

# **A Systematic Review of Epidemiology and Outcomes Associated with Local Surgical and Intersphincteric Ligation Procedures for Complex Cryptoglandular Fistulas**

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**Supplementary Table S1. Inclusion Criteria for Systematic Review**

	Inclusion criteria
Population(s)	<ul style="list-style-type: none"> <li>• Human patients with cryptoglandular or Crohn’s-related rectovaginal, anovaginal or enterocutaneous fistulas</li> </ul> <p><i>Note: The current report is limited to those studies assessing complex cryptoglandular fistula</i></p> <ul style="list-style-type: none"> <li>• No restrictions regarding other demographic characteristics</li> </ul>
Intervention(s)	<ul style="list-style-type: none"> <li>• No restrictions for assessment of incidence/prevalence or HCRU/costs</li> <li>• Treatment patterns, ClinROs and PROs will focus on the following interventions:               <ul style="list-style-type: none"> <li>- Pharmacologic: antibiotics, immunosuppressants, corticosteroids, prednisone, methotrexate, anti-tumor necrosis factor (infliximab, adalimumab, certolizumab pegol), monoclonal antibody (natalizumab, vedolizumab), interleukin antagonist (ustekinumab)</li> <li>- Surgical: fistulotomy, ligation of the intersphincteric fistula tract, cutting seton, fibrosing seton, drainage seton, sphincteric reconstruction, flap, ablation, fibrin glue, anal fistula plug, bioprosthesis plug, surgical reconstruction, sphincterotomy, sphincteroplasty, proctectomy, diversion, ostomy</li> </ul> </li> </ul> <p><i>Note: The current report is limited to the following surgical treatments: fistulotomy, fistulectomy, Modified Park’s Technique, Advancement flap, ligation of the intersphincteric fistula tract (LIFT), BioLift, and transanal opening of intersphincteric space (TROPIS).</i></p>
Comparison(s)	<ul style="list-style-type: none"> <li>• No restrictions on comparators for assessment of incidence/prevalence or HCRU/costs</li> <li>• Assessment of treatment patterns in studies that include the pharmacologic or surgical interventions listed above</li> </ul>

	<ul style="list-style-type: none"> <li>• Clinical and patient-reported outcomes in single arm or comparator studies assessing the interventions listed above</li> </ul>
Outcome(s)	<ul style="list-style-type: none"> <li>• Incidence and prevalence</li> <li>• HCRU: Outpatient physician visits, Inpatient/hospital admissions, Emergency room visits, Laboratory tests, Diagnostic and therapeutic tests, length of stay</li> <li>• Direct and indirect costs</li> <li>• Treatment patterns: Proportion of patients utilizing pre-specified surgery or pharmacologic treatment</li> <li>• Clinical outcomes: healing rate, success rate, response rate, recurrence rate, fistula closure, clinical response, clinical remission</li> <li>• Specific PRO instruments of interest: Crohn's Disease Activity Index (CDAI), Inflammatory bowel disease questionnaire (IBDQ), EQ-5D</li> <li>• Other PRO Categories: pain</li> </ul> <p><i>Note: The current report is limited to the following outcomes: healing, recurrence, incontinence, and pain.</i></p>
Time	<ul style="list-style-type: none"> <li>• No restriction on duration of illness or treatment</li> <li>• Publications within the past 10 years</li> </ul> <p><i>Note: The current report is limited to publications within the past 5 years.</i></p>
Study design	<ul style="list-style-type: none"> <li>• Observational studies: case-control studies, retrospective or prospective cohort studies/registries, cross-sectional studies</li> <li>• Pooled data excluded; however, studies identified in those reports would be reviewed as part of the manual search for inclusion. Other publications that are not original research, clinical trials, case reports/series* are excluded</li> </ul>
Other-language	<ul style="list-style-type: none"> <li>• English language</li> </ul>

\*Case series are included if they meet pre-specified criteria for cohort design.

## Supplemental Table S2. PubMed Search Strategy

PubMed search strategy (conducted on 3/25/2020)

Row	Search concept	Terms
<b>Conditions</b>		
#1	<b>Condition 1:</b> Complex cryptoglandular fistulas (CCF) and/or cryptoglandular fistulas (CF)	((cryptoglandular[Title/Abstract] OR cryptogland*[Title/Abstract] OR horseshoe[Title/Abstract] OR branching[Title/Abstract]) AND ("Fistula"[Mesh] OR "fistula"[Title/Abstract] OR fistul*[Title/Abstract])) OR  ((mid[Title/Abstract] OR high[Title/Abstract]) AND (transphincteric[Title/Abstract] OR trans- sphincteric[Title/Abstract] OR transsphincteric[Title/Abstract]) AND ("Fistula"[Mesh] OR "fistula"[Title/Abstract] OR fistul*[Title/Abstract]))
#2	<b>Condition 2a:</b> Rectovaginal fistula	"Rectovaginal Fistula"[Mesh] OR  (("rectovaginal"[Title/Abstract] OR "recto- vaginal"[Title/Abstract]) AND ("Fistula"[Mesh] OR "fistula"[Title/Abstract] OR fistul*[Title/Abstract]))
	<b>Condition 2b:</b> Anovaginal fistula	(("anovaginal"[Title/Abstract] OR "ano- vaginal"[Title/Abstract]) AND ("Fistula"[Mesh] OR "fistula"[Title/Abstract] OR fistul*[Title/Abstract]))
	<b>Condition 2c:</b> Enterocutaneous fistula	(("enterocutaneous"[Title/Abstract] OR "entero- cutaneous"[Title/Abstract]) AND ("Fistula"[Mesh] OR "fistula"[Title/Abstract] OR fistul*[Title/Abstract]))

	<b>Condition 2 combined</b> 2a OR 2b OR 2c	(((“Rectovaginal Fistula”[Mesh] OR (“rectovaginal”[Title/Abstract] OR “recto- vaginal”[Title/Abstract]) AND (“Fistula”[Mesh] OR “fistula”[Title/Abstract] OR fistul*[Title/Abstract]))) OR (((“anovaginal”[Title/Abstract] OR “ano- vaginal”[Title/Abstract]) AND (“Fistula”[Mesh] OR “fistula”[Title/Abstract] OR fistul*[Title/Abstract]))) OR (((“enterocutaneous”[Title/Abstract] OR “entero- cutaneous”[Title/Abstract]) AND (“Fistula”[Mesh] OR “fistula”[Title/Abstract] OR fistul*[Title/Abstract])))
#3	Crohns terms	“Crohn Disease”[Mesh] OR crohn*[Title/Abstract]
#4	Fistula (for title filter)	Fist*[Title]
<b>Real World Data Terms</b>		
#5	Observational studies and real world data	(“Cohort Studies”[Mesh] OR “Cross-Sectional Studies”[Mesh] OR “Longitudinal Studies”[Mesh] OR “Prospective Studies”[Mesh] OR “case-control study”[Mesh] OR “Registries”[Mesh] OR “Electronic Health Records”[Mesh] OR “Administrative Claims, Healthcare”[Mesh] OR “Observational Study” [Publication Type] OR  cohort[Title/Abstract] OR cohort*[Title/Abstract] OR cross-sectional[Title/Abstract] OR cross- section*[Title/Abstract] OR longitudinal[Title/Abstract] OR longitud*[Title/Abstract] OR prospective[Title/Abstract] OR

		<p>prospect*[Title/Abstract] OR  retrospective[Title/Abstract] OR  observational[Title/Abstract] OR  observation*[Title/Abstract] OR  registry[Title/Abstract] OR “electronic medical  record”[Title/Abstract] OR “medical  record”[Title/Abstract] OR “electronic health  record”[Title/Abstract] OR “health  record”[Title/Abstract] OR (claims[Title/Abstract]  OR “claims data”[Title/Abstract] OR “administrative  claims”[Title/Abstract] OR “systematic  review”[Title/Abstract] OR “systematic literature  review”[Title/Abstract] OR “meta-  analysis”[Title/Abstract])</p>
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**Queries of Interest**

#6	Measures of occurrence/association	<p>“Epidemiology”[Mesh] OR  “Pharmacoepidemiology”[Mesh] OR    epidemiolog*[Title/Abstract] OR  pharmacoepidemiolog*[Title/Abstract] OR  prevalence[Title/Abstract] OR  incidence[Title/Abstract] OR risk[Title/Abstract] OR  rate[Title/Abstract] OR rates[Title/Abstract] OR  proportion[Title/Abstract] OR  proportions[Title/Abstract] OR  frequency[Title/Abstract] OR  frequencies[Title/Abstract]</p>
#7	Health care resource utilization and costs	<p>“Costs and Cost Analysis”[Mesh] OR  “Economics”[Mesh] OR “Health  Expenditures”[Mesh] OR “Cost of Illness”[Mesh]  OR “Cost-Benefit Analysis”[Mesh] OR “Health Care</p>

		<p>Costs"[Mesh] OR "Direct Service Costs"[Mesh] OR "Hospital Costs"[Mesh] OR "Drug Costs"[Mesh] OR "Health Resources"[Mesh] OR "Resource Allocation"[Mesh] OR</p> <p>"cost"[Title/Abstract] OR "costs"[Title/Abstract] OR "cost analysis"[Title/Abstract] OR "resource use"[Title/Abstract] OR "resource utilization"[Title/Abstract] OR expenditure[Title/Abstract] OR expenditures[Title/Abstract] OR economic[Title/Abstract] OR economics[Title/Abstract]</p>
#8	Treatment Patterns	<p>"Immunosuppression"[Mesh] OR "Immunosuppressive Agents"[Mesh] OR "Immunosuppressive Agents" [Pharmacological Action] OR "Anti-Bacterial Agents"[Mesh] OR "Anti-Bacterial Agents" [Pharmacological Action] OR "Antibodies, Monoclonal"[Mesh] OR "Surgical Procedures, Operative"[Mesh] OR "surgery" [Subheading] OR "Ostomy"[Mesh] OR</p> <p>(immunosuppress*[Title/Abstract] OR antibiotic[Title/Abstract] OR antibiotics[Title/Abstract] OR corticosteroid*[Title/Abstract] OR prednisone[Title/Abstract] OR Methotrexate[Title/Abstract] OR anti-TNF[Title/Abstract] OR "anti-tumor necrosis factor"[Title/Abstract] OR infliximab[Title/Abstract] OR adalimumab[Title/Abstract] OR "certolizumab</p>

	<p>pegol"[Title/Abstract] OR certolizumab[Title/Abstract] OR "monoclonal antibody"[Title/Abstract] OR "monoclonal antibodies"[Title/Abstract] OR Natalizumab[Title/Abstract] OR vedolizumab[Title/Abstract] OR "interleukin antagonist"[Title/Abstract] OR "interleukin antagonists"[Title/Abstract] OR "interleukin inhibitor"[Title/Abstract] OR "interleukin inhibitors"[Title/Abstract] OR Ustekinumab[Title/Abstract] OR cyclosporine[Title/Abstract] OR tacrolimus[Title/Abstract] OR</p> <p>surgery[Title/Abstract] OR Fistulotomy[Title/Abstract] OR "ligation of the intersphincteric fistula tract"[Title/Abstract] OR LIFT[Title/Abstract] OR ligation*[Title/Abstract] OR seton[Title/Abstract] OR "cutting seton"[Title/Abstract] OR "fibrosing seton"[Title/Abstract] OR "drainage seton" OR "seton placement"[Title/Abstract] OR flap[Title/Abstract] OR "advancement flap"[Title/Abstract] OR "advancement flaps"[Title/Abstract] OR ablation[Title/Abstract] OR "fibrin glue"[Title/Abstract] OR "bioprosthetic plug"[Title/Abstract] OR "anal fistula plug"[Title/Abstract] OR plug[Title/Abstract] OR plugs[Title/Abstract] OR sealant [Title/Abstract] OR sealants[Title/Abstract] OR "surgical reconstruction"[Title/Abstract] OR "sphincteric</p>
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		reconstruction"[Title/Abstract] OR sphincterotomy[Title/Abstract] OR sphincteroplasty[Title/Abstract] OR proctectomy[Title/Abstract] OR diversion[Title/Abstract] OR therapy[Title/Abstract] OR therapeutic*[Title/Abstract] OR pattern*[Title/Abstract] OR "treatment pattern" [Title/Abstract] OR "treatment patterns" [Title/Abstract] OR "ostomy"[Title/Abstract] OR ostomies[Title/Abstract]OR enterostomy[Title/Abstract] OR enterostomies[Title/Abstract] OR Cecostomy[Title/Abstract] OR Cecostomies[Title/Abstract] OR colostomy[Title/Abstract] OR colostomies[Title/Abstract] OR duodenostomy[Title/Abstract] OR duodenostomies[Title/Abstract] OR ileostomy[Title/Abstract] OR ileostomies[Title/Abstract] OR jejunostomy[Title/Abstract] OR jejunostomies[Title/Abstract])
#9	Clinical outcomes	"Treatment Outcome"[Mesh] OR (("healing"[Title/Abstract] OR "success"[Title/Abstract] OR "response"[Title/Abstract] OR "recurrence"[Title/Abstract] OR "remission"[Title/Abstract] OR "incontinence"[Title/Abstract]) AND

		<p>(“rate”[Title/Abstract] OR  “frequency”[Title/Abstract] OR  “proportion”[Title/Abstract])) OR  closure[Title/Abstract] OR  “recurrence”[Title/Abstract] OR “recurrence  risk”[Title/Abstract] OR “remission”[Title/Abstract]  OR “remission risk”[Title/Abstract] OR  “incontinence”[Title/Abstract] OR “incontinence  risk”[Title/Abstract] OR “clinical  response”[Title/Abstract] OR “clinical  remission”[Title/Abstract]</p>
#10	Patient-reported outcomes	<p>“Patient Reported Outcome Measures”[Mesh] OR  “Fecal Incontinence”[Mesh] OR</p> <p>(“patient reported outcomes”[Title/Abstract] OR  “clinical outcome”[Title/Abstract] OR “Crohn’s  Disease Activity Index”[Title/Abstract] OR  CDAI[Title/Abstract] OR “IBD  questionnaire”[Title/Abstract] OR  “IBDQ”[Title/Abstract] OR “inflammatory bowel  disease questionnaire”[Title/Abstract] OR “Patient  reported outcome measures”[Title/Abstract] OR  PROMs[Title/Abstract] OR “perianal disease activity  index”[Title/Abstract] OR “PDAI”[Title/Abstract]  OR “CDAI”[Title/Abstract] OR “anal  pain”[Title/Abstract] OR pain[Title/Abstract] OR  “fecal incontinence”[Title/Abstract] OR “faecal  incontinence”[Title/Abstract] OR “bowel  incontinence”[Title/Abstract] OR “fecal  soiling”[Title/Abstract] OR “faecal</p>

		soiling"[Title/Abstract] OR "leakage"[Title/Abstract] OR "discharge"[Title/Abstract] OR "soiling"[Title/Abstract] OR "soil"[Title/Abstract] OR "soiled"[Title/Abstract] OR "pads"[Title/Abstract] OR "EQ-5D"[Title/Abstract] OR "EuroQol"[Title/Abstract] OR "EuroQol-5D"[Title/Abstract] OR "intercourse"[Title/Abstract] OR "sexual activity"[Title/Abstract] OR "Wexner Scale of incontinence"[Title/Abstract] OR "Revised Fecal Incontinence Scale"[Title/Abstract] OR "Revised Faecal Incontinence Scale"[Title/Abstract] OR "RFIS"[Title/Abstract])
<b>Combination Searches</b>		
#11	Condition 2 + Crohn's	(#2a OR 2b OR 2c) AND #3
#12	Combined Queries	#6 OR #7 OR #8 OR #9 OR #10
<b>SEARCH RESULTS</b>		
#13	Condition 2 + Combined Queries + RWD + filters	#11 AND #12 AND #5 +English, human, 10 years filters
#14	Condition 1 + Combined Queries + RWD + filters	#1 AND #12 AND #5 +English, human, 10 years filters
#15	Final search including fistula title search	(#13 OR #14) AND #4

### Supplemental Table S3. Embase Search Strategy

EMBASE search strategy (conducted on 3/25/2020)

Row	Search concept	Terms
<b>Conditions</b>		
#1	<b>Condition 1:</b> Complex cryptoglandular fistulas (CCF) and/or cryptoglandular fistulas (CF)	'cryptoglandular fistula':ab,ti,kw OR  ((cryptoglandular:ti,ab,kw OR horshoe:ab,ti,kw OR branching:ab,ti,kw) AND fistul*:ti,ab,kw) OR  ((mid:ab,ti,kw OR high:ab,ti,kw) AND (transphincteric:ab,ti,kw OR trans- sphincteric:ab,ti,kw OR transsphincteric:ab,ti,kw) AND fistul*:ti,ab,kw)
#2	<b>Condition 2:</b> Non Perianal Crohn's Fistula (NPCF): Rectovaginal Fistulas (RVF), Ano-vaginal Fistulas (AVF), Entero-cutaneous Fistulas (ECF)	'rectovaginal fistula':ti,ab,kw OR 'anovaginal fistula':ti,ab,kw OR 'enterocutaneous fistula':ti,ab,kw OR  ((rectovaginal:ab,ti,kw OR recto-vaginal:ab,ti,kw OR anovaginal:ab,ti,kw OR ano-vaginal:ab,ti,kw OR enterocutaneous:ab,ti,kw OR entero-cutaneous:ab,ti,kw) AND fistul*:ab,ti,kw)
#3	Crohn terms	'crohn disease':ab,ti,kw OR crohn*:ab,ti,kw
#4	Fistula (for title filter)	Fist*:ti
<b>Real World Data Terms</b>		
#5	Observational studies and real world data	'cohort analysis':ab,ti,kw OR 'cross-sectional study':ab,ti,kw OR 'cross sectional analysis':ab,ti,kw OR 'longitudinal study':ab,ti,kw OR 'prospective study':ab,ti,kw OR 'case control study':ab,ti,kw OR 'electronic medical

		<p>record':ab,ti,kw OR 'electronic health record':ab,ti,kw OR 'administrative claims':ab,ti,kw OR 'observational study':ab,ti,kw OR</p> <p>'cohort':ab,ti,kw OR cohort*:ab,ti,kw OR 'cross-sectional':ab,ti,kw OR 'longitudinal':ab,ti,kw OR 'prospective':ab,ti,kw OR 'retrospective':ab,ti,kw OR 'observational':ab,ti,kw OR 'registry':ab,ti,kw OR 'register':ab,ti,kw OR 'registries':ab,ti,kw OR 'medical record':ab,ti,kw OR 'health record':ab,ti,kw OR 'medical record':ab,ti,kw OR 'claims data':ab,ti,kw OR 'administrative claims':ab,ti,kw OR 'claims':ab,ti,kw OR 'systematic review':ab,ti,kw OR 'systematic literature review':ab,ti,kw OR 'meta-analysis':ab,ti,kw OR 'meta analysis':ab,ti,kw</p>
<b>Queries of Interest</b>		
#6	Measures of occurrence/association	<p>epidemiolog*:ti,ab,kw OR pharmacoepidemiolog*:ti,ab,kw OR prevalence:ti,ab,kw OR incidence:ti,ab,kw OR rate:ti,ab,kw OR rates:ti,ab,kw OR risk:ti,ab,kw OR proportion:ti,ab,kw OR proportions:ti,ab,kw OR frequency:ti,ab,kw OR frequencies:ti,ab,kw</p>
#7	Health care resource utilization and costs	<p>'resource utilization':ti,ab,kw OR 'resource use':ti,ab,kw OR 'resource allocation':ti,ab,kw OR 'health resources':ti,ab,kw OR 'health care utilization':ti,ab,kw OR cost:ti,ab,kw OR 'cost analysis':ti,ab,kw OR economic:ti,ab,kw OR expenditures:ti,ab,kw OR 'health care cost':ti,ab,kw</p>
#8	Treatment Patterns	<p>'immunosuppressive agent':ti,ab,kw OR 'antibiotic drug':ti,ab,kw OR</p>

		<p>corticosteroid:ti,ab,kw OR prednisone:ti,ab,kw OR methotrexate:ti,ab,kw OR ‘anti tnf’:ti,ab,kw OR ‘tumor necrosis factor inhibitor’:ti,ab,kw OR infliximab:ti,ab,kw OR adalimumab:ti,ab,kw OR ‘certolizumab pegol’:ti,ab,kw OR ‘certolizumab’:ti,ab,kw OR ‘monoclonal antibody’:ti,ab,kw OR natalizumab:ti,ab,kw OR vedolizumab:ti,ab,kw OR ‘interleukin antagonist’ OR ‘interleukin inhibitor’:ti,ab,kw OR ustekinumab:ti,ab,kw OR</p> <p>surgery:ti,ab,kw OR surgical:ti,ab,kw OR fistulotomy:ti,ab,kw OR sphincterotomy:ti,ab,kw OR sphincteroplasty:ti,ab,kw OR proctectomy:ti,ab,kw OR ligation:ti,ab,kw OR ‘ligation of the intersphincteric fistula tract’:ti,ab,kw OR lift:ti,ab,kw OR seton:ti,ab,kw OR ‘seton placement’:ti,ab,kw OR flap:ti,ab,kw OR ‘advancement flap’:ti,ab,kw OR ablation:ti,ab,kw OR ‘fibrin glue’:ti,ab,kw OR ‘bioprosthetic plug’:ti,ab,kw OR patterns:ti,ab,kw OR ‘treatment pattern’:ti,ab,kw OR ‘treatment patterns’:ti,ab,kw OR therapy:ti,ab,kw OR therapeutic:ti,ab,kw OR ‘cutting seton’:ti,ab,kw OR ‘fibrosing seton’:ti,ab,kw OR ‘drainage seton’:ti,ab,kw OR ‘sphincteric reconstruction’:ti,ab,kw OR ‘anal fistula plug’:ti,ab,kw OR sealant:ti,ab,kw OR ‘surgical reconstruction’:ti,ab,kw OR</p>
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		ostomy:ti,ab,kw OR ostomies:ti,ab,kw OR enterostomy:ti,ab,kw OR enterostomies:ti,ab,kw OR cecostomy:ti,ab,kw OR cecostomies:ti,ab,kw OR colostomy:ti,ab,kw OR colostomies:ti,ab,kw OR duodenostomy:ti,ab,kw OR duodenostomies:ti,ab,kw OR ileostomy:ti,ab,kw OR ileostomies:ti,ab,kw OR jejunostomy:ti,ab,kw OR jejunostomies:ti,ab,kw
#9	Clinical outcomes	'clinical outcome':ti,ab,kw OR 'clinical response':ti,ab,kw OR remission:ti,ab,kw OR recurrence:ti,ab,kw OR closure:ti,ab,kw OR 'incontinence':ti,ab,kw OR 'healing rate':ti,ab,kw OR 'success rate':ti,ab,kw OR 'response rate':ti,ab,kw OR 'recurrence rate':ti,ab,kw OR 'recurrence risk':ti,ab,kw OR 'remission rate':ti,ab,kw OR 'remission risk':ti,ab,kw OR 'incontinence rate':ti,ab,kw OR 'incontinence risk':ti,ab,kw
#10	Patient-reported outcomes	'patient-reported outcome':ti,ab,kw OR 'crohns disease patient-reported outcomes signs':ti,ab,kw OR 'cd pro':ti,ab,kw OR 'crohn disease activity index':ti,ab,kw OR cdai:ti,ab,kw OR 'ibd questionnaire':ti,ab,kw OR ibdq:ti,ab,kw OR 'inflammatory bowel disease questionnaire':ti,ab,kw OR 'patient reported outcome measure':ti,ab,kw OR proms:ti,ab,kw OR 'perianal disease activity index':ti,ab,kw OR PDAI:ti,ab,kw OR discharge:ti,ab,kw OR soiling:ti,ab,kw OR soil:ti,ab,kw OR soiled:ti,ab,kw OR pads:ti,ab,kw OR 'fecal incontinence':ti,ab,kw OR 'faecal incontinence':ti,ab,kw OR 'bowel incontinence':ti,ab,kw OR 'fecal soiling':ti,ab,kw

		OR 'faecal soiling':ti,ab,kw OR 'EQ-5D':ti,ab,kw OR 'EuroQol':ti,ab,kw OR 'EuroQol-5D':ti,ab,kw OR intercourse:ti,ab,kw OR 'sexual behavior':ti,ab,kw OR 'sexual activity':ti,ab,kw OR 'Wexner Scale of incontinence':ti,ab,kw OR 'Revised fecal incontinence scale':ti,ab,kw OR 'Revised faecal incontinence scale':ti,ab,kw OR 'RFIS':ti,ab,kw OR 'anal pain':ti,ab,kw OR pain:ti,ab,kw OR 'leakage':ti,ab,kw
<b>Combination Searches</b>		
#11	Condition 2 + Crohn's	#2 AND #3
#12	Combined Queries	#6 OR #7 OR #8 OR #9 OR #10
<b>SEARCH RESULTS</b>		
#13	Condition 2 + Combined Queries + RWD + filters	#11 AND #12 AND #5 +English, human, 10 years filters
#14	Condition 1 + Combined Queries + RWD + filters	#1 AND #12 AND #5 +English, human, 10 years filters
#15	Final search including fistula title search	(#13 OR #14) AND #4