# Dried blood spot microvolume sampling for DMPK

#### Cutting the cost of generating PK/TK data

Pharmacokinetic (PK) and toxicokinetic (TK) studies provide crucial insight into how drug candidates behave in the body and are, therefore, a critical step in drug development.

These analyses traditionally require large volumes of blood (generally 100–500 µl per subject per time-point) in order to provide sufficient plasma volume for quantitative bioanalysis. Because only a limited number of serial samples can be taken from each animal, composite sampling is often used, resulting in lower quality PK data and an increase in the number of animals required. Large volumes also make it difficult to conduct studies in juvenile subjects.

Additionally, the plasma must be isolated from whole blood and prepared for bioanalysis using solid phase extraction, liquid-liquid extraction, or protein precipitation. This timeconsuming process limits throughput and consequently the number of samples that can be tested.

Finally, there are the practical challenges of shipping and storing blood samples, which require controlled handling and frozen transportation and storage.

The Dried Blood Spot (DBS) method is an alternative technique that overcomes these drawbacks and delivers significant cost savings.

#### Spot. Extract. Analyze.

- Apply blood to card (Fig 1.) and let dry. Ship and store as needed at ambient temperature.
- Remove sample disc (Fig 2.) and elute with solvent.
- Analyze solution by HPLC-MS/MS.



Fig 1. Apply sample

Fig 2. Punch disc

DBS microvolume sampling using specialized Whatman<sup>™</sup> media from GE Healthcare has been shown to be both precise and accurate for a variety of compounds from different structural classes with acceptable inter- and intra-assay variability and is now being routinely employed in PK/TK studies. The FTA<sup>™</sup> DMPK-A and FTA DMPK-B cards lyse cells and denature proteins on contact. Samples can be shipped and stored at ambient temperature and long-term stability has been demonstrated for analytes and metabolites sensitive to plasma enzymes.

#### Low blood volumes

- $\bullet$  DBS microvolume sampling requires only 10 to 20  $\mu l$  per sample.
- Fewer experimental animals are required because toxicokinetic data is derived from toxicology animals, thereby eliminating satellite colonies.
- More consistent data is obtained through more serial sampling from individual animals and less reliance on composite data, further reducing the number of animals needed.
- Juvenile animals and humans can be studied.



# Whatman

#### Simple and safe processing

- The 3-step DBS procedure is much more straightforward than the cumbersome centrifugation, isolation and clean up of plasma.
- On-substrate clean up has the potential to convey greater analyte stability, especially for enzyme-sensitive compounds.

#### Easy storage and transportation

- Room temperature stability saves on cost of dry ice shipments.
- Shipping without dry ice makes it simpler and more cost effective to carry out remote sampling, or clinical studies.

#### **Getting started**

The choice of DMPK card depends on many factors such as the analyte chemical structure, extraction solvent and analysis workflow. The final decision is usually determined empirically.

### Ordering information

| Description           | Pack size         | Catalog number |
|-----------------------|-------------------|----------------|
| FTA DMPK-A Card       | 100               | WB129241       |
| FTA DMPK-B Card       | 100               | WB129242       |
| FTA DMPK-C Card       | 100               | WB129243       |
| FTA DMPK Starter Pack | 45 cards          | WB129248       |
|                       | (15 x A, B and C) |                |

#### Accessories for punching and drying

| Description           | Pack size       | Catalog number |  |  |  |
|-----------------------|-----------------|----------------|--|--|--|
| Harris Uni-Core       | 4 (with cutting | WB100039       |  |  |  |
| Punch 3.0 mm          | mat)            |                |  |  |  |
| Harris Micro-Punch    | 1               | WB100038       |  |  |  |
| 3.0 mm                |                 |                |  |  |  |
| Replacement Cutting   | 1               | WB100020       |  |  |  |
| Mat, 5-7/8" x 7-7/8"  |                 |                |  |  |  |
| Dry Rak (without      | 10 (holds 11    | 10537173       |  |  |  |
| Velcro™)              | cards/rack)     |                |  |  |  |
| Dry Rak (with Velcro) | 10 (holds 11    | 10539521       |  |  |  |
| •                     | cards/rack)     |                |  |  |  |
|                       |                 |                |  |  |  |

#### Accessories for storage and shipping

| Description                             | Pack size | Catalog number |
|---|-----------|----------------|
| Zipper Sealed Storage Bags, 4"x6"       | 100       | 10548232       |
| Foil Barrier Zipper Sealed Storage Bags | 100       | 10534321       |
| Desiccant Packets (indicating), 1g      | 1000      | WB100003       |
| Glassine Envelopes, 3-1/4" x 4-7/8"     | 100       | 10548236       |
| Biohazard Labels, 7/8" x 7/8"           | 1000      | 10534150       |

#### **Contact information**

For more information please contact your local sales representative or your local office.

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| www.whatman.<br>com/dmpk.aspx   | Contact your<br>local office  | dmpk@ge.com                                  |