



THE COMMITTEE ON ENERGY AND COMMERCE

INTERNAL MEMORANDUM

May 8, 2012

To: Energy and Commerce Committee Members

From: Majority Staff

Re: Full Committee Markup

On Wednesday, May 9, 2012, at 4:00 p.m. and Thursday, May 10, 2012, at 10:00 a.m. in room 2123 Rayburn House Office Building, the Committee on Energy and Commerce will meet in open markup session to consider legislation to reauthorize user fee programs for prescription drugs and medical devices, establish user fee programs for generic drugs and biosimilars, and reform Food and Drug Administration (FDA) programs. Opening Statements for the mark up will be held on Wednesday, May 9, 2012 at 4:00 p.m. in 2123 Rayburn House Office Building.

I. BACKGROUND

The Subcommittee on Health approved the Committee Print on May 8, 2012, by a voice vote. The following is a background on the Subcommittee approved legislation.

Title I: Prescription Drug User Fee Act (PDUFA)

The first title would reauthorize the Prescription Drug User Fee Act (PDUFA). Under the PDUFA V agreement, the industry would pay approximately \$713 million in FY 2013 (possibly more based on adjusters) and a higher amount in the remaining four years. As part of the PDUFA V agreement, the FDA would commit to attaining certain performance goals regarding the review of priority and standard drug applications. It also would foster greater interaction between drug sponsors and the FDA and more engagement with patients, including those with rare diseases.

Title II: Medical Device User Fee Act (MDUFA)

The second title would reauthorize the Medical Device User Fee Act (MDUFA). The new MDUFA agreement would provide for \$595 million in user fees for Fiscal Years 2013-2017. The user fee agreement also would include the following improvements: (1) FDA would have to report its total time for reviewing devices; (2) FDA's review process would include greater interaction between sponsors and the agency; and (3) an independent entity would review the device approval and clearance processes, and FDA would have to implement a corrective action plan to address deficiencies.

Title III: Generic Drug User Fee Act (GDUFA)

This title would authorize the new Generic Drug User Fee Act (GDUFA). The proposed generic drug user fee would provide additional resources for the review and regulation of generic drugs. Under GDUFA, the generic drug industry would pay approximately \$1.5 billion over five years. The industry agreed to this fee in return for faster and more predictable review of generic drug applications and increased inspections of drug facilities.

Title IV: Biosimilars User Fee Act (BSUFA)

Title IV contains language that would authorize the new Biosimilars User Fee Act (BSUFA). This user fee would apply to products approved under the abbreviated approval pathway for biological products shown to be biosimilar to an FDA-licensed biological product. BSUFA would authorize the following four types of fees: application, product, establishment and biosimilar product development. The first three would be set equal to the PDUFA rate for each type of fee. The product development fee would be set at 10 percent of the PDUFA application fee.

Title V: Best Pharmaceuticals for Children Act (BPCA) and Pediatric Research Equity Act (PREA)

This title includes language from legislation offered by Mr. Rogers, Ms. Eshoo and Mr. Markey (H.R. 4274) that would permanently authorize the Best Pharmaceuticals for Children Act (BPCA) and the Pediatric Research Equity Act (PREA). BPCA and PREA foster the development of prescription drugs for children and safe use of drugs by children. BPCA was established in 1997. It provides FDA with the authority to grant a six-month marketing exclusivity period to a manufacturer of a drug in return for FDA-requested pediatric use studies and reports.

The Pediatric Research Equity Act (Section 505B of the FDCA) requires a manufacturer of a drug or biologic who submits an application to market a new active ingredient, new indication, new dosage form, new dosing regimen, or new route of administration to also submit a pediatric assessment for that product.

Title VI: FDA Administrative Reforms

Section 601: The section would improve FDA's guidance process by ensuring public participation in the guidance development process and requiring FDA to finalize draft guidance documents.

Section 602: This section would improve FDA's conflict of interest rules so the agency's advisory committees have access to the most knowledgeable experts. Congress established these rules as part of the 2007 reauthorization, and they have led to significant vacancy rates, especially for advisory committees for rare diseases.

Section 603: This section would require electronic submission of drug, generic drug, biologic, and biosimilar applications following issuance of a final guidance by the Secretary.

Section 604: The section would prohibit the FDA from issuing guidance on the regulation of laboratory-developed tests unless it notifies the Committee of its intent to take such action 60 days prior to the issuance of the guidance.

Title VII: Medical Device Regulatory Reforms

Sections 701: This section would return the Investigational Device Exemption (IDE) approval process to the standard authorized by the statute and implementing regulations. IDEs are needed for manufacturers to conduct clinical trials on devices.

Section 702: This section would reaffirm the least burdensome provisions added during the Food and Drug Administration Modernization Act of 1997 (FDAMA).

Section 703: This section would require FDA reviewers to provide the scientific or regulatory rationale for major decisions, and it would establish an expedited appellate process for challenging those decisions.

Section 704: This section would require FDA to provide review decision summaries for those 510(k) clearances involving clinical data.

Section 705: This section would withdraw FDA's recent guidance documents on medical device modifications and prevent FDA from issuing any new guidance on the subject until it completes a report to Congress on the subject. Until FDA issues a new guidance on the subject, the 1997 guidance would be the standard.

Sections 711-712: The section would reaffirm the mandate of the Safe Medical Devices Act of 1990 (and recommendation of the Government Accountability Office (GAO)) that FDA require premarket approval applications from pre-amendment Class III devices or move them to Class II. Second, this legislation would codify GAO recommendations as to FDA's use of its current device recall authority.

Section 721: This section would streamline the de novo classification process, which is used for novel, innovative devices, by striking the requirement that a sponsor receive a Not Substantially Equivalent finding before entering the de novo process, thus making the process more efficient.

Sections 731-732: These sections would allow FDA to enter into agreements with foreign countries on harmonizing inspections and common international labels of medical devices. It also would provide transparency regarding FDA's involvement in international activities.

Sections 741-742: These sections would reauthorize the third party review and third party inspection programs.

Sections 751: This section would extend the exemption on profit for pediatric devices that have been granted Humanitarian Device Exemptions to include certain devices intended for use in adults. The Humanitarian Device Exemption is used to provide access to medical devices with conditions that affect 4,000 individuals or fewer.

Section 761: In 2007, FDA was required to develop a regulation that would allow for a unique identifier on each medical device. FDA has not yet promulgated the regulation. The section would require FDA to issue the regulations not later than 120 days after enactment.

Section 762: This section would extend FDA's "Sentinel" postmarket risk identification and evaluation system to include medical devices. The section would require FDA to engage stakeholders when developing the system for devices.

Section 771: This section would clarify FDA's policy on custom devices for small populations.

Section 772: This section reauthorizes, at the current authorization level, demonstration grants for non-profit consortia to promote pediatric device development.

Section 773: Under this section the Secretary of HHS, in consultation with the FDA Commissioner, the National Coordinator of Health Information Technology, and the Chairman of the Federal Communications Commission, would be charged with developing a report containing (1) recommendations on an appropriate regulatory framework for health information technology that is risk based and (2) a strategy to avoid regulatory duplication.

Title VIII: Drug Regulatory Reforms

Section 801: This section would require FDA to modernize its drug registration and listing so it has accurate and up-to-date information on drug manufacturing facilities.

Section 802: This section would allow FDA to inspect those drug facilities that pose the greatest risk to safety whenever it needs to inspect them.

Section 803: This section would force FDA to build this quality risk management into its good manufacturing practices (GMP) authority.

Section 804: This section would bar the entry of imported drugs from an establishment that is deemed to have delayed, limited or denied an inspection.

Section 805: This section would allow FDA to destroy counterfeit or adulterated imported drug products of minor monetary value.

Section 806: This section would allow FDA to detain, for a reasonable period, drugs found during inspection which the inspecting officer has reason to believe are adulterated or misbranded.

Section 807: This section would provide for 20 years of imprisonment for any person who knowingly holds, sells or dispenses a counterfeit drug.

Section 808: This section would require drug facilities to have a unique number.

Section 809: This section would require certain information from importers so FDA can implement a risk-based approach to import screening.

Section 810: This section would require drug importers to register with the FDA.

Section 811: This section would require a manufacturer to notify the FDA if it knows the use of the drug could lead to serious injury or death, or if the drug is stolen, or if it is counterfeited.

Section 812: This section would allow the FDA to share certain information with trusted foreign countries to better leverage resources.

Section 813: This section would provide for explicit extraterritorial federal jurisdiction over a violation of the Federal Food, Drug and Cosmetic Act so U.S. authorities could hold accountable those violating the Act, like those who caused the heparin deaths.

Section 814: This section would increase penalties for intentional adulteration of drugs that could cause serious adverse health consequences or death.

Section 815: This section would allow the FDA to request inspection documents in lieu of, or in advance of, conducting a physical inspection.

Section 821-823: These sections would streamline and modernize the regulation of medical gas.

Section 831-835: These sections would provide new incentives for the development of antibiotics to address the public health threat of antibiotic resistance. The provision also would require the FDA to issue guidance to improve the clinical development of antibiotic drugs.

Section 841-843: These sections would improve access to the Accelerated Approval pathway, including for drugs that treat rare diseases.

Section 851: The section would reauthorize the Critical Path Public-Private Partnerships program.

Section 861: This section would reauthorize the enantiomer exclusivity authorized in 2007. An enantiomer is one of a pair of compounds that have a mirror image relationship.

Section 862: Under current law, if a generic drug manufacturer challenges a brand manufacturer's patent as being either invalid or will not be infringed, it can be awarded 180 days of exclusivity from other generic competition. If the generic company does not market the drug within 30 months of filing its application, then it forfeits the 180 day exclusivity period. The current average time for FDA to approve a generic drug application is 31 months. The provision would temporarily increase the time period for a generic drug that is awarded the 180 day exclusivity period to market its drug to 45 months. This period would be phased back down to 30 months as the FDA eliminates the backlog of pending generic applications pursuant to the Generic Drug User Fee goals.

Section 863: Current law allows the FDA to deny a generic application if the drug the generic application is referencing was withdrawn for safety or effectiveness reasons. Current FDA

regulations allow a company to petition the FDA to determine if the reference drug was withdrawn for safety or effectiveness reasons. However, there is no timeframe under which the FDA must respond to the petition, and there have been instances of decision times lasting over two years. This section would require the FDA to respond to these petitions within 270 days, which should result in the quicker approval of generic drugs.

Section 864: Certain citizen petitions ask the agency for a stay of action related to FDA approval of a pending application submitted under section 505(b)(2) (a new drug application that relies on safety and efficacy data produced by someone else) or 505(j) (generic application) of the Act due to scientific or medical questions about the application. The FDA is required to take final agency action on the petition within 180 days. This section would require the FDA to take final agency action on the petition within 150 days.

Section 865: This section would create a demonstration project that provides priority review vouchers to companies that develop a drug for a pediatric rare disease. The voucher would be redeemed by the company for a subsequent application or could be transferred to another company. The section would require the GAO to conduct a study for Congress on the effectiveness of the program. The study is required to be completed one year after the third priority review voucher is issued. The Secretary could no longer issue new priority review vouchers one year after the third voucher is issued so Congress may determine whether to continue the incentive.

Section 866: This section would task the Secretary of HHS with reviewing and reporting to Congress on current Federal initiatives and identifying gaps and opportunities with respect to ensuring the safe use of prescription drugs with the potential for abuse.

Section 867: This section would make the Risk Evaluation and Mitigation Strategy (REMS) system more efficient by allowing for minor modifications of a drug's REMS. Certain drugs are required to have REMS to ensure that the benefit of the drug outweighs the risk. Currently, the law is unclear as to whether sponsors can make minor modifications to the REMS without the need to have a full reassessment of the REMS.

Section 868: This section would require FDA to ensure that opportunities exist for consultation with external experts on products for rare diseases.

Section 869: This section would expedite the development and review of a drug designated a "breakthrough therapy" through increased interaction with the agency during the development process. To achieve this designation, a drug must be intended to treat a serious or life-threatening disease or condition, and preliminary clinical evidence must indicate that it may demonstrate substantial improvement over existing therapies.

Section 870: This section would reauthorize FDA's orphan drug grant program at the current level for five years. This grant program helps facilitate the development of orphan disease products.

Title IX: Drug Shortages

Section 901: This section would modify existing reporting requirements for manufacturers of drugs that are life-supporting, life-sustaining, and intended for use in the prevention or treatment of a debilitating disease or condition. This will ensure better information flow to the FDA and help the agency to better marshal its resources to prevent anticipated drug shortages and mitigate existing shortages.

Section 902: The section would require the FDA to maintain a drug shortage list and provide patients, providers and the public with such information in order to prevent, mitigate, and manage drug shortages on the ground. The bill includes safeguards to prevent the release of confidential business information or information that could adversely affect public health.

Section 903: The section would require the Drug Enforcement Administration (DEA) to provide timely approvals or denials of increases in quotas of controlled substances in instances where such an increase could help address a drug shortage.

Section 904: This section would require the FDA to expedite the approval of manufacturing changes that could help prevent or mitigate a drug shortage while ensuring good manufacturing practices continue to be used.

Section 905: The section would authorize the GAO to conduct a study to examine the causes of drug shortages and issue recommendations on how to prevent or alleviate a drug shortage. This provision will provide needed data on how the regulatory framework, manufacturing challenges, or other factors contribute to drug shortages, as well as recommendations to address such issues.

Section 906-907: These sections would require the FDA and DEA to report annually on their efforts on drugs shortages based on the metrics set forth by Congress.

Section 908: The section develops interim rules to allow repackaging of drugs into smaller units by hospitals within the same hospital system for drugs that are in shortage. The interim rules will be replaced once the FDA issues guidance allowing these hospitals to continue this practice.

II. CONCLUSION

Should you have any questions regarding the hearing, please contact Clay Alspach or Ryan Long at (202) 225-2927.