DERMABOND ADVANCED®
Topical Skin Adhesive
Evidence Brief
Overview

As the final layer of wound closure, topical skin adhesives (TSAs) are an integral part of a successful clinical outcome. When deciding which TSA to use, clinical study information on closure strength, microbial protection, patient comfort, and cosmesis allows healthcare practitioners to evaluate which product will provide the greatest benefits for their patients.

DERMABOND ADVANCED® Topical Skin Adhesive is supported by an extensive body of published literature, including 53 randomized controlled trials (RCTs). DERMABOND ADVANCED Adhesive has a patented, proprietary chemical formulation\(^1\) that has been shown to provide superior strength versus other commercially available TSAs,\(^2\) and also has benefits that enhance patient comfort and cosmetic outcomes.\(^3,6\)

This Evidence Summary includes a sample of the available RCTs for DERMABOND ADVANCED Adhesive or DERMABOND® Topical Skin Adhesive. A full list of published studies can be found in the bibliography section of this document.

- DERMABOND ADVANCED Adhesive and DERMABOND Adhesive are supported by 53 published RCTs\(^*\)\(^†\)
- Total of 5,836 patients evaluated

References

1. DERMABOND ADVANCED® Topical Skin Adhesive Label. LAB-0012182 DNX12. Ethicon, Inc.

*DERMABOND ADVANCED Adhesive tests equivalent or superior to DERMABOND Adhesive in head-to-head testing for microbial barrier, wound-bursting strength, tensile strength, flexibility, durability, viscosity, drying time, water vapor transmission rate, water resistance, and physician satisfaction.

†Based on published literature in PubMed and SCOPUS, using only RCTs that evaluated the use of the product in a manner consistent with intended indication.
Summary of Key Studies

The publications that support the claims for DERMABOND ADVANCED® Topical Skin Adhesive are listed in the table below. A summary of each of these studies can be found on the subsequent pages.

<table>
<thead>
<tr>
<th>Publication Title</th>
<th>Lead Author</th>
<th>Source</th>
<th>Outcome Studied</th>
</tr>
</thead>
<tbody>
<tr>
<td>In vitro Assessment of Microbial Barrier Properties of DERMABOND® Topical Skin Adhesive</td>
<td>Bhende</td>
<td>Surgical Infections. 2002;3(3):251-257.</td>
<td>Microbial Barrier</td>
</tr>
<tr>
<td>In vitro study to determine the ability of DERMABOND ADVANCED® Topical Skin Adhesive to inhibit bacterial growth</td>
<td>Bhende</td>
<td>Internal Ethicon Study</td>
<td>Inhibition of Bacteria</td>
</tr>
</tbody>
</table>
Clinical Reference Article Summary

In Vitro Assessment of Microbial Barrier Properties of DERMABOND® Topical Skin Adhesive

Bhende S, Rothenburger S, Spangler D, Dito M

Source:
Surgical Infections. 2002;3(3):251-257

Study Objective
The purpose of this study was to evaluate the ability of DERMABOND Adhesive to provide an effective microbial barrier against the penetration of microorganisms in vitro.

Bacteria used in this study included:
Staphylococcus aureus
Staphylococcus epidermidis
Escherichia coli
Pseudomonas aeruginosa
Enterococcus faecium

Methods
Plates containing an agar media were created in a sterile environment. The agar media contained a pH-sensitive dye designed to color when exposed to the acidic metabolic products of bacteria.

DERMABOND Adhesive was applied to the agar surface. In total, 300 single-layer films and 300 triple-layer films were examined. The surface of each film was inoculated with a 10^6 L aliquot of bacteria containing at least 1x10^3 colony-forming units (cfu).

All test and control plates were incubated at 37°C for 72 hours. A change in color indicated a breach in the adhesive's microbial barrier.

Results
Single-layer films: 299 of the 300 samples retained their integrity as microbial barriers for 72 hours. All 300 samples maintained their microbial barrier for 48 hours.

For the triple-layer films, 299 of the 300 samples retained their integrity as microbial barriers for 72 hours.

Conclusion
The results of this study demonstrate that DERMABOND Adhesive provides a microbial barrier with 99% protection in vitro for at least 72 hours against organisms commonly responsible for SSIs, including: Staphylococcus epidermidis, Staphylococcus aureus, Escherichia coli, Pseudomonas aeruginosa, and Enterococcus faecium.
Clinical Reference Article Summary

In vitro study to determine the ability of DERMABOND ADVANCED® Topical Skin Adhesive to inhibit bacterial growth

Bhende S

Source:
Internal Ethicon Study

Study Objective
The purpose of this study was to demonstrate that DERMABOND ADVANCED Adhesive inhibits gram-positive bacteria and gram-negative bacteria in vitro.

Bacteria evaluated in this study:
Methicillin-resistant Staphylococcus aureus (MRSA)
Methicillin-resistant Staphylococcus epidermidis (MRSE)
Escherichia coli

Methods
Cultures of each organism were grown under sterile conditions for 18-24 hours at 35-37°C. Before being used in the experiment, each culture was diluted to achieve an approximate bacteria count of 10^5 colony-forming units (cfu)/0.04 ml.

A 2 cm diameter circle was drawn on the bottom of a sterile agar plate. In the center of this circle, 0.04 ml of the diluted inoculum was placed on the surface of the agar.

After allowing the inoculum to dry, the adhesive material was applied to the inoculated surface area, making sure to cover the area beyond the marked circle.

After 10 minutes of contact time between the adhesive and the inoculated area, the adhesive's polymerized film was removed from the surface of the agar, and the plates were incubated at 37°C for up to 48 hours.

In total, 210 samples (70 samples per organism) were evaluated. The samples were examined for bacterial growth at 24 and 48 hours. Any growth originating beneath the area of adhesive application was recorded as a positive test.

Results
After 48 hours, the test plates exhibited colony counts ranging from 0 - 59 cfu, indicating significant inhibition of the bacteria.

Each inoculated plate was declared a success if a minimum of 99.9% inhibition of the initial inoculum load was observed. For all bacteria evaluated (MRSA, MRSE, E. coli), contact with the adhesive led to a 99.9% inhibition in bacteria load from the initial inoculum.

Conclusion
In this in vitro study, DERMABOND ADVANCED Adhesive was shown to demonstrate inhibition of gram-positive bacteria (MRSA, MRSE) and gram-negative bacteria (E. coli).*

*Clinical significance is unknown.
Clinical Reference Article Summary
In Vivo Study of Wound Bursting Strength and Compliance of Topical Skin Adhesives
Singer AJ, Perry LC, Allen Jr. RL

Source:

Study Objective
The purpose of this study was to evaluate the wound-bursting strength and flexibility of five topical skin adhesives during the two-day period after wound closure.

The following adhesives were evaluated in the study:
DERMABOND® Topical Skin Adhesive
INDERMIL® Tissue Adhesive
Histoacryl® Topical Skin Adhesive
LiquiBand® Topical Skin Adhesive
GluStitch®

Methods
Using a template for incision length and location, two symmetric incisions (2 cm long each) were created over the dorsolateral flank area of 210 anesthetized, male Sprague-Dawley rats.

After achieving hemostasis and manually approximating the skin edges, a randomized computer algorithm was used to select an adhesive to close the incision. All adhesives were applied according to manufacturer instructions.

The adhesives were evaluated three times during the study—immediately after closure, 1 day after closure, and 2 days after closure.

For each evaluation, 14 samples from each adhesive group were tested for wound-bursting strength, and another 14 samples were tested for flexibility.

To test for wound-bursting strength, a vacuum chamber was placed over each sample and negative pressure was applied, stressing the wound in 3 dimensions. The pressure (mmHg) needed to cause wound failure was recorded.

To test for flexibility, a vacuum chamber was placed over the sample and negative pressure was applied to the wound while a laser measured the vertical deformation of the skin (μm). Energy absorption (mmHg x mm) was calculated to quantify the adhesives’ flexibility.
Results

In total, 210 measurements were taken on 210 incisions (5 adhesives, 3 time points, 14 samples per time point). Results are shown in Figure 1.

With the exception of the samples in the DERMABOND® Topical Skin Adhesive group, measurements could not be taken on all samples in an adhesive group because, in some samples, the adhesive’s inflexibility had caused the adhesive to fracture during testing.

As shown in Figure 2, the percent of samples in an adhesive group experiencing fractures ranged from 36% to 86%.

Conclusion

The results of this study demonstrate that DERMABOND Adhesive was significantly stronger and more flexible than the other adhesives evaluated in the study.*

*This study was funded in full or in part by an educational grant from Ethicon, Inc.
Clinical Reference Article Summary

Postoperative Outcomes Associated with Topical Skin Adhesives among Women Having Hysterectomies
Murrmann SG, Markowitz JS, Gutterman EM, Magee G

Source:
Surgical Infections. 2010;11(5):441-447

Study Objective
The purpose of this study was to evaluate the clinical and economic outcomes associated with use of a topical skin adhesive (TSA) versus traditional methods for skin closure following total abdominal hysterectomy.

Methods
The study utilized Premier Perspective™ Comparative Database, which is a large, administrative database containing clinical and economic data from all patient discharge records at more than 400 US hospitals.

Any patient in the database who was discharged from a hospital in 2005 following a total abdominal hysterectomy was included in the study.

The subjects were classified into one of four treatment groups based on the clinical method used to close the surgical incision:

- Sutures
- Staples
- TSA
- Staples and TSA

While the study was open to all commercially available TSAs, at the time of the study the only TSA used on patients in the database was DERMABOND® Topical Skin Adhesive. Thus, the TSA group only had patients treated with DERMABOND Adhesive.

All treatment groups were assessed on three continuous outcomes: length of inpatient stay, total inpatient cost, and days of antibiotic treatment. Length of stay and inpatient cost was available directly from the database; antibiotic treatment days were estimated using the last date when at dose of antibiotic was administered.

Results
In total, 46,011 patients were included in the study. The method of wound closure for these patients is summarized in Figure 1.

Due to the large sample size, there were no statistically significant differences in the clinical, demographic, or hospital characteristics of the four treatment groups.

<table>
<thead>
<tr>
<th>Skin Closure Method Evaluated in Study</th>
<th># of Patients (n)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sutures</td>
<td>21,201</td>
</tr>
<tr>
<td>Staples</td>
<td>23,441</td>
</tr>
<tr>
<td>TSA</td>
<td>880</td>
</tr>
<tr>
<td>Staples and TSA</td>
<td>489</td>
</tr>
<tr>
<td>All Methods</td>
<td>46,011</td>
</tr>
</tbody>
</table>
Length of Stay (LOS) and Total Costs

A summary of mean LOS and total hospitalization costs is shown in Figure 2.

While the difference in total costs between suture and TSA groups did not meet the significance requirement for this study ($P \leq 0.01$), the difference suggests lower total costs for the TSA group ($P = 0.039$).

<table>
<thead>
<tr>
<th>Skin Closure Method Evaluated in Study</th>
<th>Mean LOS (days)</th>
<th>Mean Total Hospitalization Costs</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sutures</td>
<td>3.9</td>
<td>$5,862</td>
</tr>
<tr>
<td>Staples</td>
<td>4.5</td>
<td>$6,965</td>
</tr>
<tr>
<td>TSA</td>
<td>3.7</td>
<td>$5,816</td>
</tr>
<tr>
<td>Staples and TSA</td>
<td>5.2</td>
<td>$9,434</td>
</tr>
</tbody>
</table>

Conclusions

The results of this study demonstrate that the clinical and economic outcomes were consistently worse when staples were used to close an incision compared with use of suture or TSA alone.

The clinical outcomes resulting from the use of DERMABOND® Topical Skin Adhesive to close wounds were at least as good as the outcomes resulting from the use of suture to close wounds.

Additionally, there is evidence that the total costs of hospitalization for total hysterectomy patients may be less when the incision is closed with DERMABOND Adhesive versus sutures or staples.*

*The study was funded in full or in part by an educational grant from Ethicon, Inc.
Clinical Reference Article Summary

A Randomized Trial Comparing Octylcyanoacrylate Tissue Adhesive and Sutures In the Management of Lacerations

Quinn J, Wells G, Sutcliffe T, Jarmuske M, Maw J, Stiell I, Johns P

Source:
JAMA. 1997;277(19):1527-1530

Study Objective
The purpose of this study was to assess whether using DERMABOND® Topical Skin Adhesive for laceration repair is an effective alternative to suturing.

Methods
Patients with non-mucosal facial lacerations as well as certain extremity and torso lacerations, but not on hands, feet or joints, were eligible for this study.

Using a computer algorithm, patients were prospectively segregated into facial and non-facial groups and randomized into two groups—DERMABOND Adhesive and sutures.

In the suture group, lacerations were anesthetized and cleaned, as needed, before repair with a 5-0 or 6-0 monofilament suture. A dressing was applied for at least 48 hours.

In the DERMABOND Adhesive group, lacerations were cleaned with chlorhexidene and hemostasis was achieved using pressure or topical 1:1000 epinephrine. The wound edges were manually approximated and the adhesive was applied to the surface of the skin, covering the wound edges. The wound was held in place for 30 seconds. No dressing was applied.

The primary outcome was the cosmetic appearance of the scar, evaluated by a blinded plastic surgeon using a photograph of the wound taken 3 months after closure.

On two occasions, the surgeon examined the photograph and provided a cosmesis score based on a validated 100-mm visual analog scale, ranging from “best scar” to “worst scar.”

Additionally, time of procedure, patient pain, and wound complications (i.e., dehiscence, infection) were recorded. Time of procedure was evaluated from start of wound care to complete closure; patient pain and wound complications were recorded on a previously validated scale.

Wound complication was initially evaluated at 3-5 days for facial and at 10-14 days for torso and extremity lacerations. A second assessment occurred 3 months after closure.

Results
In total, 130 patients with 136 lacerations were included in the study. As summarized in Figure 1, an equal number of lacerations (68 per group) were randomized to the suture and DERMABOND Adhesive groups.

<table>
<thead>
<tr>
<th>Figure 1</th>
<th>Patient Retention During Study</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td><strong>DERMABOND Adhesive</strong></td>
</tr>
<tr>
<td>Randomized</td>
<td>68</td>
</tr>
<tr>
<td>Initial follow-up</td>
<td>53</td>
</tr>
<tr>
<td>3 month follow-up</td>
<td>55</td>
</tr>
<tr>
<td>Withdrawn</td>
<td>1</td>
</tr>
<tr>
<td>Lost to follow-up</td>
<td>12</td>
</tr>
<tr>
<td>No Photographs</td>
<td>5</td>
</tr>
<tr>
<td>Completed Study</td>
<td>50</td>
</tr>
</tbody>
</table>
As shown in Figure 2, there was no significant difference in the blinded, 3-month cosmetic score of the DERMABOND® Topical Skin Adhesive group compared with the suture group. Similarly, there was no significant difference in wound complications between the suture group and the DERMABOND Adhesive group. Statistically significant differences were seen for patient pain and procedure time.

Conclusions
The results of this study demonstrate that DERMABOND Adhesive produces cosmetic results similar to suturing on certain types of lacerations.

Additionally, lacerations closed with DERMABOND Adhesive were associated with shorter procedure time and less patient pain than lacerations closed with sutures.*

*This study was funded in full or in part by an educational grant from Ethicon, Inc.

**Figure 2**
Summary of Observed Clinical Outcomes

<table>
<thead>
<tr>
<th></th>
<th>DERMABOND Adhesive</th>
<th>Suture</th>
<th>(P) Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean Cosmetic Score (mm)</td>
<td>67</td>
<td>68</td>
<td>0.65</td>
</tr>
<tr>
<td>% Optimal Wound Scores (initial eval)</td>
<td>80%</td>
<td>82%</td>
<td>0.80</td>
</tr>
<tr>
<td>% Optimal Wound Scores (3 month eval)</td>
<td>72%</td>
<td>75%</td>
<td>0.74</td>
</tr>
<tr>
<td>Mean Pain Scores (mm)</td>
<td>7.2</td>
<td>18.0</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Mean Time of Procedure (min)</td>
<td>3.6</td>
<td>12.4</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>
For More Information

Call 1-877-ETHICON (384-4266)

In addition to support from Ethicon Sales Representatives, Ethicon’s Medical Affairs team is available to provide balanced, non-promotional scientific information to healthcare professionals.

Medical information request form

To: Ethicon Medical Affairs

E-mail: Eth_Medical_Info@its.jnj.com    Voicemail: (800) 888-9234, x3800

Date: ___________________________________________________________________________________________________________________________________

From (Requestor): ___________________________________________________________________________________________________________________

Name: _________________________________________________________________________________________________________________________________

(Circle one):


Other:___________________________

Title: ___________________________________________Institution/Office: _________________________________________________________

Address: _______________________________________________________________________________________________________________________________

City: _________________________________________________________________________________ State: ______________ZIP: ________________________

Telephone:  ____________________________________________________________ Fax: _________________________________________________________

E-mail Address: _______________________________________________________________________________________________________________________

Desired Response Method (Circle one):

US Mail     Phone       E-mail      Fax  Meeting with Medical Affairs Representative

Requestor’s Signature: ____________________________

(REQUIRED FOR PROCESSING)___________________________________________________________________________________________________

Please send medical information on the following topic(s):

(Be as specific as possible with respect to product topic, area of use, outcome of interest, etc.)

________________________________________________________________________________________________________________________________________

________________________________________________________________________________________________________________________________________

________________________________________________________________________________________________________________________________________

Sales Representative: ____________________________ Territory:_____________________

PRINT FULL NAME __________________________________________________________________________________________________________________
Bibliography

Listed below are all of the currently published RCTs that have evaluated the use of DERMABOND® Topical Skin Adhesive in an application consistent with the indication in the product's label (i.e., skin closure). Studies that evaluated the use of DERMABOND Adhesive for purposes inconsistent with the intended indication were excluded from the bibliography.


