

Climate neutral print production

♣ Forest Stewardship Council (FSC")

◆ Good Manufacturing Practice guidelines (GMP)

Hygiene Directives (HACCP)

♣ DIN EN I2O 8001 Responsibility How to use the 3in1 Outsert

Use the 3in1 Outsert exactly as described.

Driving and using machines

Starting dose: read the 3in1 Outsert before using the medicine

Method of use

Use the 3in1 Outsert after opening the seal labels and unfolding the relevant leaflet.

Duration of use

Using too much 3in1 Outsert

Forgotten use

Use it immediately upon remembering.

Stop using

meter 60 gem + 450 × 596 mm ber product

🗾 **Bähren** Druck

Je banel per product

♣ 2 seal labels



Mittelstraße 62, 41236 Mönchengladbach, Germany, +49 2166 97291 0, info⊚baehren-druck.de

Bähren Druck - The Pharma Specialist



9102.20/20



What 3in1 Outsert is used for

The 3in1 Outsert is used for:

 three pieces of medical information, such as: » package leaflet, instructions for use, summary of product characteristics

» package leaflet, instructions for use, additional patient information » package leaflet, summary of product characteristics, additional patient information

trilingual medical information, such as package leaflets for

» distribution in small markets, such as the 3 Baltic countries or Finland, Norway and Sweden » cost saving distribution to summarise markets like Austria, Belgium, France, Germany, Liechtenstein, Luxembourg and Netherlands, using the Dutch, French and German languages

The number of pieces of medical information and languages can be: increased, such as to 4in1 Outsert, or

· decreased to 2in1 Outsert



Do not use does not apply to the 3in1 Outsert

Before using the 3in1 Outsert

Three products are made from only one printed sheet and sealed with perforated labels. Opening the perforated labels separates the package into its three components, which may consist of package leaflet, instructions for use and summary of product characteristics, for example.

The production of a single sheet guarantees 100% protection against mix-ups. Consistency of quality and precision is further assured by our inline measuring, in-process control and specialist staff.

Inform your management if you have recently used other outserts or package leaflets.

In comparison to other outserts and package leaflets, the 3in1 Outsert enhances:

 patient satisfaction Each single component looks more attractive and increases motivation to read, in comparison to

medical information that is three times larger. usability of provided medical information

After separation, each component can be more easily used in comparison to leaflets providing all

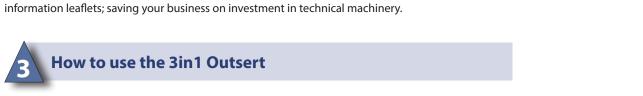
three components on one sheet.

• image of the medicine described in the 3in1 Outsert • image of the marketing authorisation holder

• satisfaction of the marketing authorisation holder See section 4.

The 3in1 Outsert has no negative influence on driving or using machines during packaging. On the

contrary, it facilitates an easier packaging process because you need only one feeder for several



Usual dose: read the 3in1 Outsert again if you have any questions Maximum dose: no limitation

Use this outsert until the updated 3in1 Outsert is produced.

An overdose does not harm; therefore, no actions are required.

Not recommended.

Possible side effects

The benefits of 3in1 Outsert prevail with correct use.

Frequencies of 3in1 Outsert side effects: **Very common**, affects more than 1 per 10 users

reduced production costs of combined medical information and multilingual package leaflets • easier, quicker and cost saving packaging of medicines with combined medical information and multilingual package leaflets

avoids necessity for new packaging machines if combined medical information and multilingual package leaflets are required avoids process and documentation changes if combined medical information is required $% \left(1\right) =\left(1\right) \left(1\right) \left($

· reduces maintenance costs of packaging machines 100% mix-up protection

Always inform your manager responsible for packaging if he/she has not yet noticed any of these side



How to store 3in1 Outsert

Store in the packaging of every relevant medicine in order to protect from moisture.

Do not use this outsert after the updated 3in1 Outsert is produced. The 3in1 Outsert is intended for you.



Further information

Composition

One 3in1 Outsert contains: • active substance: 100% innovation

 further ingredients: » 2 seal labels

» text template: the successfully readability tested 200 word model template

- Fuchs et al., The influence of the European Union's QRD template on the use of package inserts compared with a shorter model template, PharmInd 2012, 74(1): 126-136 - Wolf et al., Readability of the European QRD template - The European QRD template version 8 in

comparison to its predecessor and a shorter model template. PharmInd 2014, 76(8): 1312-1322 Free access to both studies at: http://www.paint-consult.com/en/publications/publications/ » thinstar® plus thinprint paper 60 gsm - a paper of delfortgroup

E-mail: info@baehren-druck.de

Manufacturer · 3in1 Outsert

Bähren Druck Mittelstraße 62, 41236 Mönchengladbach, Germany

Wenigenjenaer Ufer 12, 07749 Jena, Germany

T.S.P. Germanv GmbH, member of delfortgroup AG

Carl-Zeiss-Str. 2, 21465 Reinbeck, Germany



Bähren Druck

· text, layout/design

Phone: +49 3641 549396

Internet: www.paint-consult.com

E-mail: info@paint-consult.com

thinstar® plus thinprint paper

Internet: www.delfortgroup.com

E-mail: germany@delfortgroup.com

Phone: +49 40 780876 40

PAINT-Consult

Phone: +49 2166 97291 0 Internet: www.baehren-druck.de



delfortgroup PERFORMANCE BY UNDERSTANDING

Information last updated May 2016



◆ Good Manufacturing Practice guidelines (GMP)

 Climate neutral print production Forest Stewardship Council (FSC*)

♣ Hygiene Directives (HACCP)

♣ DIN EN I2O 8001

Responsibility

3. Paper selection

The paper weight chosen should be ... sufficiently thick to reduce transparency ... Glossy paper reflects" light making the information difficult to read, so the use of uncoated paper should be considered." recommends the readability guideline [1]. Opacity (light resistance) is a more appropriate parameter to assess paper quality than the paper weight [6, 7]. Thin paper with high opacity, such as thinstar® plus, ensures excellent legibility and helps to reduce the size of a folded leaflet where larger sheet formats are required.

General use of uncoated paper is unfounded. As coating can refine the surface opal as well as glossy, it is no criteria for non-reflecting surfaces. Moreover, coating can increase opacity, so that paper of lower thickness can be used than would be possible with the uncoated option. Thus, coated paper is an appropriate option and one with the added benefit of interrupting the domino effect of format increase in package leaflets.



216-page mini brochures · left coated paper with 88% opacity right uncoated paper with 87% opacity

Mittelstraße 62, 41236 Mönchengladbach, Germany, +49 2166 97291 0, info@baehren-druck.de 4. Production of the 3in1 Outsert Bähren Druck - The Pharma Specialist

thinstar plus paper 60 gsm

9102.20/20

♣ 2 seal labels 450 × 296 mm per product







Instructions for use

Many steps are required and must be carefully realised, so that users receive highly qualitative medical information. To manage these enormous tasks a finely tuned, cross-divisional cooperation is essential within each company; particularly between the departments of: • regulatory affairs/authorisation

 marketing/product design production/packaging

The advice of external specialists with regard to text, layout and packaging specifications - combined with a solid database - further ensures excellent medical information and internal resources within

1. Text creation

pharmaceutical companies.

Medical information must be legible, clear and easy to use. In the case of package leaflets, the readability guideline recommends simple words of few syllables, combined with short sentences and paragraphs, with use of bullet points and an active wording style. Abbreviations, acronyms and scientific symbols should be avoided [1]. However, the key problem of ongoing text increase is not addressed. This persistent increase in the word count of package leaflets is a main cause of larger leaflet formats and the resultant costly changes in the packaging process. At the same time, this text increase contradicts the consumer desire for shorter versions.

To this end, it is known that every increase in the number of words significantly decreases: patients' motivation to read package leaflets

locatability of provided information

patients' trust to use the respective medicine

· comprehensibility: longer leaflets contain significantly more difficult terms, abbreviations, nonquantifiable phrases, long sentences etc. than shorter versions [2, 3]

However, through systematic optimisation, package leaflet text can be substantially compressed by an average of 20%, without loss of essential medicine-related content [2]. For example, often contained repetitions are dispensable and their consistent avoidance reduces the number of words [4]. Where information is of particular importance, simply highlighting it by means of bold print or use of colour represents an effective and succinct solution.

Compressing the over 800-word QRD template for package leaflets to a text frame of headings and standard texts of around 200 words, proffers another effective and proven option [4, 5]. Authorities should take their responsibility here. Until such time as a shorter template becomes available, companies are advised to strictly avoid unnecessary optional QRD template texts.

2. Layout/design development

leaflet use, without disrupting the packaging process.

Readability guideline recommendations, such as using minimum 9pt font size, bullet points and a font in which similar letters/numbers can be easily distinguished - including avoiding words printed in uppercase letters only - improve legibility [1]. The same readability guideline recommends use of landscape format; therein denying the suitability

of all leaflets that are at least 1 mm higher than wide. Moreover, this is in the absence of any evidence that landscape outperforms the more space-saving portrait format [6].

Conclusion: All intended layout and design elements require careful selection, such that they benefit

After creating the text, typesetting the mock-up and paper selection, the 3in1 Outsert is produced in the following steps:

A) Printing one sheet



C) Margin cut of the folded sheet



D) Seal leaflets with perforated labels

B) Camera controlled folding of the sheet



B) Unfold leaflets

5. Use the 3in1 Outsert

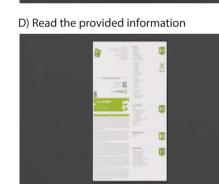
Use the 3in1 Outsert as follows: A) Open seal labels



C) Select the required leaflet

PharmInd 2015, 77(9):1380-1383





[1] European Commission. Guideline on the readability of the labelling and package leaflet of medicinal products for human use, Revision 1, 12 January 2009. http://ec.europa.eu/health/files/eudralex/ vol-2/c/2009_01_12_readability_guideline_final_en.pdf

[2] Fuchs J. The way forward in package insert user tests from a CRO's perspective. Drug Information Journal 2010, 44(2):119-129 [3] Fuchs J et al. Excessive medical information increase in package inserts. Int J Clin Pharmacol Ther

2010, 48(12):781-790 [4] Wolf et al. Readability of the European QRD template - The European QRD template version 8 in comparison to its predecessor and a shorter model template. PharmInd 2014, 76(8): 1312-1322 [5] Fuchs et al. The influence of the European Union's ORD template on the use of package inserts com-

[6] Fuchs J, Kutscha M. Handling des steten Textzuwachses in Packungsbeilagen. PharmInd 2015, 77(10):1438-1445 [7] Fuchs J, Kutscha M. How best to assess paper quality for package leaflets – weight or opacity?

pared with a shorter model template, PharmInd 2012, 74(1): 126-136

Free access to cited studies: http://www.paint-consult.com/en/publications/publications/

♣ Climate neutral print production

₽ DIN EN I2O 8001

thinstar plus paper 60 gsm

Z seal labels 420 × 296 mm per product

Je banel per product

Bahren Druck

4.5 Interaction with other medicinal products and other forms of interaction

The management must be informed if other outserts or leaflets have recently been used, to fully appreciate the benefits of the 3in1 Outsert for all distributed medicines.

Forest Stewardship Council (FSC°) In comparison to other outserts and leaflets, the 3in1 Outsert enhances:

 user satisfaction Each single component looks more attractive and increases motivation to read, in comparison to

◆ Good Manufacturing Practice guidelines (GMP) ♣ Hygiene Directives (HACCP)

medical information that is three times larger. · usability of provided medical information

After separation, each component can be more easily used in comparison to leaflets providing all Kesbousibility three components on one sheet. • image of the medicine described in the 3in1 Outsert and the marketing authorisation holder

• satisfaction of the marketing authorisation holder, see section 4.8 4.6 Fertility, pregnancy and lactation

No negative influence.

4.7 Effects on ability to drive and use machines

The 3in1 Outsert has no negative influence on driving or using machines during packaging. On the contrary, it facilitates an easier packaging process because only one feeder is required for several information leaflets; protecting businesses against unnecessary investment in technical machinery.

4.8 Desirable effects

Summary of the safety profile

The effects reported below during use of 3in1 Outsert significantly outweigh those which can be achieved with other outserts or leaflets.

Beneficial reactions reported for 3in1 Outsert are summarised in table 1. These reactions are presented by packaging system class and frequency.

Table 1: Beneficial reactions that have occurred during packaging of 3in1 Outsert

	Packaging system class	Very common (≥1 per 10 users)
	general costs	For combined medical information and multilingual package leaflets: reduced production costseasier, quicker and cost saving packaging
	packaging machines	 avoids necessity for new packaging machines if combined and multilingual medical information is required reduces maintenance costs of packaging machines
	packaging process and documentation	 avoids process and documentation changes if combined medical information is required
	mix-ups	• 100% mix-up protection

Reporting beneficial reactions after using the 3in1 Outsert is important. It allows continued monitoring of the positive benefit/risk balance with the 3in1 Outsert. Packaging specialists are asked to

Reporting of beneficial reactions

4.9 Overdose

information leaflets.

No negative overdose effects possible.

5. Pharmacological properties 5.1 Pharmacodynamic properties

Pharmacotherapeutic group: medical information leaflets, ATC code: XXL

tions) and 100% mix-up protection ensures effective and safe packaging.

report any of the above mentioned reactions to colleagues of other packaging units.

Packaging efficacy and safety Reduced costs (e.g. production and print; packaging machines, including processes and documenta-

5.2 Pharmacokinetic properties Significantly better kinetic properties exist for the 3in1 Outsert in comparison to leaflets three times

The 3in1 Outsert facilitates an easier packaging process because only one feeder is required for several

larger than the size of each single component. 5.3 Preclinical safety data

6. Pharmaceutical particulars

Adverse reactions are not observed.

6.1 List of excipients 2 seal labels

Until the updated 3in1 Outsert is produced.

 text and layout developed by PAINT-Consult • thinstar® plus thinprint paper 60 gsm - a paper of delfortgroup

6.2 Incompatibilities Not applicable.

6.3 Shelf life

6.4 Special precautions for storage

6.5 Nature and contents of container

The 3in1 Outsert is one pack intented for each medicine. It can be divided into three medical informa-

6.6 Special precautions for disposal and other handling

Bähren Druck, Mittelstraße 62, 41236 Mönchengladbach, Germany

7. Marketing authorisation holder

8. Marketing authorisation number

BD/0/00/000/0001

No special requirements.

9. Date of first authorisation/renewal of the authorisation

Phone: +49 2166 97291 0, Internet: www.baehren-druck.de, E-mail: info@baehren-druck.de

10. Date of revision of the text

Date of first authorisation: May 2013

Detailed information on this 3in1 Outsert is available at: www.baehren-druck.de.





3in1 Outsert

02/05.2016



Mittelstraße 62, 41236 Mönchengladbach, Germany, +49 2166 97291 0, info@baehren-druck.de

Bähren Druck - The Pharma Specialist

1. Name of the product

Summary of product characteristic

2. Qualitative and quantitative composition One 3in1 Outsert contains 100% innovation. For the full list of excipients, see section 6.1.

3. Pharmaceutical form The 3in1 Outsert is available in every layout/design, including colouration, as required.

4. Clinical particulars

4.1 Therapeutic indications 3in1 Outsert is indicated for all medicines to provide:

» package leaflet, instructions for use, SmPC » package leaflet, instructions for use, additional patient information

» package leaflet, SmPC, additional patient information · trilingual medical information, e.g. package leaflets for

three pieces of medical information, e.g.

» cost saving distribution to summarise markets like Austria, Belgium, France, Germany, Liechtenstein, Luxembourg and Netherlands, using the Dutch, French and German languages The number of pieces of medical information and languages can be increased, such as to 4in1 Outsert,

» distribution in small markets, such as the 3 Baltic countries or Finland, Norway and Sweden

or decreased to 2in1 Outsert. 4.2 Posology and method of administration

Starting dose: reading the 3in1 Outsert before using the medicine Usual dose: reading the 3in1 Outsert again if any question occurs

Paediatric population No limitations exist in children with sufficient reading skills.

4.3 Contraindications Does not apply to the 3in1 Outsert.

Method of administration

Maximum dose: no limitation

4.4 Special warnings and precautions for use Three products are made from only one printed sheet and sealed with perforated labels. Opening the perforated labels separates the package into its three components, which may consist of a package leaflet, instructions for use and summary of product characteristics, for example. The production of a single sheet guarantees 100% protection against mix-ups. Consistency of quality

Use the 3in1 Outsert after opening the seal labels and unfolding the relevant leaflet.

is further assured by inline measuring, in-process control and specialist staff.



Bähren Druck - The Pharma Specialist

+ Services

Launch

SMI/VMI-supplier/

specifications

Identification tags

vendor managed inventory

· Electronical document delivery

Product-/packaging development

Customised system adjustments

· Number range management

Construction of technical drawings/

+ Labels

 Cut & stack labels Sticker · Self-adhesive labels Barcode labels

· Identification labels · Clear-on-clear labels Tamper evidence labels Security features

» Micro font » Fluorescence inks

» Holograms » securPharm°

» Safety die cutting

» Coin reactive inks

» Individualization

★ Combinations/Multi-Packaging Patient Alert Cards Piggybacks

Finishing

» Braille

» Varnish

» Laminate

» Sandwich

Special materials

» No-Label-Look

» Unique Fiber-Fingerprint

» Void / Split / No-Stick

» Embossing

» Sequential back numbering







+ Leaflets

Flat

Folded

» with seal labels

» with glue points

» with perforations

» with aquascoring

· 2in1 or 3in1 Outsert

Connection sheet

t[w]o tear[®]

Outserts/Inserts [incl. Vijuk-license]

» integrated mailing solutions





Mittelstraße 62, 41236 Mönchengladbach, Germany, +49 2166 97291 0, info@baehren-druck.de



Wenigenjenaer Ufer 12, 07749 Jena, Germany, +49 3641 549396, info@paint-consult.com

Readability tests

PAINT

Package Insert Test

Readability tests of package leaflets, clinical trial and other medical information in English, German and all other official European Union languages

Bridging of package leaflets according to current guidelines, for all approval procedures

Typesetting

Layout and design for any kind of medical information and packaging materials

Medical writing

Creating and optimisation all types of medical information, based on our extensive research

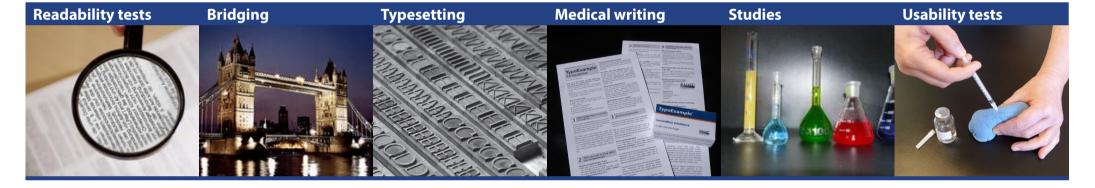
Studies of medicinal and healthcare issues

Usability tests

Tests whether patients can properly use application devices and medicines within the confines of the product specific requirements

T.S.P. Germany GmbH, member of delfortgroup AG

Carl-Zeiss-Str. 2, 21465 Reinbeck, Germany, +49 40 780876 40, germany@delfortgroup.com





PERFORMANCE BY UNDERSTANDING

thinkthin

thinprint-

thinkthin Finest quality, wood-free coated and uncoated premium papers from delfortgroup, your specialist for tailor-made papers and superior service. Be bright and keep it light!

thinprint paper

Our thinprint paper products are of unique quality, both in brightness and opacity. Our coated and uncoated, wood free paper ensures that maximum yield is achieved, saving you time and money. Constant innovation and best service give us an edge, which we wish to share with you. Contact one of our T.S.P. sales offices worldwide to experience the ultimate in thinprint paper technology: www.delfortgroup.com



Product name	3in1 Outsert (PIL, IFU, SmPC)
Size	3 × 296 mm × 420 mm
Date	24 May 2016
PAINT® Package Insert Test	PAINT-Consult®, Wenigenjenaer Ufer 12, 07749 Jena, Germany E-mail: info@paint-consult.com Tel.: +49 3641 549396

Bähren Druck - The Pharma Specialist

• Outserts/Inserts [incl. Vijuk-license]

Mittelstraße 62, 41236 Mönchengladbach, Germany, +49 2166 97291 0, info@baehren-druck.de

Services

Launch

SMI/VMI-supplier/

specifications

vendor managed inventory

Electronical document delivery

· Construction of technical drawings/

+ Labels

Bähren Druck

 Cut & stack labels Sticker · Self-adhesive labels

Security features

» Fluorescence inks

» Coin reactive inks

» Safety die cutting

» Individualization

» Micro font

» Holograms

» securPharm[®]

» Sequential back numbering Barcode labels · Identification labels Clear-on-clear labels

Tamper evidence labels

» Braille » Embossing » Varnish » Laminate Special materials » No-Label-Look » Sandwich

Finishing

» with seal labels » with glue points » with perforations » with aquascoring » integrated mailing solutions » Unique Fiber-Fingerprint · 2in1 or 3in1 Outsert » Void / Split / No-Stick t[w]o tear[®] · Connection sheet

+ Leaflets

Flat

Folded

+ Combinations/Multi-Packaging Patient Alert Cards

Piggybacks



PAINT-Consult® - The Readability Test Specialist

Wenigenjenaer Ufer 12, 07749 Jena, Germany, +49 3641 549396, info@paint-consult.com

Readability tests

Package Insert Test

Readability tests of package leaflets, clinical trial and other medical information in English, German and all other official European Union languages

Bridging of package leaflets according to current guidelines, for all approval procedures

Typesetting

Layout and design for any kind of medical information and packaging materials

Medical writing

Creating and optimisation all types of medical information, based on our extensive research

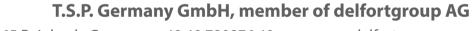
Studies of medicinal and healthcare issues

Usability tests

Tests whether patients can properly use application devices and medicines within the confines of the product specific requirements







Carl-Zeiss-Str. 2, 21465 Reinbeck, Germany, +49 40 780876 40, germany@delfortgroup.com

delfortgroup

thinkthin®

Finest quality, wood-free coated and uncoated premium papers from delfortgroup, your specialist for tailor-made papers and superior service. Be bright and keep it light!

thinkthin thinprint-

thinprint paper

Our thinprint paper products are of unique quality, both in brightness and opacity. Our coated and uncoated, wood free paper ensures that maximum yield is achieved, saving you time and money. Constant innovation and best service give us an edge, which we wish to share with you. Contact one of our T.S.P. sales offices worldwide to experience the ultimate in thinprint paper technology: www.delfortgroup.com



Bähren Druck

Bähren Druck - The Pharma Specialist

Mittelstraße 62, 41236 Mönchengladbach, Germany, +49 2166 97291 0, info@baehren-druck.de

+ Labels

· Cut & stack labels

· Self-adhesive labels

· Identification labels

» Fluorescence inks

» Coin reactive inks

Barcode labels

 Finishing » Sequential back numbering

» Braille

· Clear-on-clear labels · Tamper evidence labels Security features » Micro font

Sticker

 Number range management · Product-/packaging development

· Customised system adjustments Identification tags

» Holograms + Combinations/Multi-Packaging » securPharm[®] » Safety die cutting Patient Alert Cards Piggybacks » Individualization

» Embossing » Varnish » Laminate Special materials » No-Label-Look » Sandwich » Unique Fiber-Fingerprint

» Void / Split / No-Stick

» with seal labels » with glue points » with perforations » with aquascoring » integrated mailing solutions • 2in1 or 3in1 Outsert

Outserts/Inserts [incl. Vijuk-license]

vendor managed inventory Construction of technical drawings/ specifications

SMI/VMI-supplier/

★ Services

Launch

· Electronical document delivery Number range management Product-/packaging development · Customised system adjustments

Identification tags



Leaflets

Flat

Folded

t[w]o tear[®]

· Connection sheet



PAINT-Consult® - The Readability Test Specialist Wenigenjenaer Ufer 12, 07749 Jena, Germany, +49 3641 549396, info@paint-consult.com

Creating and optimisation all types of medical information, based on our extensive

Readability tests of package leaflets, clinical trial and other medical information in

English, German and all other official European Union languages

Bridging of package leaflets according to current guidelines, for all approval procedures

Typesetting

Layout and design for any kind of medical information and packaging materials

Usability tests

research

Medical writing

Tests whether patients can properly use application devices and medicines within

Studies of medicinal and healthcare issues

the confines of the product specific requirements





T.S.P. Germany GmbH, member of delfortgroup AG Carl-Zeiss-Str. 2, 21465 Reinbeck, Germany, +49 40 780876 40, germany@delfortgroup.com

thinkthin®

Finest quality, wood-free coated and uncoated premium papers from delfortgroup, your specialist for tailor-made papers and superior service. Be bright and keep it light!

thinkthin thinprint-

Our thinprint paper products are of unique quality, both in brightness and opacity. Our coated and uncoated, wood free paper ensures that maximum yield is achieved, saving you time and money. Constant innovation and best service give us an edge, which we wish to share with you. Contact one of our T.S.P. sales offices worldwide to experience the ultimate in thinprint paper technology: www.delfortgroup.com

