

Foreign Material in Postoperative Adhesions

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Objective

The authors determined the prevalence of foreign body granulomas in intra-abdominal adhesions in patients with a history of abdominal surgery.

Patients and Methods

In a cross-sectional, multicenter, multinational study, adult patients with a history of one or more previous abdominal operations and scheduled for laparotomy between 1991 and 1993 were examined during surgery. Patients in whom adhesions were present were selected for study. Quantity, distribution, and quality of adhesions were scored, and adhesion samples were taken for histologic examination.

Results

In 448 studied patients, the adhesions were most frequently attached to the omentum (68%) and the small bowel (67%). The amount of adhesions was significantly smaller in patients with a history of only one minor operation or one major operation, compared with those with multiple laparotomies ($p < 0.001$). Significantly more adhesions were found in patients with a history of adhesions at previous laparotomy ($p < 0.001$), with presence of abdominal abscess, hematoma, and intestinal leakage as complications after former surgery ($p = 0.01$, $p = 0.002$, and $p < 0.001$, respectively), and with a history of an unoperated inflammatory process ($p = 0.04$).

Granulomas were found in 26% of all patients. Suture granulomas were found in 25% of the patients. Starch granulomas were present in 5% of the operated patients whose surgeons wore starch-containing gloves. When suture granulomas were present, the median interval between the present and the most recent previous laparotomy was 13 months. When suture granulomas were absent, this interval was significantly longer—*i.e.*, 30 months ($p = 0.002$). The percentage of patients with suture granulomas decreased gradually from 37% if the previous laparotomy had occurred up to 6 months before the present operation, to 18% if the previous laparotomy had occurred more than 2 years ago ($p < 0.001$).

Conclusions

The number of adhesions found at laparotomy was significantly larger in patients with a history of multiple laparotomies, unoperated intra-abdominal inflammatory disease, and previous postoperative intra-abdominal complications, and when adhesions were already present at previous laparotomy. In recent adhesions, suture granulomas occurred in a large percentage. This suggests that the intra-abdominal presence of foreign material is an important cause of adhesion formation. Therefore intra-abdominal contamination with foreign material should be minimized.

Intra-abdominal adhesions are strands or membranes of fibrous tissue that can be attached to the various intra-abdominal organs, gluing them strongly together. It has been estimated that one third of intestinal obstructions¹⁻⁵ and 15% to 20% of female infertility is caused by adhesions.^{6,7} The amount of effort general surgeons put into work because of adhesions is large. Approximately 1% of all surgical admissions and 3% of laparotomies are the result of intestinal obstruction from adhesions.² The treatment of patients with symptoms caused by adhesions also will generate extra costs.

Adhesions are congenital or develop after an abdominal operation or infection.^{2,3} Intraoperative tissue damage, infections, tissue ischemia, and intra-abdominal presence of foreign material, blood, or bile,^{1,8-15} all have been shown to be potent causes of peritoneal adhesions. Foreign materials, such as glove powder,¹¹ fluff from surgical packs (gauze lint),¹² sutures,^{13,14} and material extruded from the digestive tract, cause a peritoneal inflammatory reaction.¹⁵ This reaction potentiates adhesion formation, especially with concomitant peritoneal damage, as has been demonstrated in various animal models.^{1,11,13,15-18} Such adhesions often contain multiple foreign body granulomas.^{10,16,17,19-23} This strongly suggests a relation between foreign material, foreign body granulomas, and adhesion formation. Foreign bodies contaminating the peritoneal tissues also might be a cause of adhesion formation in humans. Reported studies are small or date from the time when the contemporary advanced hygienic surgical techniques were not in use.²⁴⁻²⁶ In the current series, the presence of foreign body granulomas and adhesions after laparotomy was studied in humans.

PATIENTS AND METHODS

Adult patients with a history of previous abdominal surgery and scheduled for laparotomy between 1991 and 1993 at the University Hospital Rotterdam (Rotterdam, The Netherlands = NL), Groupe Hospitalier Pitié-Salpêtrière (Paris, France = F1), Hôpital d'Instruction des Armées du Val-de-Grâce (Paris, France = F2), Hospital de la Esperanza (Barcelona, Spain = SP), and the University of Göteborg Östra Hospital (Göteborg, Sweden = SW), were examined during surgery. Patients were selected for study if adhesions appeared to be present. During the operation, the quantity, distribution, and quality of adhesions were scored, and adhesion samples were

taken for histologic examination. Quantity was grouped as 1 to 3, 4 to 10, and more than 10 adhesions.²³ To score distribution, adhesion adherence to the following structures was determined: laparotomy scar, omentum, abdominal wall, liver, spleen, stomach, small intestine, colon, retroperitoneum, and in female reproductive organs. Whenever structures were absent or not explored, this was noted. In addition, adhesions were scored using the *macroscopic* classification according to Zühlke (Table 1).²⁷ One to three samples of adhesions were taken. These samples were fixed in 4% formaldehyde, put in paraffin, sectioned (six micron), and stained with hematoxylin and eosin (H&E) and periodic acid Schiff's reagents. Then, the adhesions were scored microscopically, using the *histologic* classification according to Zühlke (Table 2),²⁷ and the slides were screened for granulomas and foreign material in or near histiocytes or giant cells, using both normal and polarized light. Using polarized light, glove starch powder can be recognized by the typical birefringent Maltese crosses.

Patient-related factors of gender, age, history of previous abdominal surgery, radiotherapy, and prior inflammatory disease and operation-related factors including the presence of adhesions at prior surgery, postoperative complications (abscess, hemorrhage, anastomotic bowel leakage), use of prosthetic material, and use of powdered or unpowdered gloves were analyzed. An appendectomy, hysterectomy, incisional hernia repair, or diagnostic laparotomy was scored as minor operation. All other abdominal operations were scored as major operations.

Statistical methods used included the Kruskal-Wallis test or the Mann-Whitney test for the comparison of ordered classifications and time intervals. Percentages were compared using the chi square test. Logistic regression²⁸ was used to evaluate various factors simultaneously with regard to the percentage of patients who had more than ten adhesions. This technique also was used to assess the relation between the percentage of patients with granulomas and the (logarithmically transformed) time interval from the previous operation. P values given are two-sided; 0.05 was considered the limit of significance.

Table 1. MACROSCOPIC CLASSIFICATION ACCORDING TO ZÜHLKE²⁷

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- | | |
|-----|-------------------------------------------------------------------------------------------------------------------------------|
| I | Filmy and easy to separate by blunt dissection |
| II | Blunt dissection possible, partly sharp dissection necessary, beginning vascularization |
| III | Lysis possible by sharp dissection only, clear vascularization |
| IV | Lysis possible by sharp dissection only, organs strongly attached with severe adhesions, damage of organs hardly preventable. |
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RESULTS

Quantity of Adhesions

A total of 448 patients with adhesions was included (222 NL, 139 F1, 14 F2, 44 SP, 29 SW). There were 202 men and 246 women; 27% had 1 to 3 intra-abdominal adhesions, 30% had 4 to 10 intra-abdominal adhesions, and 43% had more than 10 intra-abdominal adhesions. A history of one minor, one major, or multiple operations was present in 24%, 33%, and 43% of the patients, respectively.

The number of adhesions was significantly lower in patients with a history of only one minor or one major operation, compared with those with multiple laparotomies ($p < 0.001$). No significant difference was found between one minor and one major operation ($p = 0.15$, Table 3).

The number of adhesions was significantly larger in patients with adhesions at previous laparotomy (Table 4) compared with those without ($p < 0.001$). Also, significantly more adhesions were found in patients with a history of a surgical complication, such as abdominal abscess ($p = 0.01$), hematoma ($p = 0.002$) and intestinal leakage ($p < 0.001$), compared with patients who had an uncomplicated course. Significantly more adhesions also were found in patients with a history of an unoperated inflammatory process compared with patients without ($p = 0.04$). No significant difference in amount of adhesions was found between patients who had prosthetic material intra-abdominally or in those who had received abdominal radiotherapy.

Using multivariate analysis, it was found that, besides the type of previous operation and the presence of adhesions at previous operation, intestinal leakage was the most important factor regarding the presence of more than ten adhesions. None of the other factors that were significantly related to the amount of adhesions when considered alone (abdominal abscess, hematoma and previously unoperated inflammatory process) appeared of importance when taking into account the presence of adhesions and intestinal leakage at previous operations and type of previous operation.

Table 2. HISTOLOGIC CLASSIFICATION ACCORDING TO ZÜHLKE²⁷

I	Loose connective tissue, cell-rich, old and new fibrin, fine reticulin fibers
II	Connective tissue with cells and capillaries, few collagen fibers
III	Connective tissue more firm, fewer cells, more vessels, few elastic and smooth muscle fibers
IV	Old firm granulation tissue, cell-poor, serosal layers hardly distinguishable.

Table 3. THE NUMBER OF ADHESIONS IN PATIENTS (%) CLASSIFIED BY THE HISTORY OF OPERATIONS

	Previous Operation(s)		
	One Minor (A)	One Major (B)	Multiple (C)
No. of adhesions			
1-3	43 (41)	51 (35)	22 (12)
4-10	37 (36)	46 (32)	48 (26)
>10	24 (23)	47 (33)	118 (63)
Total	104 (100)	144 (100)	188 (100)

A vs. B: $p = 0.15$.
A vs. C: $p < 0.001$.
B vs. C: $p < 0.001$.

Distribution

In calculating the percentages of patients with adhesions per organ, patients were excluded if the organ was not visualized during surgery. The distribution was as follows: omentum, 293 of 430 patients (68%); small bowel, 288 of 428 patients (67%); abdominal wall, 193 of 432 patients (45%); female reproductive organs, 49 of 212 patients (23%); colon, 205 of 419 patients (41%); liver, 118 of 352 patients (34%); stomach, 67 of 342 (20%); retroperitoneum, 56 of 388 patients (14%); and spleen, 28 of 309 patients (9%). The laparotomy scar was attached to one or more of the following organs in 314 out of 440 cases (71%): omentum, small bowel, colon, or abdominal wall. The site of previous surgery was attached to one or more of these organs in 229 of 434 cases (53%). Remote adhesions (to structures not previously operated on) were adherent to one or more of these organs in 156 of 436 cases (35%).

Nature

Of 24 patients with adhesions, either no biopsy material or unsuitable biopsy material was taken, leaving 424 patients for microscopic examination. The percentages of adhesions scored in the macroscopic ($n = 429$) and histologic ($n = 415$) Zühlke classification I, II, III, and IV were, respectively, 11%, 30%, 49%, and 10% and 7%, 49%, 37%, and 7%. No relationship was found between 1) both classifications and 2) the time interval to the most recent operation and presence of granulomas.

Granulomas

Granulomas were found in 112 of 424 patients (26%). Suture granulomas (Fig. 1) were found in 105 of 424 pa-

Table 4. THE INFLUENCE OF VARIOUS FACTORS ON THE NUMBER OF ADHESIONS*†

Factor	No. of Adhesions			Value
	1-4	4-10	>10	
Adhesions at previous operation				
Yes (n = 126)	12 (9)	26 (21)	88 (70)	<0.001
No (n = 119)	32 (27)	40 (34)	47 (39)	
Postoperative complications				
Abdominal abscess				
Yes (n = 44)	8 (19)	8 (19)	28 (66)	0.01
No (n = 390)	108 (28)	123 (31)	159 (41)	
Hematoma				
Yes (n = 15)	0 (0)	3 (20)	12 (80)	0.002
No (n = 419)	116 (28)	128 (30)	175 (42)	
Intestinal leakage				
Yes (n = 42)	5 (12)	3 (7)	34 (81)	<0.001
No (n = 392)	111 (28)	128 (33)	153 (39)	
Other risk factors				
Prosthetic material used intra-abdominally				
Yes (n = 22)	4 (18)	6 (27)	12 (55)	0.25
No (n = 302)	111 (27)	125 (30)	176 (43)	NS
Abdominal radiotherapy				
Yes (n = 18)	2 (11)	6 (33)	10 (56)	0.16
No (n = 412)	113 (27)	121 (29)	178 (43)	NS
Previously unoperated inflammatory process				
Yes (n = 64)	12 (19)	15 (23)	37 (58)	0.04
No (n = 353)	100 (28)	109 (31)	144 (41)	
Gloves used in previous operations				
Contained starch (n = 319)	74 (23)	98 (31)	147 (46)	0.13
Starch free (n = 45)	16 (35)	12 (27)	17 (38)	NS

NS = not significant.

* Data given are numbers of patients (%).

† Totals do not add up to total amount of patients because of occasionally missing data.

tients (25%). Starch granulomas (Fig. 1) were found in 14 of 309 patients (5%) with a history of being operated on with starch-containing gloves.

In those patients with suture granulomas (n = 105), the median interval between the present and previous operation was 13 months. This interval was significantly longer (p = 0.002) for patients with no suture granulomas (median 30 months). Table 5 shows the percentage of patients with suture granulomas according to the interval from the previous operation. The percentage decreased gradually from 37% if the previous operation had occurred up to 6 months before the present operation, to 18% if the previous operation had occurred more than 2 years ago. Figure 2 shows the percentages according to the interval on a continuous basis.

In patients with starch granulomas (n = 14), the median interval between the present and previous operation was 14 months. This interval showed a trend to be longer (p = 0.09) for patients with no suture granulomas with a median of 42 months. The percentage of patients with

starch granulomas according to the interval from previous operation also is shown in Table 5.

DISCUSSION

More than 25 years ago, Myllärniemi reported that 61% of 309 patients with postoperative adhesions showed reactions to foreign material. Substances identified included talc (31%), gauze lint (16%), a combination of talc and gauze lint (11%), starch (1%), and sutures (< 1%). In contrast, in the current series 25% of 424 patients had suture granulomas, and 5% of 309 patients being operated on with starch containing gloves had starch granulomas. This difference might be explained partially by the stringent criteria used for identification of foreign body granulomas, which reduces the chance of contamination being erroneously scored as such. Further, talc glove powder no longer is in use, and has been replaced by starch. Another explanation might be a different dis-

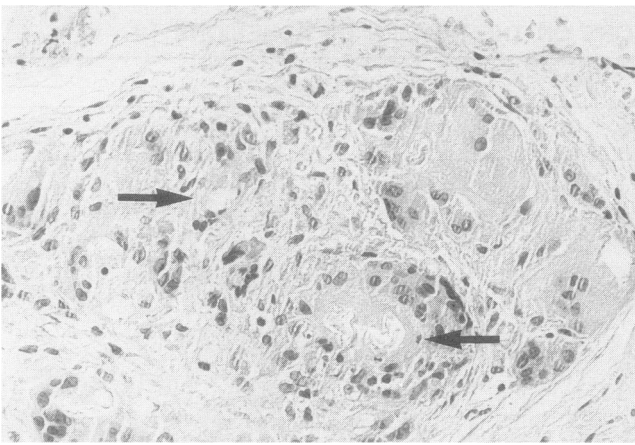


Figure 1. Adhesion with granulomas; left arrow: starch granuloma with typical Maltese cross; right arrow: suture granuloma. Birefringent and H&E, magnification 400 X.

tribution of patients with respect to the interval between the present and the most recent previous operation. Foreign body granulomas are found more often in patients recently operated on. In the current study, this was demonstrated for suture granulomas, but not for starch granulomas. An explanation for this difference might be the ability of the body to resorb starch and suture materials. The largest extent of this resorption of suture materials seems to take place during the first year.

The number of granulomas found at laparotomy might be even larger than accounted for in the current study. First, in patients with a history of multiple laparotomies, the adhesion removed for microscopic exami-

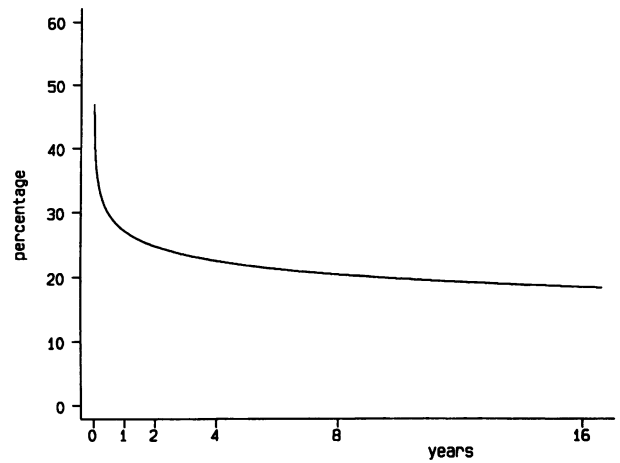


Figure 2. Percentage of patients with suture granulomas with respect to the interval between present and the most recent operation.

nation might have formed before the most recent operation. This reduces the chance of finding granulomas because of resorption. Second, granulomas might be present in the adhesion but not in the sample or histologic slide. Third, resorption of foreign material might have been completed, leaving an empty granuloma or, when the granuloma itself also has disappeared with time, an adhesion without granulomas.¹⁵ This resorption of foreign material, however, comes too late to prevent adhesion formation. The organization of fibrinous adhesions starts at 3 days.²⁹

The resorption rate strongly influences the chance of finding foreign material. The resorption rate of starch powder is not known, but will depend on the glove powder used (kind, composition, amount, clumping) and the host (intra-abdominal conditions, individual inflammatory reaction/sensitivity).²³ Experiments on animals suggest that powder can be resorbed within 24 hours, leaving granulomas and firm adhesions long thereafter.¹⁶ At the latest, starch granulomas still were present at 15 months in experimental studies¹¹ and at 23 months in clinical studies.²⁴ In the current study, starch granulomas were found mostly up to 48 months. However, in four patients, starch granulomas were found much later—up to 32 years after the last operation. We do not have an explanation for this finding.

The question remains: why were suture granulomas found so often in this study (105/424; 25%), whereas in other studies, almost no suture granulomas were found? Specimens were taken at random, and the presence of a granuloma or suture seldomly was suspected macroscopically. Remarkably, in 39 of 211 patients (18%) who had the last operation more than 2 years previously *versus* 66 of 213 (31%) who had had their last operation less than 2 years previously, suture granulomas were present.

Table 5. DISTRIBUTION OF FOREIGN BODY GRANULOMAS WITH RESPECT TO THE INTERVAL BETWEEN CURRENT AND THE MOST RECENT OPERATION*

Type of Granuloma	Interval (mos)				Totals
	0-6	>6-12	>12-24	>24	
Suture granulomas†					
Yes	33 (37)	17 (30)	16 (26)	39 (18)	105 (100)
No	56 (63)	46 (70)	45 (74)	172 (82)	319 (100)
Starch granulomas‡					
Yes	5 (10)	2 (5)	1 (2)	6 (3)	14 (100)
No	47 (90)	39 (95)	42 (98)	167 (97)	295 (100)

* Data given are numbers of patients (%).

† p < 0.001.

‡ p = 0.09.

Therefore, the large amount of suture granulomas cannot be explained exclusively by use of nonabsorbable suture material. Unfortunately, we were not able to differentiate between absorbable or nonabsorbable suture in our material.

Apart from the aforementioned, the presence of suture material and the ischemia caused by tightening the suture can potentiate adhesion formation.^{23,30} This may explain why the laparotomy scar so often is attached to its surroundings by adhesions.

As a consequence to these results, the use of suture material should be minimized to avoid the intra-abdominal presence, as well as the ischemia it causes. Closure of the peritoneum, for instance, is unnecessary^{1,29} and in the aforementioned context, unwanted. A meticulous technique can limit foreign body contamination and subsequent granuloma formation and peritoneal damage. This may reduce adhesion formation.⁵

Powder-free gloves prevent starch granulomas, and thus are recommended. Powdered gloves can be made powder free by a 1-minute washing with 10 mL of povidone/iodine, followed by a 30-second rinse with sterile water.³¹ However, this method is time consuming and costly, and deviating from this procedure can lead to clumping of starch granules, which may give rise to a more intense tissue reaction.¹⁷

CONCLUSION

The amount of adhesions found at laparotomy is significantly larger in patients with a history of multiple laparotomies, with adhesions already present at previous laparotomy, with previous intra-abdominal complications (abdominal abscess, hematoma and intestinal leakage), and with unoperated intra-abdominal inflammatory disease.

Microscopic examination of adhesions showed granulomas in a large percentage (26%), with suture granulomas in 25% and starch granulomas in 5%. Suture granulomas were found significantly more often in patients recently undergoing surgery. Therefore, modern surgical techniques are not as meticulous as we might think. If the causative relation between foreign material, foreign body granulomas, and adhesion formation shown in animals also exists in humans, the operative contamination with foreign material is an important cause of adhesion formation. In view of the serious consequences, such as intestinal obstruction and infertility, prevention of contamination during surgery with foreign material is of considerable importance.

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