC	100	66.7
Regimbeau et al. [53]	2001	1984–1998	172	60	36	47.1	Both	100	100
Blumberg et al. [12]	2001	1982-1995	145		34	49	Both		15.2
Dayton et al. [16]	2002	1982-2001	644	79	37.2	39.4	UC		
Rossi et al. [56]	2002	1989–2000	75		40	33.3	Both	54.7	100
Bullard et al. [13]	2002	1980–1992	154	151	34	44.2	Both	64	
Gignoux et al. [22]	2002	1993–1998	84	22	33	42.9	Both	100	
Mukhtar et al. [47]	2002	1990–1997	50		35	44	Both	2	98
Rudolph et al. [57]	2002	1991–1999	71	47	38	56	UC		84.5
Carmon et al. [14]	2003	1990–2001	77	51	38	55.8	UC		72.7
Teixeira et al. [63]	2003	1986–2000	80	108		50	UC	11.3	77.5
Michelassi et al. [46]	2003	1987-2002	391	37	33.7	44.2	Both	29.9	65
Holubar and Hyman [32]	2003		51	85	42.1	51	UC		70.6
Fowler et al. [21]	2003	1984-2001	106		40	50	Both	39.4	41.3
Parc et al. [49]	2004	1983-1990	110	175	26.5	43.6	FAP	100	
Lin et al. [43]	2004	1985-2002	61	60	37	54	Both	67.2	
Hueting et al. [34]	2004	1989–2000	111	42	35.4	68.5	UC		
Gosselink et al. [23]	2004	1989–2001	127	56	36	42.5	UC		21.6
Krausz et al. [40]	2005	1984–2004	174	64	34.1	47.7	Both	54	87.4
Hallberg et al. [26]	2005	1990–1997	100	53	32	39	UC		17
Swenson et al. [62]	2005	1995-2003	54	22	35	35.2			57.4
Chapman et al. [15]	2005	1981-2000	2002	121	33.5	46.6	Both		5
Kok Sun Ho et al. [30]	2005	1989–2001	330	32		59.1	UC		
Ikeuchi et al. [35]	2005	1999–2003	242		32.1	50.8	UC	100	38
Arai et al. [3]	2005	1993-2003	296	53	33.8	43.2	UC	96.3	55.4
Araki et a l. [4]	2006	1998-2003	220		34.5	46.8	Both		47.3
Rickard et al. [54]	2006	1982–2002			33	45.2	Both	32.4	88
Bednarz et al. [6]	2007	1994–2005	108	60		50	Both	18.5	100
Berndtsson et al. [8]	2007	1982–1995	370	180	34	45.7	UC		
Gualdrini et al. [24]	2007	1987–2005	391	45.6		51.7	Both		
Kiran et al. [38]	2008	1907 2000	2016	180	37.7	43.9	Both	15.4	85.9
Tulchinsky et al. [66]	2008	1986–2005	211	124.6	35.4	58.3	UC	16.5	79
Walker and Bulow [67]	2008	1987–2004	178	84	31	44.9	Both	6.7	100
Ferrante et al. [18]	2008	1990–2004	173	78	39	38.7	UC	2.3	85
Leal et al. [41]	2008	1984-2006	69	82	29.6	59.4	FAP	100	100
Zarate et al. [71]	2008	1984-2006	107		34.4	57.4 57	Both	12.1	100
Pinto et al. [50]	2008	2001–2008	66	21.1	52.8	66.7	Both	18.2	100
Leowardi et al. [42]	2009	1988–1996	197	147.6	33	45.2	UC	100	100
Norwood et al. [48]	2009	1988 1990	107	77	38	45.8	Both	100	99
Loftus et al. [44]	2009	2001–2005	237	5.9	38.9	45.8	UC		<i>,,</i>
Tekkis et al. [64]	2009	1976–2006	2491	5.9 74.4	35.9	47.9	Both	33.3	67.3
									01.5
Scarpa et al. [58]	2009	1984–2004	58	136	38	34.5	UC	0	

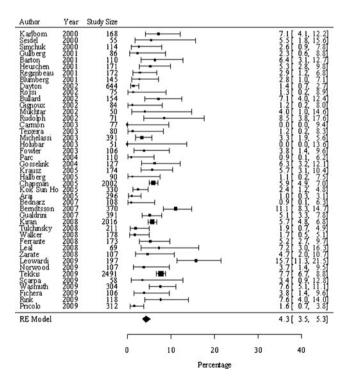
Table 2 (continued)

Author	Year of publication	Inclusion period	Number of patients	FU (months)	Age	Gender (% female)	Type of disease	Handsewn anastomosis (%)	With diverting ileostoma (%)
Wasmuth et al. [68]	2009	1984–2006	304	120	34.4	38.8	Both	40.1	84
Andersson et al. [2]	2009	1984-2003	110	144	35	45.5	Both		47.3
Fichera et al. [19]	2009	2002-2007	106	24.8	36.9	42.2	UC		71.6
Rink et al. [55]	2009	1990-2002	131	85	33	46.6	UC		84
Pricolo [51]	2009	1991–2008	312		36	46.8	Both		8

We compared incidences of the most important complications reported in studies published prior to 2000 versus the incidences reported in studies published since 2000 (Table 3). Pouch failure decreased between the two time periods from 6.8% to 4.3% (p=0.0038). Pelvic sepsis showed a decrease from 9.5 to 7.4% (p=0.1284). Other complications and the functional results showed no significant differences between the two groups of studies.

The effect of time of study conductance on IPAA outcomes

Results of both reviews were combined to analyze the effect of time of study conductance on the outcomes of the IPAA procedure. For this, the relationship between the median



year of inclusion of each study and the outcomes observed in that study was evaluated. Regression analysis showed that studies with an earlier median year of inclusion have a higher rate of pouch failure than studies with a more recent median year of inclusion (p<0.001). Figure 4 is a scatter plot depicting this relationship. In a sensitivity analysis excluding the five studies with a pouch failure rate of >12.5%, the relationship remained statistically significant (p=0.009). Additionally, we categorized the median year of inclusion of all studies (published before and after 2000) into four interquartile ranges in order to evaluate the median rate of pouch failure over these four time periods. Figure 5 shows a consecutive decline in the pouch failure rate over these time periods (p=0.005, one-way ANOVA analysis).

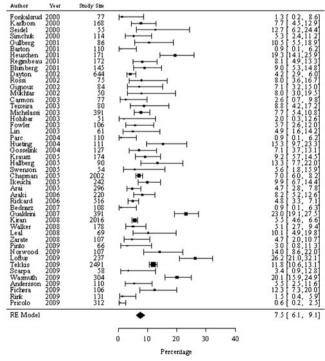


Fig. 2 Proportions of pouch failure following an IPAA procedure in studies published since 2000. The *size of the dot* correlates with the population size of the study. The *lines* represent the 95% CI. The pooled incidence of all studies is 4.3%

Fig. 3 Proportions of pelvic sepsis following an IPAA procedure in studies published since 2000. The *size of the dot* correlates with the population size of the study. The *lines* represent the 95% CI. The pooled incidence of all studies is 7.5%

	Since 2000			Before 2000	P value		
	Number of studies	Number of patients	Pooled incidences	Number of studies	No. of patients	Pooled incidences	
Complications			% (95% CI)			% (95%CI)	
Pouch failure	43	13,249	4.3 (3.5–5.3)	39	8,877	6.8 (5.8-8.4)	0.0038
Pouch failure: FU>5 years	22	9,873	4.7 (3.4–6.4)	11	3,198	8.5 (5.4–13.2)	
Pelvic sepsis	46	13,450	7.5 (6.1–9.1)	41	9,082	9.5 (8.2–10.9)	NS
Fistula	38	12,398	4.5 (3.5–5.7)	30	5,120	5.5 (4.3-7.0)	NS
Stricture	35	12,219	10.7 (8.2–13.8)	28	5,185	9.2 (6.8–12.4)	NS
Pouchitis	39	12,685	26.8 (21.0-33.5)	33	7,289	18.8 (15.7–22.4)	NS
Sexual dysfunction	13	6,131	3.0 (1.7-5.2)	21	5,112	3.6 (2.7-4.7)	NS
Small bowel obstruction	34	11,895	11.4 (9.1–14.1)	27	5,853	13.1 (11.0–15.7)	NS
Fecal incontinence			%(95%CI)			% (95%CI)	
Mild day incontinence	21	6,988	14.3 (7.3–25.9)	31	4,313	17.0 (12.8-22.2)	NS
Severe day incontinence	13	3,718	6.1 (2.9–12.3)	27	3,914	3.7 (2.8-4.8)	NS
Mild night incontinence	9	5,423	17.3 (4.7–46.8)	17	2,582	13.1 (9.5–17.9)	NS
Severe night incontinence	10	3,614	7.6 (2.5–21.3)	9	1,271	4.5 (3.0-6.7)	NS
Frequency			Mean (95%CI)			Mean (95%CI)	
Frequency day	26	5,321	5.7 (4.9-6.7)	13	2,277	5.2 (4.0-6.7)	NS
Frequency night	22	7,117	1.5 (1.0-2.1)	20	2,950	1.0 (0.6–1.6)	NS
Frequency 24 h	26	5,132	5.9 (5.0-6.9)	20	3,547	5.2 (4.4-6.1)	NS

 Table 3
 Pooled incidences of complications and functional outcome following an IPAA procedures in studies published since 2000 compared to studies published before 2000

CI confidence interval, FU follow-up period

Centralization: trends over time

The median inclusion rate for all studies was 14 patients per year (range from 2.9 to 105). We evaluated the effect of time

**Fig. 4** Scatter plot depicting the median year of inclusion and the rate of pouch failure of studies included in both reviews

of study conductance (median year of inclusion) on the inclusion rate (Fig. 6). The figure shows that between 1991 and 2000, an increase was seen in studies from centers including less than one patient per month, from eight studies

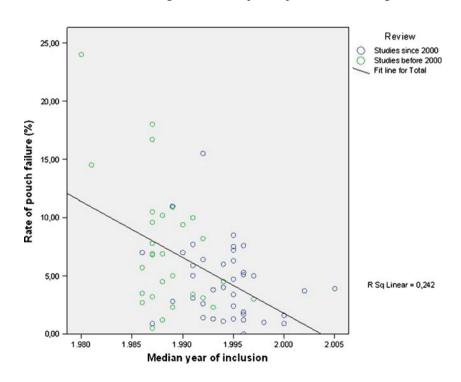
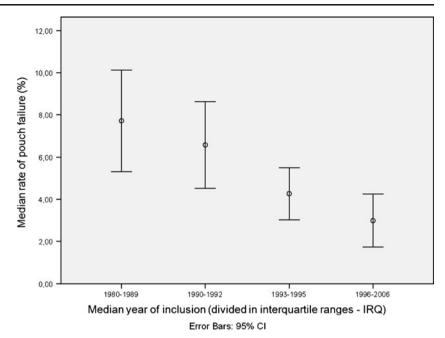


Fig. 5 Error bar chart depicting the median rate of pouch failure compared to the median year of inclusion (divided into interquartile ranges)



(30%) before 1991 to 23 studies (46%) between 1991 and 2000. It also shows that there are not yet enough studies available that have been conducted after 2000 to draw conclusions about the current situation. Analysis of the effect of the inclusion rate on the outcomes of the IPAA did not show any benefit on the outcomes of the IPAA.

## Technical aspects of the IPAA

Overall, 43 out of 78 studies (55%) used a protective ileostomy as a routine part of the IPAA procedure (routine use defined as ileostomy performed in more than 80% of patients). The percentage has decreased from almost 70% (16 out of 23) in studies conducted between 1980 and 1990 to 47% (21 out of 44) in studies conducted after 1990. The number of studies performing an ileostomy in <50% of patients increased from 11% to 32% over the same periods. The rate of ileostomy was not associated with any of the outcomes of the IPAA. Type of anastomosis (hand-sewn vs. stapled) was only available from 31 studies published since 2000 (the previous review did not register this characteristic). Fifteen studies (48%) still performed a hand-sewn anastomosis in more than 50% of

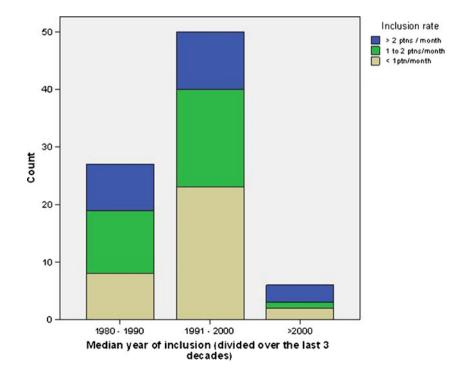


Fig. 6 Relationship between moment of study conductance (median year of inclusion) and inclusion rate of patients, divided in the past three decades patients. There were no differences in functional outcome or complications between studies performing different types of anastomosis.

#### Discussion

The open IPAA operation is the most common surgical reconstruction procedure in patients with UC and FAP. Randomized trials on IPAA are rare [1] due to low incidences as well as the complexity of patient selection and the intervention. Therefore, most studies are retrospective case series. Estimates of the outcomes of the IPAA procedure vary considerably. Acceptable complication and functional outcome rates are, therefore, difficult to set. This systematic review provides up to date and reliable incidences of important outcomes of the IPAA procedure, which could provide a reference values for these outcomes. Additionally, this review may provide insights in changes of practice over time and their effects on outcomes.

Several important finding are provided. First, it shows that the rate of pouch failure after an IPAA procedure has declined continuously over time. This decline was larger in the earlier period of the IPAA but seems to continue over time. Secondly, a large proportion of studies with this complex type of surgery are conducted in centers that include less than one patient per month. Thirdly, the functional outcome of the IPAA seems not to improve over time, despite several developments in surgical technique and perioperative care. This may represent an intrinsic limitation of the IPAA technique regarding functional outcome.

This review has some limitations. First, many of the included studies are retrospective cohort series, which may limit the reliability of the results. However, the overwhelming majority of the evidence in this field consists of this type of studies. This review provides an overview of this evidence, thereby increases its utility for clinicians and researchers. Moreover, the statistical techniques used are rather conservative to allow for incorporation of the heterogeneity of the studies.

Second, publication bias is a factor that should be taken into consideration. For this purpose, we excluded series with <50 patients. Studies with a small sample size do not only increase heterogeneity of results but are also more subject to publication bias; i.e., small studies with "unfavorable" results are less likely to get published. By introducing a threshold, we aimed to prevent including small studies with overestimated intervention effects.

Finally, the lack of individual patient data limits the analyses that could be performed. For example, many studies report a cohort in which several types of operative techniques have been used (e.g., several types of anastomses). This prohibits analyzing the effect of variations in technique on a patient level, since only aggregate outcomes of the complete cohorts are available. Collecting individual patient data of these large numbers of studies, especially considering the dated studies, is not feasible. We, therefore, limited our analyses to factors that could be analyzed reliably on an aggregate level.

Many factors may have contributed to the reduction of the rate of pouch failure observed. The additional analyses in this review were unable to pinpoint factors responsible for the observed reduction in pouch failure. This is probably caused by the lack of individual patient data (as discussed above), and we believe that general improvements in care over the past decades and the increased experience of surgeons with this approach are the most important factors contributing to this finding.

Another explanation that could be considered is the variation in the surgical techniques that have been introduced and sometimes largely employed over the last decades. Such variations include the various types of pouches used and the use of the double-stapled anastomosis. The routinely constructed deviating ileostoma does not seem to improve the frequency of postoperative complications but only their severity. Despite comparative studies and several large meta-analyses published previously, none of these variations in technique could be identified as factors associated with the reduction of postoperative complications, including pouch failure. [45, 69] This is in line with the findings in this review.

Fewer studies reported on other outcomes, which decreased the power of the analyses concerning these other outcomes. Secondly, it could be that the observed decrease in the rate of pouch failure may be, at least in part, a result of smaller decreases in various other postoperative complications. Complications like pelvic sepsis, fistulae, and strictures could all result in pouch failure, provided they were severe enough. Smaller decreases in these outcomes can be difficult to show statistically. Overall, however, they may have caused the observed substantial decrease in pouch failure rate.

Recently, discussions on the relation between volume of complex surgical interventions and outcomes have gained wide attention [7, 17, 27, 37]. Certain complex surgical interventions have been centralized in high-volume hospitals with improved outcomes [9–11, 31]. Centralization still has to be reflected in the data in the years to come. Therefore, this review was unable to evaluate the potential influence of this development on current practice. However, in the period from 1991 to 2000, nearly 50% of studies were performed in "low inclusion rate" centers (including less than one patient per month). This is a substantial increase from the 30% observed in the period from1980 to 1990. Taking into account that we only included series of 50 patients or more (Fig. 1) may mean that the true proportion

of studies conducted in such "low inclusion rate" centers may be much higher. With the repeatedly established relationship between volume and outcome in several fields of surgery, this is a troublesome observation. Centralization of the complex IPAA procedure may offer a way to further improve outcomes in future.

Finally, we found no statistically significant differences in pooled estimates of the functional outcome over time. There were also no differences in functional outcomes between various surgical factors studied in this review. This is in agreement with findings in literature showing no benefits in terms of functional outcome of technical developments of the IPAA procedure, like type of anastomosis [59] or laparoscopic approach [1]. In most patients, however, functional outcome of the IPAA is considered to be highly satisfactorily, allowing for an acceptable quality of life and social functioning [28, 65]. Therefore, reducing complications, especially severe complications like pelvic sepsis and pouch failure, seems to be a more important goal for future surgical developments.

#### Conclusions

This systematic review and meta-analysis summarizes the available evidence and provides up to date estimates for complications and functional outcome after an IPAA procedure that can be useful as reference values for practice and research. It also shows a reduction in pouch failure over time in patients operated on using the IPAA procedure. The functional results remain unchanged over time and between various operative factors.

Acknowledgment We did not receive any financial or material support for the research and the work. The first two authors have full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis.

**Open Access** This article is distributed under the terms of the Creative Commons Attribution Noncommercial License which permits any noncommercial use, distribution, and reproduction in any medium, provided the original author(s) and source are credited.

#### References

- Ahmed AU, Keus F, Heikens JT, Bemelman WA, Berdah SV, Gooszen HG, and van Laarhoven CJ (2009) Open versus laparoscopic (assisted) ileo pouch anal anastomosis for ulcerative colitis and familial adenomatous polyposis. Cochrane Database Syst Rev, CD006267
- Andersson T, Lunde OC, Johnson E, Moum T, and Nesbakken A (2011) Long-term functional outcome and quality of life after restorative proctocolectomy with ileo-anal anastomosis for colitis. Colorectal Dis 13:431–437
- Arai K, Koganei K, Kimura H, Akatani M, Kitoh F, Sugita A, Fukushima T (2005) Incidence and outcome of complications following restorative proctocolectomy. Am J Surg 190:39–42

- Araki T, Parc Y, Lefevre J, Dehni N, Tiret E, Parc R (2006) The effect on morbidity of mesentery lengthening techniques and the use of a covering stoma after ileoanal pouch surgery. Dis Colon Rectum 49:621–628
- Barton JG, Paden MA, Lane M, Postier RG (2001) Comparison of postoperative outcomes in ulcerative colitis and familial polyposis patients after ileoanal pouch operations. Am J Surg 182:616–620
- Bednarz W, Olewinski R, Woldan J (2007) Ileal-pouch-anal anastomosis after restorative proctocolectomy in patients with ulcerative colitis or familial adenomatous polyposis—11 years of experience. Polski Przeglad Chirurgiczny 79:652–662 (Portuguese, English)
- Begg CB, Cramer LD, Hoskins WJ, Brennan MF (1998) Impact of hospital volume on operative mortality for major cancer surgery. JAMA 280:1747–1751
- Berndtsson I, Lindholm E, Oresland T, Borjesson L (2007) Longterm outcome after ileal pouch-anal anastomosis: function and health-related quality of life. Dis Colon Rectum 50:1545–1552
- Birkmeyer JD, Siewers AE, Finlayson EV, Stukel TA, Lucas FL, Batista I, Welch HG, Wennberg DE (2002) Hospital volume and surgical mortality in the United States. N Engl J Med 346:1128–1137
- Birkmeyer JD, Sun Y, Wong SL, Stukel TA (2007) Hospital volume and late survival after cancer surgery. Ann Surg 245:777–783
- Birkmeyer NJ, Goodney PP, Stukel TA, Hillner BE, Birkmeyer JD (2005) Do cancer centers designated by the National Cancer Institute have better surgical outcomes? Cancer 103:435–441
- Blumberg D, Opelka FG, Hicks TC, Timmcke AE, Beck DE (2001) Restorative proctocolectomy: Ochsner Clinic experience. South Med J 94:467–471
- Bullard KM, Madoff RD, Gemlo BT (2002) Is ileoanal pouch function stable with time? Results of a prospective audit. Dis Colon Rectum 45:299–304
- 14. Carmon E, Keidar A, Ravid A, Goldman G, Rabau M (2003) The correlation between quality of life and functional outcome in ulcerative colitis patients after proctocolectomy ileal pouch anal anastomosis. Colorectal Dis 5:228–232
- Chapman JR, Larson DW, Wolff BG, Dozois EJ, Cima RR, Pemberton JH, Crownhart BS, Larson DR (2005) Ileal pouchanal anastomosis: does age at the time of surgery affect outcome? Arch Surg 140:534–539
- Dayton MT, Larsen KR, Christiansen DD (2002) Similar functional results and complications after ileal pouch-anal anastomosis in patients with indeterminate vs ulcerative colitis. Arch Surg 137:690–694
- Dudley RA, Johansen KL, Brand R, Rennie DJ, Milstein A (2000) Selective referral to high-volume hospitals: estimating potentially avoidable deaths. JAMA 283:1159–1166
- Ferrante M, Declerck S, De HG, Van AG, Geboes K, Rutgeerts P, Penninckx F, Vermeire S, D'Hoore A (2008) Outcome after proctocolectomy with ileal pouch-anal anastomosis for ulcerative colitis. Inflamm Bowel Dis 14:20–28
- Fichera A, Silvestri MT, Hurst RD, Rubin MA, Michelassi F (2009) Laparoscopic restorative proctocolectomy with ileal pouch anal anastomosis: a comparative observational study on long-term functional results. J Gastrointest Surg 13:526–532
- Fonkalsrud EW, Thakur A, Roof L (2000) Comparison of loop versus end ileostomy for fecal diversion after restorative proctocolectomy for ulcerative colitis. J Am Coll Surg 190:418–422
- Fowler AL, Turner BN, Thomson WH (2003) A study of the complications and pelvic visceral function after restorative proctocolectomy and W pouch construction. Colorectal Dis 5:342–346
- Gignoux B-M, Dehni N, Parc R, Tiret E (2002) Ileal pouch analanastomosis without covering ileostomy. [French]. Gastroenterol Clin Biol 26:671–674
- 23. Gosselink MP, Schouten WR, van Lieshout LM, Hop WC, Laman JD, Ruseler-van Embden JG (2004) Delay of the first onset of

pouchitis by oral intake of the probiotic strain *Lactobacillus rhamnosus* GG. Dis Colon Rectum 47:876–884

- Gualdrini U, Coraglio M, Filippa L, Masciangioli G, Gutierrez A, Lumi C, Graziano A (2007) Long term functional outcomes after ileoanal pouch in patients with and without postoperative complications]. Acta Gastroenterol Latinoam 37(3):143–149
- Gullberg K, Liljeqvist L (2001) Stapled ileoanal pouches without loop ileostomy: a prospective study in 86 patients. Int J Colorectal Dis 16:221–227
- Hallberg H, Stahlberg D, Akerlund JE (2005) Ileal pouch-anal anastomosis (IPAA): functional outcome after postoperative pelvic sepsis. A prospective study of 100 patients. Int J Colorectal Dis 20:529–533
- Halm EA, Lee C, Chassin MR (2002) Is volume related to outcome in health care? A systematic review and methodologic critique of the literature. Ann Intern Med 137:511–520
- 28. Heikens JT, de VJ, and van Laarhoven CJ (2011) Quality of life, health related quality of life and health status in patients having restorative proctocolectomy with ileal pouch-anal anastomosis for ulcerative colitis: a systematic review. Colorectal Dis 2010 Dec 22 doi:10.1111/j.1463-1318.2010.02538.x
- Heuschen UA, Hinz U, Allemeyer EH, Lucas M, Heuschen G, Herfarth C (2001) One- or two-stage procedure for restorative proctocolectomy: rationale for a surgical strategy in ulcerative colitis. Ann Surg 234:788–794
- 30. Ho KS, Chang CC, Baig MK, Borjesson L, Nogueras JJ, Efron J, Weiss EG, Sands D, Vernava AM III, Wexner SD (2006) Ileal pouch anal anastomosis for ulcerative colitis is feasible for septuagenarians. Colorectal Dis 8:235–238
- Hollenbeck BK, Wei Y, Birkmeyer JD (2007) Volume, process of care, and operative mortality for cystectomy for bladder cancer. Urology 69:871–875
- Holubar S, Hyman N (2003) Continence alterations after ileal pouch-anal anastomosis do not diminish quality of life. Dis Colon Rectum 46:1489–1491
- 33. Hueting WE, Buskens E, van der Tweel I, Gooszen HG, van Laarhoven CJ (2005) Results and complications after ileal pouch anal anastomosis: a meta-analysis of 43 observational studies comprising 9,317 patients. Dig Surg 22:69–79
- 34. Hueting WE, Gooszen HG, van Laarhoven CJ (2004) Sexual function and continence after ileo pouch anal anastomosis: a comparison between a meta-analysis and a questionnaire survey. Int J Colorectal Dis 19:215–218
- Ikeuchi H, Nakano H, Uchino M, Nakamura M, Noda M, Yanagi H, Yamamura T (2005) Safety of one-stage restorative proctocolectomy for ulcerative colitis. Dis Colon Rectum 48:1550–1555
- 36. Karlbom U, Raab Y, Ejerblad S, Graf W, Thorn M, Pahlman L (2000) Factors influencing the functional outcome of restorative proctocolectomy in ulcerative colitis. Br J Surg 87:1401–1408
- Killeen SD, O'Sullivan MJ, Coffey JC, Kirwan WO, Redmond HP (2005) Provider volume and outcomes for oncological procedures. Br J Surg 92:389–402
- Kiran RP, Remzi FH, Fazio VW, Lavery IC, Church JM, Strong SA, Hull TL (2008) Complications and functional results after ileoanal pouch formation in obese patients. J Gastrointest Surg 12:668–674
- 39. Knapp G, Hartung J (2003) Improved tests for a random effects meta-regression with a single covariate. Stat Med 22:2693–2710
- 40. Krausz MM, Duek SD (2005) Restorative proctocolectomy with ileal pouch-anal anastomosis for ulcerative colitis and familial adenomatous polyposis: twenty years follow-up in 174 patients. Isr Med Assoc J 7:23–27
- Leal RF, Ayrizono MDLS, Coy CSR, Fagundes JJ, Goes JRN (2008) Short-term and long-term postoperative complications after ileal pouch-anal anastomosis in familial adenomatous polyposis. Arq Gastroenterol 45:106–110 (Portuguese)
- 42. Leowardi C, Hinz U, Tariverdian M, Kienle P, Herfarth C, Ulrich A, Kadmon M (2010) Long-term outcome 10 years or more after

🖄 Springer

restorative proctocolectomy and ileal pouch-anal anastomosis in patients with ulcerative colitis. Langenbecks Arch Surg 395:49–56

- Lin JJ, Song ZF, Xu JH (2004) [The prognosis of the total proctocolectomy and ileal-pouch anal anastomosis. Zhonghua Wai Ke Za Zhi 42(14):861–863
- 44. Loftus EV Jr, Friedman HS, Delgado DJ, Sandborn WJ (2009) Colectomy subtypes, follow-up surgical procedures, postsurgical complications, and medical charges among ulcerative colitis patients with private health insurance in the United States. Inflamm Bowel Dis 15:566–575
- 45. Lovegrove RE, Constantinides VA, Heriot AG, Athanasiou T, Darzi A, Remzi FH, Nicholls RJ, Fazio VW, Tekkis PP (2006) A comparison of hand-sewn versus stapled ileal pouch anal anastomosis (IPAA) following proctocolectomy: a meta-analysis of 4183 patients. Ann Surg 244:18–26
- 46. Michelassi F, Lee J, Rubin M, Fichera A, Kasza K, Karrison T, Hurst RD (2003) Long-term functional results after ileal pouch anal restorative proctocolectomy for ulcerative colitis: a prospective observational study. Ann Surg 238:433–441
- Mukhtar H, Zak A, Berry AR (2002) Restorative proctocolectomy: one surgeon's experience of a now routine procedure in a district general hospital. JR Coll Surg Edinb 47:400–406
- Norwood MG, Mann CD, West K, Miller AS, Hemingway D (2009) Restorative proctocolectomy. Does ethnicity affect outcome? Colorectal Dis 11:972–975
- Parc Y, Piquard A, Dozois RR, Parc R, Tiret E (2004) Long-term outcome of familial adenomatous polyposis patients after restorative coloproctectomy. Ann Surg 239:378–382
- 50. Pinto RA, Canedo J, Murad-Regadas S, Regadas SF, Weiss EG, Wexner SD (2009) Ileal pouch anal anastomosis (IPAA) in the elderly: is there difference in the morbidity compared to younger patients? Colorectal Dis 13:177–183
- Pricolo VE (2009) Ileal pouch-anal anastomosis for ulcerative colitis—the Rhode Island experience. Med Health R I 92:100–102
- 52. R Development Core Team (2010) R: A language and environment for statistical computing. R Foundation forStatistical Computing, Vienna
- 53. Regimbeau JM, Panis Y, Pocard M, Hautefeuille P, Valleur P (2001) Handsewn ileal pouch-anal anastomosis on the dentate line after total proctectomy: technique to avoid incomplete mucosectomy and the need for long-term follow-up of the anal transition zone. Dis Colon Rectum 44:43–50
- Rickard MJFX, Young CJ, Bissett IP, Stitz R, Solomon MJ (2007) Ileal pouch-anal anastomosis: the Australasian experience. Color Dis 9:139–145
- Rink AD, Radinski I, Vestweber KH (2009) Does mesorectal preservation protect the ileoanal anastomosis after restorative proctocolectomy? J Gastrointest Surg 13:120–128
- Rossi HL, Brand MI, Saclarides TJ (2002) Anal complications after restorative proctocolectomy (J-pouch). Am Surg 68:628–630
- Rudolph WG, Uthoff SM, McAuliffe TL, Goode ET, Petras RE, Galandiuk S (2002) Indeterminate colitis: the real story. Dis Colon Rectum 45:1528–1534
- 58. Scarpa M, Mescoli C, Rugge M, D'Inca R, Ruffolo C, Polese L, D'Amico DF, Sturniolo GC, Angriman I (2009) Restorative proctocolectomy for inflammatory bowel disease: the Padova prognostic score for colitis in predicting long-term outcome and quality of life. Int J Colorectal Dis 24:1049–1057
- 59. Schluender SJ, Mei L, Yang H, Fleshner PR (2006) Can a metaanalysis answer the question: is mucosectomy and handsewn or double-stapled anastomosis better in ileal pouch-anal anastomosis? Am Surg 72:912–916
- Seidel SA, Newman M, Sharp KW (2000) Ileoanal pouch versus ileostomy: is there a difference in quality of life? Am Surg 66:540–546
- Simchuk EJ, Thirlby RC (2000) Risk factors and true incidence of pouchitis in patients after ileal pouch-anal anastomoses. World J Surg 24:851–856

- Swenson BR, Hollenbeak CS, Poritz LS, Koltun WA (2005) Modified two-stage ileal pouch-anal anastomosis: equivalent outcomes with less resource utilization. Dis Colon Rectum 48:256–261
- 63. Teixeira MG, Ponte AC, Sousa M, Almeida MG, Silva FE, Calache JE, Habr-Gama A, Kiss DR (2003) Short- and long-term outcomes of ileal pouch-anal anastomosis for ulcerative colitis. Rev Hosp Clin Fac Med Sao Paulo 58:193–198
- 64. Tekkis PP, Lovegrove RE, Tilney HS, Smith JJ, Sagar PM, Shorthouse AJ, Mortensen NJ, and Nicholls RJ (2010) Longterm failure and function after restorative proctocolectomy—a multi-centre study of patients from the UK National Ileal Pouch Registry. Colorectal Dis 12:433–441
- 65. Thirlby RC, Sobrino MA, Randall JB (2001) The long-term benefit of surgery on health-related quality of life in patients with inflammatory bowel disease. Arch Surg 136:521–527
- 66. Tulchinsky H, Dotan I, Alper A, Brazowski E, Klausner JM, Halpern Z, Rabau M (2008) Comprehensive pouch clinic concept for follow-up of patients after ileal pouch anal anastomosis: report

of 3 years' experience in a tertiary referral center. Inflamm Bowel Dis 14:1125–1132

- Walker LR, Bulow S (2008) Restorative proctocolectomy with an ileoanal pouch. Postoperative course and long-term functional results. Ugeskr Laege 170(20):1721–1725
- Wasmuth HH, Trano G, Endreseth B, Rydning A, Wibe A, Myrvold HE (2009) Long-term surgical load in patients with ileal pouch-anal anastomosis. Color Dis 11:711–718
- 69. Weston-Petrides GK, Lovegrove RE, Tilney HS, Heriot AG, Nicholls RJ, Mortensen NJ, Fazio VW, Tekkis PP (2008) Comparison of outcomes after restorative proctocolectomy with or without defunctioning ileostomy. Arch Surg 143:406–412
- Viechtbauer W (2010) Conducting meta-analyses in R with the Metafor Package. J Stat Softw 36:1–48
- Zarate CA, Zuniga DA, Pinedo MG, Lopez KF, Molina PM, Viviani GP (2008) Ileal pouch-anal anastomosis for ulcerative colitis: complications and long term functional results. Rev Med Chil 136:467–474