Valuing R&D Projects in a Portfolio: Evidence from the Pharmaceutical Industry

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Understanding the value of a product development project is central to a firm’s choice of project portfolio. The value of a project to a firm depends not only on its properties but also on the other projects being developed by the firm. This is due to interactions with the other projects that address the same consumer need and require the same development resources. In this study, we empirically investigate the structure and significance of these portfolio-level project interactions. Using a self-developed pharmaceutical industry data set, we conduct an event study around the failure of phase III clinical trials and their effect on the market valuation of the firm. The study exploits the natural experiment of a product development failure to give us a measure of the value of a drug development project to a firm. We then explain the variance in the value of projects based on interactions with other projects in the firm’s portfolio. We find that the presence of other projects targeting the same market and a build-up of projects that require the same development resources reduce the value of a development project. In addition to providing evidence on the significance and structure of these portfolio-level project interactions, the empirical model estimated in this paper also provides a data-driven approach to valuing projects that may be relevant to licensing transactions.

Key words: product development; pharmaceuticals; development pipeline; portfolio properties; backup projects; portfolio management

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1. Introduction

Understanding the value of a product development project is central to the scientific management of the product development process. The value of a project to a firm depends not only on its properties but also on those of the other projects the firm is developing. This is due to interactions with the other projects in the firm’s same development portfolio. Understanding these portfolio-level project interactions is central to a firm’s choice of project portfolio and development capacity (Loch et al. 2001a, Kavadias and Loch 2003). Decision support models for portfolio choice have provided analytical models of these interactions (cf. Ding and Eliashberg 2002, Loch and Kavadias 2002). In this study, we empirically investigate the structure and significance of these interaction effects.

We use the natural experiment of a product development failure to estimate the value of an individual project. We design an event study around the failure of a late-stage development project. This event study gives us a metric of the change in the firm’s value (as measured by the stock markets) due to the failure of a development project with all other factors affecting the firm’s value being held constant (MacKinlay 1997). This change in firm value is an empirical measure of the value of the failed project to the firm. We then explain the variance in the value of all failed projects in our sample based on the interactions of the project with other projects in the portfolio. Specifically, we investigate how the value of a project to a firm may depend on the presence of other projects in the firm’s portfolio that address the same customer need or utilize the same development resources.

The specific context of our empirical examination is the pharmaceutical industry. New product development in the pharmaceutical industry is regulated and thus, proceeds along a series of well-defined steps illustrated in Figure 1. Drug development starts with an investigation of the chemical and biological properties of a compound in the lab (basic research), followed by animal trials (preclinical studies) and, finally, three stages of clinical trials or trials in human subjects (phase I, II, and III). Our study is designed around the failure of development projects currently undergoing phase III clinical trials, which is the final stage in the development process, where the efficacy
Figure 1  Drug Development Process

<table>
<thead>
<tr>
<th>Stage</th>
<th>Key issue</th>
<th>Average duration (years)</th>
<th>Average output rate (compounds/year)</th>
<th>Average cost of phase per unit of output (MMS/compound)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Basic research</td>
<td>Compound discovery and screening</td>
<td>1.5</td>
<td>12.5</td>
<td>12</td>
</tr>
<tr>
<td>Preclinical</td>
<td>Animal testing for safety and metabolism</td>
<td>1</td>
<td>6.25</td>
<td>13</td>
</tr>
<tr>
<td>Phase I</td>
<td>Safety in small sample of healthy volunteers</td>
<td>2</td>
<td>3.75</td>
<td>32</td>
</tr>
<tr>
<td>Phase II</td>
<td>Safety and efficacy in sample of patients</td>
<td>2</td>
<td>1.25</td>
<td>141</td>
</tr>
<tr>
<td>Phase III</td>
<td>Safety and efficacy in large sample of patients</td>
<td>1</td>
<td>1.13</td>
<td>220</td>
</tr>
<tr>
<td>NDA review</td>
<td>FDA review of filling</td>
<td>1</td>
<td></td>
<td>24</td>
</tr>
<tr>
<td>Medical indications</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Notes. Symbols  are different patient needs (clinical target indicators). The values are for a typical pharmaceutical company and are normalized based on an average annual output of one compound. Development projects that target the same market or indication are denoted with the same symbol. Estimates were obtained from the Parexel pharmaceutical R&D Statistical Source Book, 2002/2003.

and safety of the drug is investigated in a large sample of patients. Common causes for failures at this stage include adverse side effects of the drug and harmful interactions with other drugs. For a detailed description of the drug development process, the reader is referred to Pisano and Rossi (1994) and Girotra et al. (2004).

The pharmaceutical industry presents an ideal domain of enquiry for our study. It is a large industry where product development is central. Although the product development process closely resembles the classic stage-gate development process prevalent in most industries, the role of regulators in the later phases of the drug development process significantly simplifies the empirical design of our study. The different stages in the product development process are clearly and uniformly defined by the regulator, and all pharmaceutical firms must pass their development projects through the same development stages. This allows us to identify projects at the same stage of development across different firms in the industry. The results of each stage of the development process are public knowledge. This allows us to create our data set of product development failures from public sources. Finally, the late-stage product development portfolio of each firm is public knowledge. Thus, when the stock markets value failures, they have the information on the portfolio-level project interactions which we are investigating.

We find empirical evidence for two portfolio-level project interactions. First, we find that the impact of the failure, which is our measure for the value of the project to the firm, is smaller when the firm is developing other projects for the same market as the failed project. When the firm is developing multiple projects for the same market, the failure of one of them does not preclude the firm from earning sales in that market. Thus, the marginal value of any of the multiple projects is smaller than the value of a single project being developed for the market.

Second, we find evidence that the value of a compound or the impact of a failure is smaller if the firm has more projects than the anticipated number in its portfolio, which require the same resources as those used by the failed project (even if the other projects are not “backups” for the same market). A failure leads to the freeing up of resources shared by the failed and other projects. These freed-up resources can be redirected to other projects, which may then be brought to the market sooner than they would be if there was no failure. Thus, failures in portfolios with more than the anticipated number of projects that utilize the same resources as the failed project lead to the
acceleration of the other compounds in the portfolio and have a smaller impact.

This study enhances our understanding of portfolio-level project interactions. We build and validate a theory on the financial effect of these interactions. We find that these interactions significantly alter the value of a development project to a firm and are thus crucial to portfolio choices. Our empirical evidence also allows for a critical examination of the existing analytical literature on portfolio and capacity choices with respect to the modeling of project interactions. This can help us understand the reasons behind the limited application of this literature in practice (Loch et al. 2001a, Loch and Kavadias 2002, Shane and Ulrich 2004) and inspire new improved analytical models, which take into account the empirical regularities that we find. Finally, our results also provide a data-driven model that aids in valuing individual projects in the context of the product development portfolio of a firm. This is useful in valuing development projects available for in-licensing and comparing alternative development projects.

2. Prior Literature

Two streams of academic work are relevant to this study: (1) research on portfolio choices and (2) research on the financial impact of product development outcomes. An established body of literature in operations research attempts to provide optimal product portfolio decisions. Initially, optimization models were developed in a static and deterministic setting, with the decision modeled as one-shot choice under complete information, often with a mathematical programming formulation (see e.g., Lucas 1971). More recent work has emphasized the stochastic, dynamic, or process nature of the problem and has analyzed capacity and congestion effects (Loch and Terwiesch 1999) as well as strategies for search and information gathering (Loch et al. 2001b, Dahan and Mendelson 2001).

Portfolio-level project interactions are central in many of the contemporary models on portfolio selection. Loch and Kavadias (2002) present a dynamic model of portfolio selection within a budget constraint, taking into account multiple project interactions, including those arising out of shared markets in a general setting. In their study on the number of development approaches to pursue for a given market, Dahan and Mendelson (2001) model the interactions between projects of different quality that target the same market. Ding and Eliashberg (2002) investigate the number of development approaches to pursue for a given market in a staged development process. They build an analytical model of the interactions between projects targeting the same market. In their model, unlike Dahan and Mendelson (2001), all successful projects are assumed to have identical quality. In our paper, we empirically examine interactions similar to those investigated by Ding and Eliashberg (2002).

Adler et al. (1995) build a model of project interactions due to shared development resources. Using a development project as their unit of analysis they find that if development resources are stretched, the project completion times are longer. In contrast to Adler et al. (1995), we take an empirical approach and study the effect of shared development resources at the portfolio level. We examine the impact of one project on other projects in the portfolio. Further, we find the impact of these interactions on the financial value of the project as opposed to the development lead time.

Multiple studies have focused on the impact of product development events on financial value, notably Hendricks and Singhal (1997) on the impact of product development delays. They find significant negative stock returns associated with the announcement of product introduction delays. Industry competitiveness and the firm’s degree of diversification influence the size of this impact. Chaney et al. (1991) and Chaney and Devinney (1992) study the stock market reaction to announcements of new products across a wide range of industries. Bayus et al. (2003) study the impact of new product introductions in the personal computer industry on profit rate, profit rate persistence, and asset growth. Robertson et al. (1995) and Chen et al. (2005a) study the impact of new product announcements on competing firms. Chen et al. (2005b) examine the effect of product introduction delays on industry rivals. Sharma and Lacey (2004) compare the impact of pharmaceutical successes and failures on firm value. This body of work quantifies the impact of these product development events. Further, they explain the variance in the financial impact of product development with product or industry properties, but not the portfolio.

We build on the rigorous methodologies developed in this literature to empirically value projects. However, in contrast to this literature, we relate the impact of the product development outcomes (our measure for the financial value of projects) to key properties of the product development portfolio—the presence of other compounds in the portfolio that target the same unmet market need, and the availability of research opportunities that can utilize resources freed up due to the failure.

3. Theory Development

Failure of a late-stage development project represents the loss of potential future sales for a firm, which should lead to a decrease in the value of the firm.
In the pharmaceutical industry, a drug undergoing phase III clinical trials has an average approval probability of about 80% (Parexel 2002/2003). On approval, an average drug has the potential to generate sales of hundreds of millions of dollars. When a phase III failure occurs, these potential sales are lost; the 20% probability of failure is updated to certainty. This gives us our baseline hypothesis:

**Hypothesis 1.** The value of a pharmaceutical firm falls when a compound fails in phase III clinical trials.

Not surprisingly, previous research has proposed and found evidence for similar hypotheses. Sharma and Lacey (2004) propose a similar hypothesis in their study of stock market reactions to news from the pharmaceutical industry.

### 3.1. Effect of Projects Targeted at the Same Market

Drug development, like most other product development, is associated with long development lead times and low odds of success. The clinical trials phase of the drug development process alone takes an average of six years to complete, and only one out of six drugs that enter clinical trials makes it to the market. Fortunately, there are often multiple, unrelated technological approaches available to address the same medical need. For instance, there are multiple chemical compounds that pharmacologically inhibit the COX enzymes, which provide relief from the symptoms of inflammation and pain. These compounds differ in their side effects and thus, their success or failure in late-stage clinical trials are largely unrelated. In such settings, firms follow a parallel development strategy that increases the likelihood of developing a viable product for a given lucrative market within a reasonable time frame. The candidate compound farthest along in the development process is referred to as the lead compound, and the other compounds are referred to as backup compounds. Such parallel development strategies have been shown to be optimal in a variety of product development settings where the odds of success are low, the development lead times are long, and the correlation between the successes of different concepts is low (Loch et al. 2001b, Ding and Eliashberg 2002).

Although firms often develop multiple compounds to address a given market opportunity, firms rarely market more than one of them. Introducing several drugs for the same medical need earns the same sales as introducing only one successful compound (Ding and Eliashberg 2002, Girotra et al. 2004). Firms typically introduce the first one that passes clinical trials, and then cease development of all other drugs targeted toward this market. Consequently, the probability of having at least one successful drug for a given market is a crucial metric related to the financial value of a product development pipeline. By developing multiple compounds for the same market, firms increase this probability. Thus, the marginal value of a compound to a firm is proportional to how much the compound increases the probability of having a successful drug in its market for the firm. To illustrate this concept consider the two scenarios provided in Figure 2.

In Figure 2, the compounds of concern have a probability of failure of 20%. In failure 1, the firm is

**Figure 2** Example Illustrating the Role of Backup Compounds

<table>
<thead>
<tr>
<th>Failure 1</th>
<th>Before failure:</th>
<th>After failure:</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Probability of having at least one successful drug-0.8</td>
<td>Probability of having at least one successful drug-0</td>
</tr>
<tr>
<td></td>
<td>Change in probability-0.8</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Failure 2</th>
<th>Before failure:</th>
<th>After failure:</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Probability of having at least one successful drug-0.96</td>
<td>Probability of having at least one successful drug-0.8</td>
</tr>
<tr>
<td></td>
<td>Change in probability-0.16</td>
<td></td>
</tr>
</tbody>
</table>
developing only one compound for the indication. Prior to the failure, the firm has an 80% probability of obtaining at least one successful drug. After the failure, this probability changes to 0%. Due to the failure, the probability of earning the sales from this indication falls by 80 percentage points. Equivalently, the change in the expected future cash flows from the pipeline due to the failure is proportional to 80 points. Alternatively, consider failure 2, where the firm is developing two compounds for the indication. Prior to the failure, the firm had a 96% \( (1-(0.2)^2) = 0.96 \) chance of having at least one successful drug in the indication. After the failure, this changes to 80%. As a result of failure 2, the change in probability is 16 percentage points. The change in the expected future cash flows from the pipeline is thus proportional to 16 points. The presence of a backup compound in the case of failure 2 mitigates the impact of the failure. This should be reflected in the stock market reaction to the failures, or the valuation of these development projects. Next, we extend this logic to a general portfolio with compounds in each of the three phases of development.

Consider a pipeline with \( n_i \) candidate drugs undergoing phase \( i \) \((i=1, 2, 3)\) development for the market in question. Let \( p_i \) denote the probability of a drug currently in phase \( i \) trials during phase \( i \) or in any subsequent phase.\(^1\) From this pipeline the firm could earn the sales from the market under three mutually exclusive realizations of the clinical trials: (1) One of the compounds currently undergoing phase III trials succeeds\(^2\) (probability \( (1-p_3^{n_3}) \)); (2) all the compounds currently undergoing phase III trials fail, but one of the compounds currently undergoing phase II trials succeeds in phase II trials and in phase III (probability \( p_3^{n_3} (1-p_3^{n_3}) \)); (3) all the compounds currently undergoing phase II trials fail, but one of the compounds currently undergoing phase I trials succeeds in phase I trials and all subsequent phases (probability \( p_1^{n_1} p_2^{n_2} (1-p_3^{n_3}) \)).

Under each of the three scenarios, the firm earns the sales from the market; however, scenarios 2 and 3 have longer times to market. Thus, their sales should be discounted using appropriate discount factors, \( \alpha_2 \) and \( \alpha_1 \), respectively. The expected net present value of sales from this indication is thus given as

\[
E[NPV] = E[M] \times [ (1-p_3^{n_3}) + \alpha_2 p_3^{n_3} (1-p_3^{n_3}) + \alpha_1 p_1^{n_1} p_2^{n_2} (1-p_3^{n_3}) ].
\]

Here, \( E[M] \) is the expected value of sales from the target market conditional on having a successful drug. Equivalently,

\[
E[NPV] \propto (1-p_3^{n_3}) + \alpha_2 p_3^{n_3} (1-p_3^{n_3}) + \alpha_1 p_1^{n_1} p_2^{n_2} (1-p_3^{n_3}),
\]

where the expected sales serve as the proportionality constant. We refer to the right-hand side of the above expression as the time-adjusted probability of indication success (or TAPIS).

\[
TAPIS(p_1, p_2, p_3; n_1, n_2, n_3) = [(1-p_3^{n_3}) + \alpha_2 p_3^{n_3} (1-p_3^{n_3}) + \alpha_1 p_1^{n_1} p_2^{n_2} (1-p_3^{n_3})].
\]

TAPIS is proportional to the expected financial returns from the development projects. The marginal value of a phase III compound, or the impact of losing a phase III compound in the portfolio, is thus proportional to the consequent change in TAPIS \((n_3 \rightarrow n_3-1)\):

\[
\Delta TAPIS = TAPIS(n_3) - TAPIS(n_3-1) = (P_3^{n_3-1} - P_3^{n_3}) \left[ 1 - \alpha_2 (1-P_3^{n_3}) + \alpha_1 P_1^{n_1} P_2^{n_2} (1-P_3^{n_3}) \right].
\]

Equation (2) is the change in the time-adjusted probability of earning the sales from the target market. This is proportional to the reduction in the expected value of the future cash flows from the sales in the target market. Thus, failures for which the expression in (2) is large should have a higher financial impact.

**Hypothesis 2.** The change in the value of a pharmaceutical firm when a phase III compound fails is negatively correlated with the change in the time-adjusted probability of success for an indication (\( \Delta TAPIS \) in Equation (2)).

3.2. Effect of Other Compounds in the Portfolio That Utilize the Same Development Resources

Phase III trials involve the examination of the safety and efficacy of the drug in a large sample of patients. The primary resources required at this stage are clinical trial sites and bio-statisticians. Irrespective of the disease or indication, all compounds draw from the same pool of resources. It is quite costly to scale the capacity of these resources up or down in the short term because these resources are mostly professionals hired with a multi-year expectation of employment, or fixed assets, which take time to set up. Consequently, firms must make long-run commitments to the capacity of these sticky phase III resources. Firms set up phase III capacity well in advance of observing the results of the most recent phase II trials or the current demand for phase III resources. On the basis of the long-run or expected probability of success in phase II and the phase II capacity, firms forecast the demand for phase III...
resources, and make long-term commitments to R&D capacity (Girotra et al. 2004).

For instance, consider a firm that from its past experience estimates that the average phase II success rate across all diseases is 50%. If the firm has the capacity to process four compounds per year in phase II clinical trials, it expects that two of the four compounds will succeed and proceed to phase III resources. Thus it establishes phase III capacity as 50% of phase II capacity, two compounds per year.

The actual demand for phase III resources at any point in time is a function of the actual realizations of recent phase II trials. Continuing with our example of the firm that has set phase III capacity to two compounds per year, consider two possible realizations of phase II trials (Figure 3): (1) The realized success rate is 50% (two of the four compounds in phase II trials succeed); (2) the realized success rate is 75% (three of the four compounds in phase II trials succeed). In case 2, the firm has more compounds than the available phase III resources, thus one compound has to wait in a "buffer" before phase III.

Now consider that out of the two compounds currently undergoing phase III trials in each of the above scenarios, one compound fails; e.g., the compound denoted by the rhombus in Figure 3. In both cases, the firm loses the potential future sales from the failed compound; however in case 2, there is a mitigating effect: One compound is waiting in the buffer (denoted by the square in Figure 3), which can now take advantage of the freed-up phase III resources. The value of this waiting compound actually increases as it can now enter phase III trials and be brought to the market earlier. There is no such mitigating effect of the failure in case 1. Thus, the impact of the failure in case 2 should be smaller than the impact of the failure in case 1.

Failures that come at a time when the realized phase II success rate in the firm’s pipeline is higher than expected lead to an acceleration of the other compounds in the pipeline, and their impact should therefore be lower. Further, the benefit associated with the acceleration depends on the unanticipated demand, or the number of compounds that are waiting in the “buffer,” captured by the degree to which the realized success rate was higher than the expected phase II success rate.

Although the above example uses the notion of a buffer to illustrate this effect, literal presence of a buffer is not required for the acceleration. As long as a higher resource utilization of resources leads to longer processing times, the freeing up of resources due to failures will have the beneficial impact of accelerating compounds in the pipeline. The higher the utilization, the higher the benefit, because the extent of the acceleration and the number of projects that benefit from this acceleration are both higher at higher levels of utilization.

Hypothesis 3. The decrease in firm value from a phase III failure is lower (higher) if the firm has experienced an above (below) average phase II success rate in the period prior to the failure.

Note that Hypothesis 3 is based on the number of successful phase II compounds in the recent past across all target markets; whereas Hypothesis 2 is concerned with “backup” compounds in all phases but only in the target market of the failed compound.

4. Data Source
We use drug development data from the R&D Insight database developed by ADIS international. It is compiled by a team of scientific editors that track more than 17,000 drugs in active development from over 200 pharmaceutical companies. The primary sources of information are: direct contact with companies, information collected from medical and biomedical journals, attendance at international meetings and conferences, company annual reports, news services, press releases, licensed Lehman Brothers’ Pharma-Pipelines data, and public information from the Food and Drug administration. Drug development is tracked from the earliest laboratory report and continues through world market launch. Every scientific or commercial development advancing the drug’s progress to market is assessed, evaluated, and verified for authenticity before being reported in the database. The database is used by many leading pharmaceutical companies to monitor the competitive landscape. A sample entry for a failed drug from the ADIS database
is provided in §EC.4 of the online supplement (provided in the e-companion).³

We use the ADIS database to identify dates of drug failures, the associated indication, the ownership pattern of the drug, and the portfolio properties (other compounds in development \((n_1, n_2, n_3)\), and the recent success rates) on the failure date. Finally, we look at industry-wide historical data on successes and failures from the ADIS database to estimate the success probability of compounds in each indication \((p_1, p_2, p_3)\).

To verify the data on the firms’ portfolio \((n_1, n_2, n_3)\) properties, we imputed the pipeline for one firm in our data set (Merck & Co), and compared it with information obtained from within the firm. The two portfolios were identical. To verify the authenticity of the failure announcements, we checked a sample of failures with the lead pharmaceutical analyst at a financial firm. Failures were found to be accurate in both date and indication specification. Some of the more prominent failures in our sample received extensive coverage in the popular press. We compare the dates of these failures from the ADIS database with news reports (in the Factiva database) and find the data from ADIS to be accurate. A news report and the associated ADIS database entry is provided in §EC.5 of the e-companion.

³ An electronic companion to this paper is available as part of the online version that can be found at http://mansci.journal.informs.org.

We restrict our attention to phase III failures that originated at publicly traded firms with common stock listed on any U.S. market at the time of the failure. We get stock price data from the CRSP financial database.⁴ We identify the ownership and holding pattern of the originator firm(s) at the time of the failure using the Corporate Affiliations data set maintained by the Lexis-Nexis group. Some descriptive statistics on the firms included in this study are provided in Table 1. The annual sales (averaged over the period of the study) for firms in our data set range from over US$20 billion for the big pharmaceutical firms such as Merck, Pfizer, and Bristol Myers Squibb to just under US$1 billion for biotechnology firms such as Chiron. The median firm in our data set has annual sales of US$13.26 billion, employs 46,560 employees, spends US$1.6 billion annually on R&D (14.10% of sales), and experiences 6.5 phase III failures during the period of our study.

During the time period of the study, 1994–2004, there were 132 phase III failures for publicly traded pharmaceutical firms in our database; they represent our sample. Less than 2% of the events are within a month of other related events, thus assuaging any concerns about clustering.

⁴ The CRSP Database provides access to NYSE, AMEX, and NASDAQ daily and monthly securities prices, as well as to other historical data related to over 20,000 companies. The data is produced and quarterly updated by the Center for Research in Security Prices (CRSP), a financial research center at the Graduate School of Business at the University of Chicago.
5. Methodology and Variables

5.1. Measuring the Impact of Drug Failures
To measure the implications of a late-stage failure, we use an event study methodology (MacKinlay 1997, Kothari and Warner 2007). Event studies have been applied to quantify the impact of a wide variety of firm-specific and economy-wide events. Notable applications from the finance and accounting literature involve measuring the impact of mergers and acquisitions, earnings announcements, issue of new debt or equity, and announcements of macroeconomic variables (trade deficits, unemployment data, interest rates). Notable applications from the strategy literature include studies on the impact of CEO succession, name changes, diversification, takeovers, and competitive entry. In the product development and supply chain management literature, they have been employed to estimate the impact of new product introductions (Chaney et al. 1991), delays in new product introductions (Hendricks and Singhal 1997), supply chain disruptions (Hendricks and Singhal 2005), ISO 9000 certification (Corbett et al. 2005), and of excess inventory (Singhal 2005).

Using the prices of a firm’s tradable securities in financial markets, an event study measures the impact of a specific event on the value of a firm as measured by the price of its common stock. The logic behind this approach is the efficient-market hypothesis: Given rationality and information in the marketplace, the impact of an event should be reflected by the change in the stock price of the firm.

Event study methodologies provide a rigorous foundation to isolate the change in stock price due to the event from the change in stock price due to other factors known prior to the event. A model for the returns given the information prior to the event is first estimated using historical data over the estimation period for each event in the study. This estimated model is then used to predict the expected returns on the day of the event, conditional on no new information or events. To ensure robustness of our findings, we use three alternative return-generating models for predicting the expected returns, the comparison period model (CP), the market model (MM), and the Fama-French three-factor model (FF). Details of the three models are provided in §EC.2.1 of the e-companion. These models give us the expected returns on the day of the failure, taking into account the impact of market and firm specific factors, but in absence of the failure.

The component of the return that cannot be explained by the return-generating models, or the difference between the actual return and the expected return, is attributed to the event—in our case the failure of the phase III clinical trials. This component is commonly referred to as the abnormal return. If no economically relevant information is available, we expect this abnormal return to be zero.

Often, the impact of the event is not limited to the day of the occurrence, but extends a few days before and after the event. This is called the event window. We use multiple event windows, including those suggested by looking at trading volumes using the techniques proposed by Tkac (1999) (detailed in §EC.2.2 of the e-companion). We then aggregate the abnormal return for each day in the respective event windows to obtain our main dependent variable, the cumulative abnormal return or CAR. This variable captures the financial impact of losing a compound while controlling for other factors that influence firm value. This is an empirical measure of the value of each compound to the associated firm, our dependent variable.

For Hypothesis 1, we test the null hypothesis that $\text{CAR}_i = 0$. We report the cross-sectional standard deviation test (the “standard approach” from MacKinlay 1997), the standardized Patell-Z test statistic (Patell 1976), a test that controls for cross-sectional dependence between individual security returns (“crude dependence adjustment test” from Brown and Warner 1980, p. 233), a nonparametric generalized sign-z statistic (Sprent 1989), and the nonparametric Wilcoxon signed rank test. We test this hypothesis for several typical event windows as well as for the event window implied by excessive trading volumes.

To test Hypotheses 2 and 3, we run a linear regression with $\text{CAR}_i$ as the dependent variable, the two explanatory variables, $\Delta TAPIS$ and the phase II buffer, in addition to the control variables. We describe the construction of the two explanatory variables and the control variables in the next three subsections.

5.2. Explanatory Variable: $\Delta TAPIS$
To test Hypothesis 2, we need to construct our explanatory variable, $\Delta TAPIS$ (Equation (2)). $\Delta TAPIS$ is a function of the number of compounds at each stage of development ($n_1$, $n_2$, $n_3$) and the probabilities of success of each compound ($p_1$, $p_2$, $p_3$). $n_1$, $n_2$, and $n_3$ are obtained from the ADIS database as the number of distinct compounds in each of the three stages of trials for the same market as the failed trial....
compound. To estimate the probabilities, we use data on all clinical trials in the ADIS database. A vast majority of these trials are run by firms that are not publicly traded and do not otherwise appear in our sample. Danzon et al. (2005) find that the indication explains the largest fraction of the variance in success probabilities between different drugs. Thus, we estimate \( p_i \) at the level of an indication and assume that all drugs for an indication at the same stage of development have the same odds of failure. For example, to determine \( p_i \) for an Alzheimer’s drug, we look at the performance of all Alzheimer’s drugs irrespective of originating firm. The estimated-indication phasespecific probabilities and the detailed procedure are provided in \( \text{SEC.3} \) of the e-companion. Finally, we use an annual discount rate of 12%, to compute \( \alpha_1 \) and \( \alpha_2 \). A minority of compounds (25 out of 116) in our sample fail for more than one indication at the same time (often due to safety concerns), thus they have more than one \( \Delta TAPIS \) value associated with them. For these compounds, we compute an aggregated \( \Delta TAPIS \) as the sum of the multiple \( \Delta TAPIS \) values and present our results using the same. We also tested our results using the average and the maximum of the multiple \( \Delta TAPIS \) values and found similar results.

5.3. Explanatory Variable: Phase II Buffer

Hypothesis 3 posits that the impact of the phase III failure is proportional to the unanticipated demand for phase III resources at the time of the failure. The unanticipated demand is captured by the difference between the recent and the expected phase II success rate for the firm in question (phase II buffer). To compute the phase II success rates, we divide the number of phase II successes (across all indications) by the total number of phase II trials (sum of the number of successes and failures across all indications) over the relevant time period for the firm in question. For the expected success rate, we look at the number of successes and failures for the firm over all periods of the data set, 1994 to 2004. To compute the recent success rate, we look at the number of successes and failures in the 300-day period preceding the day of the failure announcement. The difference of the two is used as the explanatory variable in our regressions (Equation (3)).

\[
\text{Phase II buffer} = \frac{\text{Recent success rate} - \text{Long-run success rate}}{\text{# of Successes}_{t[300]} - \text{# of Failures}_{t[300]}}
\]

To test the robustness of our results, we also run our regression models with just the recent success rate, the absolute number of recent successes, and the log of the recent success rate minus the log of the long-run success rate. Our results are robust to all these formulations.

5.4. Control Variables

We control for the properties of the compound in question and the firm in question. At the compound level, we include three variables: First, the number of active trials at the time of the failure in the same indication as the focal compound across all firms present in the ADIS database (\( \text{NActiveTrials} \)). Previous research (Nicholson et al. 2005) finds that this variable is highly correlated with the revenue potential of the compound. Second, the number of licensees for the compound in question (\( \text{NLicenses} \)). Depending on the structure of the licensing agreement, this variable is associated with the firm’s financial stake in the compound. Third, the number of originating firms associated with the compound (\( \text{NOriginators} \)). This variable is also related to the firm’s financial stake.

We also include two firm-specific control variables, sales in the quarter of the failure (\( \text{Sales} \)) and R&D expenses incurred by the firm in the quarter of the failure (\( \text{R&D Expenses} \)). These capture the firm-specific heterogeneity, namely the size of the firm and the R&D organization associated with the failure. We estimate the model in Equation (4).

\[
\text{CAR}_i = a_0 + a_1(\text{NActiveTrials}) + a_2(\text{NOriginators}),
+ a_3(\text{NLicenses}) + a_4 \text{Sales}_i,
+ a_5 \text{R&D Expenses}_i + a_6 \Delta TAPIS_i,
+ a_7(\text{Phase II buffer}) + e_i
\]

Hypothesis 2 implies negative and significant estimates for the coefficient \( a_6 \). Hypothesis 3 implies positive and significant estimates for the coefficient \( a_7 \).

Descriptive statistics of the explanatory variables are provided in Table 2. The median failed compound has one originator firm, is targeted at one indication, has one backup compound, and has no licensees. Table 3 shows the Pearson correlation coefficients among the variables.
6. Results and Discussion

6.1. Identification of Event Window: Abnormal Trading Volume

We first estimate a model for the benchmark trading volume for each firm and find the days associated with unexpectedly high trading. The abnormal trading volume data are illustrated graphically in Figure 4. Abnormal trading volume peaks at 816 million shares over the expected volume two days before the announcement of the failure. Abnormal trading volumes in the time period (−2, 4) are found to be statistically different from zero. This implies that most of the information about the event failure is incorporated in the value of the firm during this period. Thus, the abnormal returns in the period (−2, 4) should capture the effects of the event on the firm’s valuation.

To ensure robustness of our results with respect to the choice of the event window, we also conduct all further analyses for alternative event windows ((−3, 3) and (−4, 4)), which are often used in the event study literature.

6.2. Impact of Phase III Failures: The Average Cumulative Abnormal Return

We estimate three benchmark expected return models—the comparison period model (CP), the market model (MM), and the Fama-French three-factor model (FF). These models are estimated individually for each security event in our database using data from the event window (−255, −10). For three data points, fewer than 10 trading days of market data are available in this estimation period. These points are excluded from our sample. The estimated benchmark return models are used to compute the daily and cumulative abnormal returns as described in §5.1.

Table 4 reports the results for the mean and median cumulative abnormal returns. The mean cumulative abnormal returns are negative and significant for a wide variety of model specifications, event windows, and test specifications. These results provide evidence for Hypothesis 1 predicting a negative effect of a drug failure on firm value. Over the time period of the event window, (−2, 4), a phase III drug failure leads to an average loss of 1.46% in the value of the firm (according to the Fama-French model, estimates range from −1.07% to −1.61% using different models and event windows). In dollar terms, these losses correspond to a decrease in the firm value by US$405 million.

Chaney et al. (1991) report in their study of announcements of product successes a cumulative abnormal return of 0.21% using the market model for the pharmaceutical firms in their sample. For an average phase III compound, with an 80% chance of success, the increase in probability on completion of a successful trial is 20% as opposed to an 80% reduction in probability on failure. Thus, we expect the financial impact from the study by Chaney et al. (1991) to be four times smaller. Our results from the market model are consistent with this plausibility check.

Sharma and Lacey (2004) construct in their comparative study of drug development failures and successes a data set of 41 FDA rejections of new drug approval (NDA) applications (the last stage in Figure 1, subsequent to phase III success). An NDA rejection indicates managerial failure in implementing a firm’s internal controls and systems to ascertain drug safety. This is a more serious and rarer event than the medical failure of a drug undergoing phase III clinical trials. Thus, the expected impact of these events should be larger than that of our event. They find abnormal returns as high as 21% associated with the announcement. This corresponds to a financial impact of over US$7 billion, which is much higher than the net present value of the sales of even the

<table>
<thead>
<tr>
<th>Table 2 Independent Variables</th>
</tr>
</thead>
<tbody>
<tr>
<td>Independent variables</td>
</tr>
<tr>
<td>NActiveTrials</td>
</tr>
<tr>
<td>NLicenses</td>
</tr>
<tr>
<td>NOriginators</td>
</tr>
<tr>
<td>Sales (MMS)</td>
</tr>
<tr>
<td>R&amp;D expenses (MMS)</td>
</tr>
<tr>
<td>ΔTAPIS</td>
</tr>
<tr>
<td>Phase II buffer</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Table 3 Pearson Correlation Between Independent Variables</th>
</tr>
</thead>
<tbody>
<tr>
<td>NActiveTrials NLicenses NOriginators Sales (net) R&amp;D expense Phase II buffer ΔTAPIS</td>
</tr>
<tr>
<td>----------------------------------------------------------</td>
</tr>
<tr>
<td>NActiveTrials 1</td>
</tr>
<tr>
<td>NLicenses     −0.18*</td>
</tr>
<tr>
<td>NOriginators  −0.14*</td>
</tr>
<tr>
<td>Sales (net) (MMS) 0.29*** −0.28*** −0.29*** 1</td>
</tr>
<tr>
<td>R&amp;D expense (MMS) 0.2* −0.22** −0.27** 0.61 1</td>
</tr>
<tr>
<td>ΔTAPIS        0.33*** −0.07 −0.08 0.22** −0.08 1</td>
</tr>
<tr>
<td>Phase II buffer 0.12 0.14* −0.08 0.06 0.02 0.04 1</td>
</tr>
</tbody>
</table>

*Significant at the p < 10% level, **significant at the p < 1% level, ***significant at the p < 0.1% level.
biggest blockbuster drugs. This finding suggests that investors may be losing confidence in the firm’s management on account of this kind of rare failure and may be penalizing it for much more than just the lost compound.

The cross-industry study of Hendricks and Singhal (1997) reports that on announcement of product development delays firm values drop by an average of 5.25%, or US$119 million. Chen et al. (2005b) report that on announcement of delays firm values drop by 11.4%.

6.3. Effects of Backup Projects and Recent Success Rate

Table 5 provides results from the OLS estimation of the model in Equation (4). Results are provided for the three return-generating models described in §5.1 and three event windows. We obtain similar results using a WLS regression, with the precision of the estimated abnormal returns as the weights. The $R^2$ for our models ranges from 12% to 19%, which is comparable to other studies of this type (e.g., Chaney et al. 1991). The regressions using the dependent variable created from the market model and the Fama-French model are all significant at the $p < 0.01$ or higher level. The regressions using the comparison period model are significant at the $p < 0.05$ level. Diagnostic tests reveal no problems with heteroskedasticity or multicollinearity. We also examine our estimation procedures for influential observations (Belsley et al. 1980) and find our results to be robust to outliers.

<table>
<thead>
<tr>
<th>Model</th>
<th>Window</th>
<th>Mean cumulative abnormal return (%)</th>
<th>Cross sectional std. dev test</th>
<th>Patell-Z statistic*</th>
<th>Crude dependence adjustment test$^a$</th>
<th>Generalized sign-Z$^a$</th>
<th>Median cumulative abnormal return (%)</th>
<th>Wilcoxon signed rank test</th>
<th>Change in market capitalization (MMS)</th>
<th>Median change in market capitalization (MMS)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Comparison</td>
<td>(−2, 4)</td>
<td>−1.14</td>
<td>−2.18**</td>
<td>−2.20**</td>
<td>−2.15**</td>
<td>−1.64**</td>
<td>−1.12</td>
<td>−1.53*</td>
<td>−143.29</td>
<td>−32.63</td>
</tr>
<tr>
<td>period model</td>
<td>(−3, 3)</td>
<td>−1.19</td>
<td>−2.17**</td>
<td>−2.69***</td>
<td>−2.25**</td>
<td>−0.77</td>
<td>−1.08</td>
<td>−1.59*</td>
<td>−12.91</td>
<td>−16.52</td>
</tr>
<tr>
<td>Market model</td>
<td>(−2, 4)</td>
<td>−1.23</td>
<td>−2.16**</td>
<td>−2.31***</td>
<td>−2.06**</td>
<td>−1.46*</td>
<td>−1.52</td>
<td>−1.90*</td>
<td>−184.17</td>
<td>−40.12</td>
</tr>
<tr>
<td></td>
<td>(−3, 3)</td>
<td>−1.20</td>
<td>−2.27**</td>
<td>−2.76***</td>
<td>−2.41***</td>
<td>−0.53</td>
<td>−0.48</td>
<td>−2.09*</td>
<td>−95.41</td>
<td>−5.07</td>
</tr>
<tr>
<td></td>
<td>(−4, 4)</td>
<td>−1.38</td>
<td>−2.44***</td>
<td>−2.58***</td>
<td>−2.44***</td>
<td>−1.93**</td>
<td>−0.85</td>
<td>−2.81***</td>
<td>−210.28</td>
<td>−36.12</td>
</tr>
<tr>
<td>Fama-French</td>
<td>(−2, 4)</td>
<td>−1.46</td>
<td>−3.17***</td>
<td>−2.90***</td>
<td>−1.79**</td>
<td>−0.90</td>
<td>−3.54***</td>
<td>−404.99</td>
<td>−21.77</td>
<td></td>
</tr>
<tr>
<td>three-factor</td>
<td>(−3, 3)</td>
<td>−1.48</td>
<td>−3.14***</td>
<td>−2.95***</td>
<td>−1.44*</td>
<td>−1.13</td>
<td>−3.89***</td>
<td>−383.86</td>
<td>−51.77</td>
<td></td>
</tr>
<tr>
<td>model</td>
<td>(−4, 4)</td>
<td>−1.61</td>
<td>−3.18***</td>
<td>−2.83***</td>
<td>−1.79**</td>
<td>−1.11</td>
<td>−3.90***</td>
<td>−451.02</td>
<td>−36.94</td>
<td></td>
</tr>
</tbody>
</table>

Note. Significance levels from a one-tail $t$-test: *10% significance, **5% significance, ***1% significance, ****0.1% significance. $N = 132$.

*Unlike the cross-sectional standard deviation test, in computing the Patell-Z statistic, each abnormal return is standardized using the estimated variance of the abnormal return (Patell 1976), standardized tests are not available for the Fama-French Model.

This test uses a single variance estimate for the entire portfolio thereby, avoiding the potential problem of cross-sectional correlation of security returns.

The nonparametric sign-Z test tests the null hypothesis that the number of positive and negative return is the same (Sprent 1989).
We find support for Hypothesis 2. The coefficient for the variable $\Delta TAPIS$ is found to be significant. A 1% difference in the change in TAPIS leads to 1.86 basis points difference in the abnormal return associated with the failure (according to the Fama-French $(-2, 4)$ model; estimates from other models range from 1.86 to 3.02 basis points). In the illustrative example of Figure 2, our regressions suggest that failure 1 would hurt the firm by an extra 1.19%, or US$334.1 million, compared to failure 2. Further, for one standard-deviation difference in the value of $\Delta TAPIS$ (calculated from our sample), the difference in financial impact corresponds to 0.92%, or US$258 million. Support for Hypothesis 2 demonstrates that the presence of other compounds for the same market leads to lower financial impact of a failure or lower valuation of a compound.

We also find support for Hypothesis 3. The coefficient for the variable $\text{phase II buffer}$ is significant. Failures that follow a period where the phase II success rate was higher than the long-run average success rate lead to a less negative impact on the firm value. Conversely, failures that follow periods of below average phase II success hurt firm value more. A 1 percentage point difference in success rate (phase II buffer) leads to a 6.04 basis point difference in the abnormal return associated with the failure. (According to the Fama-French $(-2, 4)$ model, estimates from other models range from 4.2 basis points to 6.5 basis points.)

In the illustrative example of Figure 3, our regressions suggest that failure 2 would hurt the firm by an additional 1.52%, or US$419.5 million as compared to failure 1. Further, for one standard deviation difference in the phase II buffer, the difference in financial impact corresponds to 1.27%, or US$351.7 millions.

Support for Hypothesis 3 demonstrates that presence of additional projects that utilize the same development resources as the failed project mitigates the impact of a failure. Put differently, the value of a compound for a portfolio in which there are many other projects that utilize the same resources is smaller than for a portfolio in which it is the sole claimant to these resources.

Support for this hypothesis also empirically validates the anecdotal phenomena that a failure (an in-licensing opportunity) at a time when the product development pipeline is “congested” or has more compounds than expected hurts (helps) the firm less vis-à-vis a failure (an in-licensing opportunity) at a time when the development pipeline is lean or has fewer compounds than expected (cf. Landers and Lublin 2003 on the impact of failures and a lean pipeline on Merck Pharmaceuticals).

In developing the $\Delta TAPIS$ variable in §3.1, we ignored the effects of compounds for the same market that are present in the pipelines of competing firms. If the competition has late stage compounds in development, we would expect the mitigating effect of the

Table 5: Regression Model: Parameter Estimates

<table>
<thead>
<tr>
<th>Variables</th>
<th>Comparison period model</th>
<th>Market model</th>
<th>Fama-French model</th>
</tr>
</thead>
<tbody>
<tr>
<td>Intercept</td>
<td>(-2, 4) 0.0307 (0.76)</td>
<td>(-2, 4) 0.0186 (0.78)</td>
<td>(-2, 4) 0.0346 (1.45)</td>
</tr>
<tr>
<td>$N_{\text{ActiveTrials}}$</td>
<td>(-1.25) -0.7618 (-0.4597)</td>
<td>(-1.25) -0.9947** (-1.09**</td>
<td>(-1.25) -0.5481 (-0.8145**</td>
</tr>
<tr>
<td>1,000s</td>
<td>(-0.32) -0.0555 (-2.12)</td>
<td>(-0.32) -0.0606 (-0.0004)</td>
<td>(-0.32) -0.68 (-0.257)</td>
</tr>
<tr>
<td>$N_{\text{Originators}}$</td>
<td>(-1.27) -0.17 (-1.96)</td>
<td>(-1.27) 0 (-1.21)</td>
<td>(-1.27) -0.17 (-0.88)</td>
</tr>
<tr>
<td>$N_{\text{Licensees}}$</td>
<td>(-1.17) -0.219 (-0.526)</td>
<td>(-1.17) -0.908** (-0.606)</td>
<td>(-1.17) -0.908** (-0.606)</td>
</tr>
<tr>
<td>1,000,000s</td>
<td>(2.11) (2.24) (1.86)</td>
<td>(2.11) (2.24) (1.86)</td>
<td>(2.11) (2.24) (1.86)</td>
</tr>
<tr>
<td>100,000s</td>
<td>(-2.46) (-2.93) (-2.14)</td>
<td>(-2.46) (-2.93) (-2.14)</td>
<td>(-2.46) (-2.93) (-2.14)</td>
</tr>
<tr>
<td>$\Delta TAPIS$</td>
<td>(-2.61) (-2.05) (-2.12)</td>
<td>(-2.61) (-2.05) (-2.12)</td>
<td>(-2.61) (-2.05) (-2.12)</td>
</tr>
<tr>
<td>F-value</td>
<td>2.12 2.57 3.63 2.63 3.31 3.32 2.56 2.97</td>
<td>2.12 2.57 3.63 2.63 3.31 3.32 2.56 2.97</td>
<td></td>
</tr>
<tr>
<td>Pr &gt; $F$ (%)</td>
<td>4.79 1.75 0.15 1.50 0.31 0.31 1.77 0.70</td>
<td>4.79 1.75 0.15 1.50 0.31 0.31 1.77 0.70</td>
<td></td>
</tr>
</tbody>
</table>

Note. Significance levels from a two-tail $t$-test: *10% significance, **5% significance, ***1% significance.

*Of the 132 failures for publicly traded firms, pipeline data is available only for 116 failures.
firm’s early stage backup compounds to be smaller. To capture this effect, we construct an alternate version of \( \Delta TAPIS \) that quantifies the change in the odds of being the first firm to launch a compound (as opposed to the odds of launching a compound, in the original metric). In our sample, this metric is highly correlated with the original metric suggesting that there is not significant variation in the competitive situation for the failed compounds. Not surprisingly, we find that this metric also has a statistically significant effect on the impact of a failure.\(^{10}\)

In testing Hypothesis 3, we argued that the difference in the phase II success rate measures the excess of work or shortfall of phase III resources. Alternately, this variable may also measure a perception of the firm’s capabilities: A higher than average phase II success rate may indicate high or improving capabilities, and vice versa. To test this alternate interpretation, we construct a variable measuring the difference in the recent phase III success rate versus the long-run phase III success rate, and run our regression with this variable instead of the phase II buffer variable used in the study. This variable is arguably a more direct measure of firm capabilities, but does not measure the work build-up for phase III. We find no statistically significant impact of this variable on the impact of the failure. This discredits the alternate interpretation of the variable used to test the hypothesis.

Taken together, support for Hypotheses 2 and 3 suggests that portfolio-level project interactions significantly alter the value of a project.Ignoring these interactions would lead to estimation errors in a project’s value by an order of millions of dollars when the average values of a project is approximately US$500 million. This could lead to highly suboptimal portfolio and capacity choices. Thus, a product development manager interested in maximizing shareholder returns would benefit from using decision support systems that acknowledge and model these interactions based on our empirical observations.

Although we developed a detailed nonlinear model for Hypothesis 2, relating the number of backup compounds to the financial value of a compound, we did not do so for Hypothesis 3. The financial impact of the congestion effects central to Hypothesis 3 may not be linear in the proxy for utilization.\(^{11}\) An appropriate queuing model that analytically captures these effects remains the subject of future work.

7. Conclusion

The results of our empirical investigation suggest that a late-stage failure of a project is associated with a significant decline in firm value; for an average failure in our data set, this corresponds to a decline in value of US$405 million. We find support for our hypothesis predicting that decline in firm value is mitigated by the presence of backup projects. Put differently, the value of a project within a portfolio that contains multiple projects targeting the same market is smaller than within a portfolio in which it is the sole claimant to the market. We also find support that this impact is mitigated if the firm has an above average phase II success rate prior to the failure leading to a more than expected number of compounds that will utilize the same development resources as the failed project. Put differently, the value of a project within a portfolio containing more projects that utilize the same resources as the failed project is smaller than a project within a portfolio where it is the sole claimant to the resources.

In addition to validating our intuition about portfolio-level projects interactions, support for our hypotheses validates our theoretical metrics for the financial impact of these interactions. Finally, the magnitude of our results suggests that these portfolio-level project interactions significantly alter the value of a project.

Our method offers a data-driven approach for valuation and comparison of in-licensing opportunities available to a pharmaceutical firm. Using the natural experiment of failures, we have built a predictive model of the impact of different compounds on the firm’s valuation, taking into account the portfolio-level project interactions or the fit of the compound in the firm’s portfolio. This model can be used to predict the increase in a firm’s value if a particular compound is added to its portfolio. This should be the maximum fair price that the firm should pay for this compound.

Although the coefficients estimated in this paper are applicable only for late-stage failures in the pharmaceutical industry, the insights, framework, and empirical methodology can be employed more generally for product development portfolios in environments with high uncertainty, which gets resolved in consecutive phases of testing. The development of alternate approaches akin to backup compounds is common in many product development settings. The “winner-takes-all” payoff structure is typical for industries where alternate approaches are investigated to address one user need. The notion of a shared, fixed development capacity and the associated economics are also typical of many R&D environments. Product development environments such as consumer packaged goods with test markets, multiphase defense development contracts, etc. are all amenable to the methods and insights developed in this paper.

Of course, a generalization of our results must be approached with caution. Our methodology rests

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\(^{10}\) We thank an anonymous referee for suggesting this.

\(^{11}\) We thank the anonymous referee for highlighting this.
on the assumption that markets accurately estimate the factors that influence profits from drug development. This is a reasonable assumption for the pharmaceutical industry, which has high investments by sophisticated institutional investors, extensive regulatory and scientific scrutiny, high levels of disclosure, and exogenously defined and publicly measured metrics of product performance. However, this assumption may not apply equally well to all industries. Although the actual impact predicted from the failure may be less accurate when the assumption does not hold, the insights into the relationship between firm value and properties of the product development portfolio should remain applicable as long as there is no systemic irrationality correlated with our product development variables.

This study empirically identifies a direction for developing decision support models for portfolio and capacity choice in risky development environments. Decision support models that realistically represent the portfolio-level project interactions we identify in this paper can be useful for product development managers and could help address the hitherto limited relevance of academic research for product development choices in industrial practice.

8. Electronic Companion
An electronic companion to this paper is available as part of the online version that can be found at http://mansci.journal.informs.org.

Acknowledgments
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References