

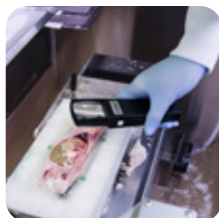
A flexible approach to  
**DMPK**

**QPS DMPK PROVIDES A DEDICATED  
TEAM OF SENIOR SCIENTISTS TO HELP SELECT,**  
design and conduct the appropriate ADME studies  
for your specific compounds and therapeutic targets.  
Working with QPS DMPK is a collaborative and consultative  
endeavor that also incorporates our operational effectiveness  
and dedication to customer service.

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**TIME IS OF THE ESSENCE IN DRUG DEVELOPMENT.  
CONTACT THE QPS BUSINESS DEVELOPMENT TEAM TODAY!**

**CALL** +1 512 350 2827 **EMAIL** [infobd@qps.com](mailto:infobd@qps.com)



## Macro & Micro Autoradiography

One of QPS' main services in DMPK is the combined mass balance, biliary excretion, PK, and QWBA studies.

- ▶ Increased efficiency by using a single study protocol
- ▶ Better study oversight with an integrated study report
- ▶ Elimination of the possibility of radiolabel degrading over an extended period of time between the two studies
- ▶ Metabolite profiling and identification can be done using the samples from the same study
- ▶ Micro-autoradiography provides insight into sub-cellular tissue distribution

## Biotransformation

QPS biotransformation studies determine how a molecule may be altered by the action of enzymes. These study types include:

- ▶ *In vitro* metabolic stability in hepatic subcellular fractions to determine intrinsic clearance
- ▶ *In vitro* comparison of metabolite formation in animal and human hepatic preparations, using non-labeled and radio-labeled test articles
- ▶ *In vivo* metabolite profiling, identification and quantification using samples collected from animal PK, mass balance excretion studies, and human AME studies to satisfy Metabolites in Safety Testing (MIST)

## Protein Binding

QPS protein binding studies determine the extent of protein binding in various plasma species and tissues.

- ▶ Methods: equilibrium dialysis (RED, Harvard device, 96-well HTD), ultrafiltration, and ultracentrifugation
- ▶ Discovery screening, *in vitro* protein binding for IND, and *ex vivo* studies in clinical phases
- ▶ Experience in compounds with very high binding, stability issues, or non-specific issues

## Drug-Drug Interaction

QPS drug interaction studies determine the potential of a substance to alter cytochrome P450 activity. Studies conducted to assess inhibition and induction potential to support discovery and development of new drug candidates are:

- ▶ *In vitro* inhibition characterization in human liver microsomes or hepatocytes to determine reversible or time dependent IC50
- ▶ *In vitro* mechanistic characterization of reversible inhibitory rate constants  $K_i$
- ▶ Identification of the reversible mechanism of inhibition as competitive, non-competitive or uncompetitive
- ▶ *In vitro* mechanistic characterization of time dependent inhibition  $K_{inact}$  and  $K_i$
- ▶ Identification of the mechanism of time dependent inhibition as metabolite mediated via covalent modification or due to tight binding effect of substrate
- ▶ *In vitro* characterization of CYP1A2, CYP2B6, and CYP3A4 induction potential in human hepatocytes based on mRNA and/or CYP activity using isoform selective probe substrates

## QPS is a Global CRO with locations around the world

