



RESEARCH PAPER

An overview of intellectual property within agricultural biotechnology in Brazil



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Abstract Brazilian agricultural biotechnology has seen great advances in recent decades, especially in the development of GM crops, including soybean, cotton, and maize, which has placed Brazil in second place since 2013 in the ranking of countries with the greatest GM-cultivated area. However, patenting these technologies is somewhat more restrictive in Brazil than in other countries, such as the USA and Japan, especially concerning isolated biological material from nature. Hence, the intellectual protection of crops in Brazil is encompassed by *sui generis* rights and/or the patenting of only the development process. Given the current scenario and the importance of biotechnology for the Brazilian agriculture sector, it is necessary to deeply study the patent system for recently developed technologies to identify opportunities for enterprises and national institutes to act in this area. The application of novel biotechnological strategies to agriculture will contribute to the expanding agriculture sector and become part of the solution to global challenges. Through this study, we can identify the major companies developing and protecting their agrobiotechnologies. Additionally, a more detailed analysis verifies

Abbreviations: BGMV, bean golden mosaic virus; CMV, cucumber mosaic virus; CNBS, National Biosafety Council; CRISPR/Cas9, clustered regularly interspaced short palindromic repeats-associated protein-9 nuclease; CTNBio, National Biosafety Technical Commission; CTP, chloroplast transit peptides; DII, Derwent Innovation Index; DNA, deoxyribonucleic acid; EMBRAPA, Brazilian Agricultural Research Corporation; EMBRAPAII, Brazilian Agency for Industrial Research and Innovation; FAO, Food and Agriculture Organization of the United Nations; FAPESP, São Paulo Research Foundation; GM, genetically modified; INPI, National Institute of Industrial Property; IPC, International Patent Classification; ISAAA, International Service for the Acquisition of Agri-Biotech Applications; MAPA, Brazilian Ministry of Agriculture Livestock and Food Supply; OECD, Organisation for Economic Co-operation and Development; PLRV, potato leafroll virus; PRSV, papaya ringspot virus; PVY, potato virus Y; RNAi, RNA interference; TALEN, transcription activator-like effector nuclease; UFRGS, Federal University of Rio Grande do Sul; UnB, University of Brasília; UNICAMP, State University of Campinas; UPOV, International Union for the Protection of New Varieties of Plants; USA, United States of America; USP, University of São Paulo; WIPO, World Intellectual Property Organization; WMV, watermelon mosaic virus; ZFN, zinc-finger nuclease.

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that although there are some restrictions in Brazilian laws, GM patent applicants find ways to obtain intellectual protection for the tools they use in the development of GM crops, which include regulatory sequences, gene constructs and production methodologies. Mechanisms to stimulate investment in Brazilian research companies and public policies must be consolidated, allowing investment and public-private partnerships in this sector, with the aim of applying biotechnological knowledge and turn it into products demanded by society.

Introduction

According to data from FAO (United Nations, 2017), in 2050, the world population will be 29% greater than today, and 70% will be urban. The increasing demand for food in emerging countries, the volatility of food prices, climate change, soil degradation and water shortages will be among the major challenges to fighting hunger, which affects 795 million people worldwide, including 780 million in developing regions (FAO, 2015). These issues confirm the importance of investing in technological alternatives in the agriculture sector that can be tailored to local conditions, aiming to increase the amount of food and to exploit conditions to promote inclusive growth (FAO, 2015).

One of these alternatives continues to be the production of GM plants with traits that confer higher productivity and better alimentary and crop conditions allied with suitable soil exploitation. The first GM crop developed was a tobacco (*Nicotiana tabacum*) variety produced by Monsanto scientists 35 years ago that conferred resistance to aminoglycoside antibiotics (Fraley et al., 1983). Recently, EMBRAPA developed a common bean variety (event 5.1) resistant to BGMV, which shows the impact of this technology in Brazil. Despite the importance of virus-resistant GM plants, most transgenic-growing fields also develop herbicide-tolerant and insect-resistant plants. In this way, different strategies have been employed to genetically modify plants and obtain varieties that are resistant to insect pests, most often by transforming the hosts' genes for protein expression that interferes in the life cycle of insects or that is lethal to them, such as enzyme inhibitors and lectin genes. Cry toxins from *Bacillus thuringiensis* have also been expressed in transgenic cultures and substantially contributed to the efficient control of insect pests, dramatically reducing the use of chemical pesticides (Bravo, Likitvivatanavong, Gill, & Soberon, 2011).

According to the ISAAA, from 1996 to 2016, there was an accumulation of 2 billion hectares dedicated to GM crop production for the market (ISAAA, 2016). The main GM-growing countries in 2016 were the USA (72.9 million hectares), Brazil (49.1 million hectares), Argentina (23.8 million hectares), Canada (11.6 million hectares) and India (10.8 million hectares) (ISAAA, 2016). In Brazil, there was an increase of 11% in the area used for transgenic crop cultivation from 2015 to 2016, of which soybean, maize and cotton represent the majority of crops, corresponding to 93.4% of the total area growing GM plants (ISAAA, 2016). ISAAA also demonstrated in 2016 that, for soybeans, growers obtained adherence of 96.5% for GM varieties. For maize and cotton, adherence was 88.5% and 79%, respectively. Regarding the area increase relative to each crop, from 2015 to 2016, soybean, maize and cotton saw 7.5%, 16.1% and 6.3% increases,

respectively (ISAAA, 2016). Hence, the adhesion of GM technology and the cultivation area increased simultaneously. This, however, should be prevented because there is a limit to the area available for cultivation, focusing on environmental preservation.

To couple the adhesion to GM crops to economic development, there are 2 intellectual protection mechanisms used in Brazil: the patent system (Brasil, 1996) and the *sui generis* rights of cultivars (Brasil, 1997). Once protected and released to the market, these plants ensure considerable profits for their technology owners though the payment of royalties.

Regarding the first mechanism, the Law of Industrial Property (N. 9279 from May 14th 1996) specifies that "the whole or a part of natural living beings and biological materials, even when their genome or germplasm are isolated, as well as natural biological processes" are not considered inventions. It is important to remark that GM plants and GM animals are considered inventions in Brazil, but they aren't patentable according Article 18 (III according the Law N. 9279/96). The only living beings that can be considered patentable inventions under Brazilian law are transgenic microorganisms, which can be patented if they meet the two conditions of patentability (clarity and descriptive sufficiency) and the 3 patentability requirements (novelty, inventive step and industrial application).

In addition, processes involving living organisms (e.g., methods for the development of transgenic plants), gene constructs (e.g., expression vectors and cassettes), recombinant proteins and compositions of biological extracts are also patentable according to the Brazilian law. This means that in the case of GM plants, the tools and methods for their development can be protected, but not the plants themselves. In this case, it is possible to resort to the second mechanism, known as crop protection, based on *sui generis* rights and ruled by Law N. 9456/97 (Brasil, 1997), which grants owners crop protection for 15 years, except for vines, fruit trees, forest species and ornamental plants, whose protection period is 18 years. Accordingly, considering the impact of GM plants on the Brazilian agriculture sector in the worldwide context, in this work, we provide an outlook on intellectual property in agricultural biotechnology from 2010 to 2016.

Methods

To analyse the situation for patented agrobiotechnologies in Brazil, we first reported the main transgenic events approved by CTNBio based on scientific papers and reports. Next, we searched the database and reports of MAPA, in 2017, for protected and registered crops. We also performed

an analysis of patent requests (biotechnology area) from 2000 to 2015 in the WIPO database (WIPO, 2017a) based on the following IPC codes (WIPO, 2017b): C07G, C07K, C12M, C12P, C12Q, C12R and C12S. In this case, two search strategies were performed: (a) patents applied in Brazil and (b) patents of Brazilian origin. A comparative analysis was also performed analysing different countries. In this way, we used strategy “(b)” selecting the options for “Office” as Brazil/USA/India/Argentina/China/Canada. Finally, we analysed the patent protection system for agrobiotechnologies in the Derwent Innovation Index (DII) database using search strategy based on the International Patent Classification (IPC).

Aiming to provide an overview of patent applications in agricultural biotechnology in Brazil, we created a search strategy based on the IPC used by the OECD from 2010 to 2016, focused on the agriculture sector and restricted to Brazilian documents. Hence, the query sentence used was as follows: IP=(A01H-001/00 OR A01H-004/00 OR A61K-038/00 OR A61K-039/00 OR A61K-048/00 OR C02F-003/34 OR C07G-011/00 OR C07G-013/00 OR C07G-015/00 OR C07K-004/00 OR C07K-014/00 OR C07K-016/00 OR C07K-017/00 OR C07K-019/00 OR C12M* OR C12N* OR C12Q* OR C12S* OR C12P* OR G01N-027/327 OR G01N-033/53 OR G01N-033/54 OR G01N-033/55 OR G01N-033/57 OR G01N-033/68 OR G01N-033/74 OR G01N-033/76 OR G01N-033/78 OR G01N-033/88 OR G01N-033/92) AND PN=BR*. Data retrieved from DII were analysed using Vantage Point software (Georgia Tech/Search Technology Inc., Atlanta, USA).

Results and discussion

Approved GM plants for the Brazilian market

Following the approval of the Law of Crop Protection in Brazil (N. 9456/97), the first GM crop event was released in the Brazilian market in 1998 – the soybean variety Roundup Ready containing the *cp4epsps* gene that conferred tolerance to glyphosate (Brasil, 1998). Thus, it became critical to elucidate the complex intellectual property context for glyphosate-tolerant soybean cultivars and protection strategies in agricultural biotechnology to promote the legal safety of those involved in these technologies, since for the farmers it is hard to correlate sometimes the royalties payment with the intellectual property enterprises right (Rodrigues, Lage, & Vasconcellos, 2011). The Law of Biosafety (N. 11105) was thus created in 2005 (Brasil, 2005a) and provided safety rules and surveillance mechanisms for activities involving GM organisms and their derivatives; it also instituted the CNBS and restructured CTNBio.

In 2005, the first commercial GM cotton (Bolgard I – Monsanto) was also approved, containing the *cry1ac* gene that conferred insect resistance (Brasil, 2005b). In 2007, Monsanto, Bayer and Syngenta companies developed and received approval for their first commercial transgenic maize events in Brazil: MON810 (containing the *cry1ab* gene for insect resistance), Libert Link (containing the *pat* gene for herbicide tolerance) and TL (containing both *cry1ab* and *pat* genes), respectively. Following this, many other GM events were released to the Brazilian market, suggesting that the referred laws were definitive in paving the way

for these processes. In addition, the transgenic common bean event 5.1 from EMBRAPA and the higher-productivity eucalyptus from FuturaGene were also approved by CTNBio (Brasil, 2011, 2018a).

In 2016, CTNBio approved 18 GM organisms for the market, of which 9 were GM plants. According to the commission, there are currently 66 transgenic plant events commercially approved in Brazil, of which 40 are maize events, 13 cotton events, 11 soybean events, 1 common bean event and 1 eucalyptus event (CTNBio, 2016; 2017). CTNBio reports (CTNBio, 2016; 2017) show increasing interest from multinational companies in the development of GM plants characterised by insect resistance and herbicide tolerance, especially using gene pyramiding.

Despite a growing number of Brazilian published patents in the general area of biotechnology since the Law of Industrial Property (1996), we notice that Brazil is far behind the USA and China (Table 1), which reveals an urgent need to invest much more in innovation, including patent protection, to help Brazil be more competitive in the biotechnology sector.

Brazilian protected GM crops

According to internal reports from MAPA and an analysis of its protected crop database (MAPA, 2017) there were 2318 protected crops in Brazil, of which 628 had a definitive protection certification: 593 soybean crops, 34 cotton crops and 1 common bean crop (BRS FC401 RMD from EMBRAPA). By June 2017, there were 2214 GM crops registered in MAPA, which means they are ruled for commercial purposes. Of these crops, 51% represent soybean, 45% maize, 3.7% cotton, 0.15% common bean and 0.15% eucalyptus.

Data from Fig. 1 show the tendency towards an increase in the number of registered GM crops in Brazil – especially soybean – and highlight the impact of this technology on Brazilian agribusiness. The decrease shown in Fig. 1 for the more recent years is probably due to a delay in indexing data. In the case of maize, although there are a reasonable number of GM crops, owners usually prefer to protect their technology through secret industrial systems because this species provides hybrid crops.

Agricultural Biotechnology: analysis of Brazilian patent protection

OECD data for 2015 identifies the USA, EU members and Japan as the main countries with patent applications in the biotechnology area for the period from 2010 to 2013 (OECD, 2015).

An analysis of published patents in the biotechnology area in Brazil (Fig. 2) shows that despite the large number of patent applications, most of the patenting effort is of foreign origin, which are simply represented by “Brazilian documents”, in contrast to “documents of Brazilian origin”, since Brazilian documents include patents of foreign origin but applied in Brazil, whereas documents of Brazilian origin refer to patents that have first application in Brazil and are usually done by Brazilian. This demonstrates the interest of several international actors in protecting their inventions in the Brazilian market and indicates that given

Table 1 Number of patent documents in biotechnology for the main GM plant-producing countries (2000–2015).

| Country | 2000 | 2001 | 2002 | 2003 | 2004 | 2005 | 2006 | 2007 | 2008 | 2009 | 2010 | 2011 | 2012 | 2013 | 2014 | 2015 |
|--------------------------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|
| Argentina | 6 | 6 | 16 | 20 | 24 | 18 | 20 | 25 | 22 | 18 | 16 | 26 | 19 | 21 | 36 | 18 |
| Brazil | 30 | 42 | 72 | 49 | 87 | 70 | 79 | 75 | 107 | 128 | 110 | 114 | 97 | 155 | 165 | 168 |
| Canada | 727 | 777 | 1019 | 1065 | 1047 | 916 | 711 | 709 | 692 | 736 | 703 | 748 | 644 | 629 | 706 | 453 |
| China | 424 | 2563 | 2169 | 1291 | 1508 | 2266 | 2433 | 2746 | 3835 | 4511 | 5573 | 7211 | 8840 | 10,091 | 11,695 | 6374 |
| India | 48 | 65 | 103 | 172 | 185 | 203 | 224 | 317 | 298 | 231 | 249 | 276 | 298 | 252 | 271 | 318 |
| United States of America | 11,011 | 13,134 | 17,486 | 18,578 | 15,894 | 15,308 | 13,350 | 13,564 | 13,845 | 13,929 | 13,732 | 14,598 | 14,190 | 14,957 | 17,139 | 12,905 |

Source: Created by the authors based on WIPO data (WIPO, 2017a).

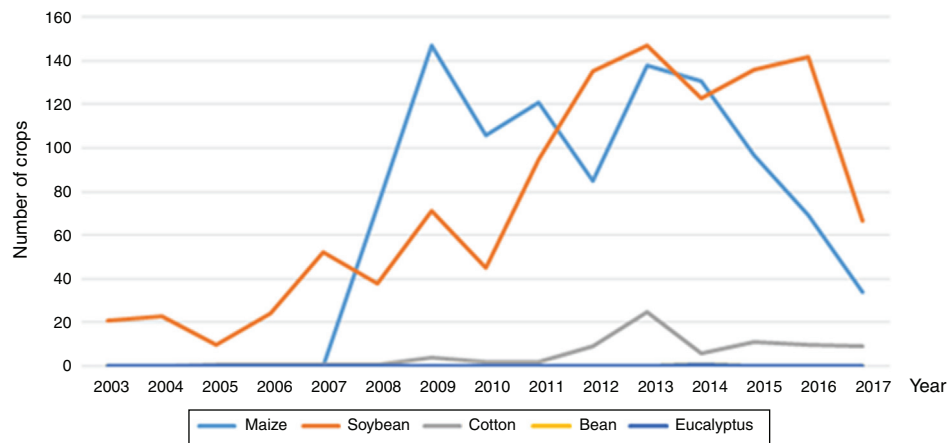


Figure 1 Evolution of the number of GM crops registered in MAPA.

Source: Created by the authors based on MAPA data (WIPO, 2017a).

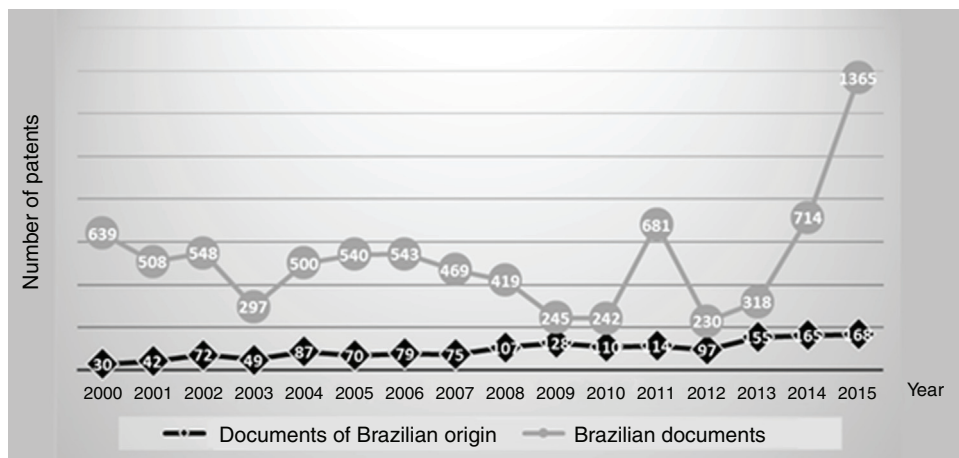


Figure 2 Number of published patents of Biotechnology from 2000 to 2015 in Brazil.

Source: Created by the authors based on WIPO data (WIPO, 2017a).

the important knowledge base built over the years in the field of biotechnology in Brazil, national inventors could demonstrate more significant participation in patenting in this technological field.

A study performed by INPI concerning applied biotechnology patents in Brazil by Brazilian owners from 2009 to 2013 concluded that most of the applications were from companies and universities and were related to the agroindustrial sector, especially those involving GM plants (Verde, Dos Santos, & Guerrante, 2015). INPI also showed UNICAMP as the ranking's leading applicant, with USP and EMBRAPA also being important.

Using the above-mentioned strategy to analyse Brazilian documents representing patent applications in the biotechnology sector from 2010 to 2016, 3711 results were retrieved, of which 916 were retained after refining the search to agricultural biotechnology. Of these, 47% were related to plants, 32% to animals and 15% to microorganisms. Those related to microorganisms mostly involve biofuel production processes using algae but also involve molecule biosynthesis, cultivation methods, fermentation processes, vaccines and cosmetics composition, probiotics and compositions for the biological control of plant pests. Those

documents for animals mostly involve vaccines and methods for treating human and animal diseases but also involve GM animals, diets, disease detection methods, synthetic genes for veterinary applications, methods for animal reproduction (e.g., artificial ovaries, sperm enrichment), genotyping chips, molecular markers, cryopreservation, among others. In the case of plants, most (approximately 93%) involve the development of GM plants but also involve culture media, bioreactors, gene mapping, seed analysis, fertilizers, embryo analysis, hybrid production, molecular markers, and grafting.

An analysis of priority countries revealed that most of the Brazilian patent documents from 2010 to 2016 were first applied in Canada (523), the USA (511) and South Korea (260), which strongly suggests that these countries are the main research development sites resulting in patent applications in Brazil. Brazil was the fourth in this ranking (203). The results showed that Australia, China, Japan and the Philippines are also interested in the Brazilian market, possibly due to the extensive area used for agriculture in the country and to well-established GM cultivation. The USA, EU countries and China are other targets of market interest because the USA and China are worldwide leaders

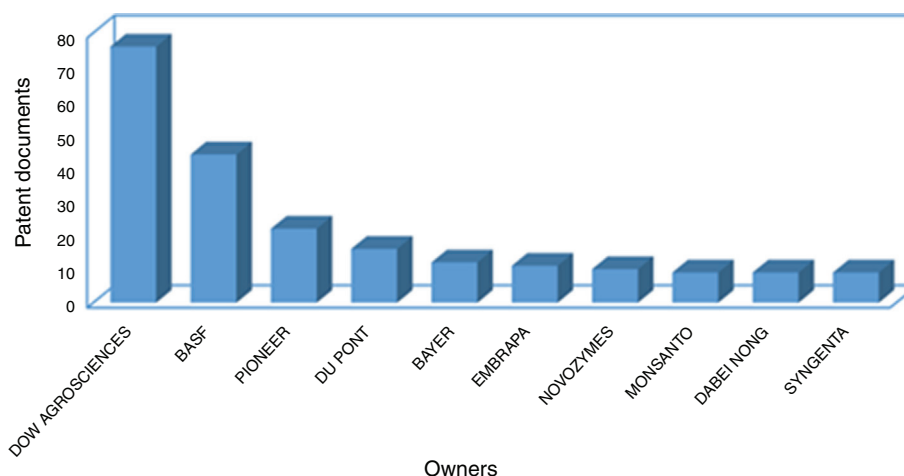


Figure 3 Ranking (Top 10 – legal persons) of the main owners of Brazilian patents involving GM plants (2010–2016).

in most technologies, and EU countries have powerful biotechnology research centres.

Examining the major patent applicants in Brazil, Dow AgroSciences, BASF and DuPont Pioneer were the main patent owners in agrobiotechnologies, and in the national context, EMBRAPA was the main applicant, with 17 published patent documents, representing 6th place in the ranking, followed by the Danish company NOVOZYMES (Fig. 3).

We must also emphasise the acquisition of companies over the years. In 2017, for instance, Dow AgroSciences was merged with DuPont Pioneer (Globo, 2017), Monsanto was bought by Bayer (Valor Econômico, 2017a) and Syngenta was sold to ChemChina (Valor Econômico, 2017b). These three large companies established in the agriculture sector control more than 60% of the worldwide seed and agrochemical markets (Valor Econômico, 2017c).

Considering the 30 main applicants in agricultural biotechnology, EMBRAPA stands out as the most linked with different institutions through partnerships, either Brazilian (e.g., UnB, FAPESP and UFRGS) or foreign (e.g., with BASF). BASF also has a striking link with natural persons, probably because individual inventors are considered owners in the USA. Abbott and Abbvie were also found to be linked, likely because the pharmaceutical company Abbott spun off Abbvie. Danisco was linked to DuPont Pioneer because it was incorporated into DuPont in 2011 (Supplementary Figure 1).

Kenneth E. Narva from Dow AgroSciences is the inventor who figures in the largest number of patent applications of this study; these usually involve insect (Cry 1F, RPA70, RPS6, Cry1Ab, Cry2Aa, Cry1C, Cry1Da, Cry1Be, Cry1Fa, Cry2Aa, Cry1I, Cry1E, Cry-1Da and Cry1 Ca proteins) and nematode (Cry14, Cry 5 and Cry 6 proteins) control.

Analysing these 916 agricultural biotechnology patent documents according to the IPC (WIPO, 2013), we found that most of them were classified as C12N 15/82 (vectors or expression systems especially adapted to plant cells) or A01H 5/00 (angiosperms), confirming results indicating that patent applications mostly involve GM plants. This shows despite the restrictions in industrial property legislation regarding the protection of plants by the patent system, many companies and institutions are interested in protecting related biotechnological tools necessary for the development of GM plants in Brazil.

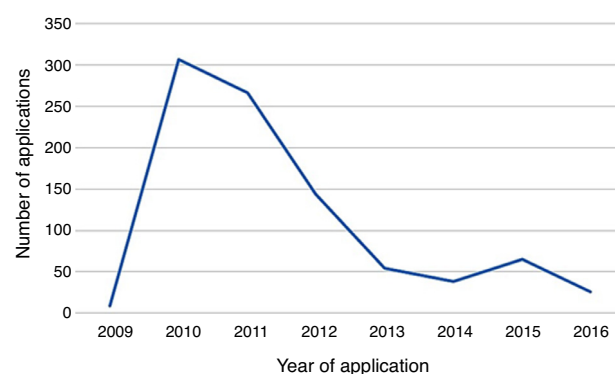


Figure 4 Evolution of Brazilian patent requests per year in Agricultural Biotechnology (2010–2016).

Analysing the evolution of patent applications in Brazil in the field of agricultural biotechnology (Fig. 4), a progressive decline was shown, with some interfering factors: patents still under the secrecy period; a delay in database updates; or disinterest in searching for protection because of, among other factors, (1) the restrictive scope of patent protection from Brazilian law in the biotechnology field or (2) the very slow rate of patent analysis in Brazil. When focusing on requested patents for GM plants in the same period (Fig. 5), it was found that they follow almost exactly the same pattern as that for agrotechnologies in general (Fig. 4).

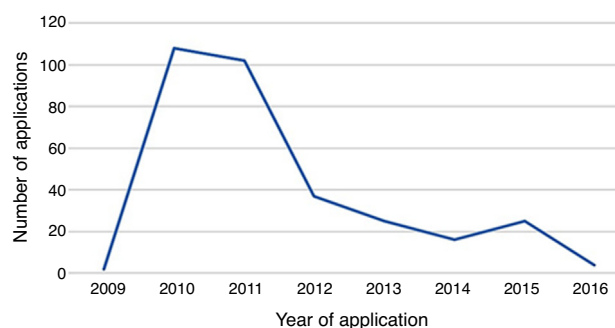


Figure 5 Evolution of Brazilian patent requests per year involving GM plants (2010–2016).

Patenting process for GM crops in Brazil

Despite restrictions in the laws concerning the patent protection of plants, interest clearly remains in protecting biotechnological tools useful for the development of GM crops in Brazil, as 318 out of the 916 refined documents (35%) comprised the development of GM crops. A more detailed analysis places Brazil in the 5th place of applicant countries, with the USA, Canada and South Korea being the main applicants, indicating other countries' growing interest in the Brazilian market.

Regarding the type of protected GM crops, most are related to genes or proteins protecting against insecticidal activity (especially against Lepidoptera, Coleoptera and Hemiptera) and providing tolerance to herbicides (e.g., 2,4-D and glyphosate). In the first case, the most cited target insect-pest species are *Ostrinia nubialis* (Lepidoptera), *Spodoptera frugiperda* (Lepidoptera), *Diabrotica speciosa* (Coleoptera), *Pseudoplusia includens* (Lepidoptera), *Anticarsia gemmatalis* (Lepidoptera) and *Euschistus heros* (Hemiptera), and their control is based on the insertion of genes coding for Cry toxins.

Table 2 lists patent requests from the main owners related to insect resistance. Although requests referring to Dabeji Nong relate to Chinese insect pests, these patent applications in Brazil may aim to protect a possibly useful technology if this pest spreads to Brazil. This table also indicates different protection strategies used by owners: gene stacking technologies, transgenic events containing stacked genes and methods for developing insect-resistant GM plants (e.g., through RNAi).

The documentation for the listed patent requests also describes other GM-related technologies such as plant transformation methods, GM organism detection kits, site-directed integration methods such as gene editing tools (e.g., ZFNs and TALENs), function restoring genes for plant male sterility, target loci detection methods for foreign DNA, methods of introgression for transgenic events, plant regulatory sequences (e.g., promoters), devices and GM microorganisms used in plant transformation, GM plants with oil production improvement, CTPs, RNAi strategies, fusion proteins, proteins for nematode control and genes for yield improvement (e.g., increase in biomass and/or seed productivity). Although the use of other genome engineering technologies such as CRISPR/Cas9 is on the rise, few documents based on these were found, perhaps because they were developed recently, and many patent documents may be still within the secrecy period.

Analysing the patent requests related to GM crops made it possible to observe that owners try to include not only the central invention (usually a gene or a protein) but also any surrounding technologies with possible commercial importance, such as the GM plant itself or its parts, gene constructs or cassettes, transgenic-derived products, methods involving invented molecules and even fields containing these organisms. Otherwise, most of these items will have their protection denied in Brazil by the law, although INPI holds that protection of items containing natural biological products must not be considered solely dilutions of non-patentable technologies (INPI, 2015a).

It was also observed that approximately 22% of the identified patent applicants have a "filed" status in INPI most

of the time due to the lack of payment of annuities. This status may have a number of causes: (1) temporary problems in the INPI's processing of the fees paid due to changes in their value and the need for proof by the applicant, (2) withdrawal by owners or (3) management trouble for active patents. The last cause is less likely because owners have a long protection history. In the cases of withdrawal, a patent examination resulting in the absence of patenting criteria for a technology that was previously experienced abroad may be a reason. This explanation corresponds with the observation that most of the documents are of foreign origin – especially being from the USA – often first applied in the USA, and thus able to retrieve examination results more quickly. Likewise, some patents were requested almost 10 years ago, and none had yet been deferred yet or had technical requirements, confirming the long-lasting patent examination process in Brazil, although efforts are being made in INPI in order to accelerate this process, which can overcome this obstacle and result in a very different scenario in the future.

Conclusion

There is too high an interest among multinational companies in obtaining patent protection for their biotechnological products in Brazil, especially as concerns GM plants. This interest is likely due to (1) Brazil's status as an agricultural power; (2) the progressive adhesion to GM crops in the last decade; (3) the role of GM plants in supporting the development of agriculture; and (4) the capacity of GM plants to reduce environmental damage by diminishing the use of soil and decreasing the use of chemicals such as insecticides.

However, national companies are not able to compete with multinational ones, with the exception of EMBRAPA, which was 6th place in terms of patent applications for agrobiotechnologies. Thus, Brazil is not yet strongly competitive in this area, particularly compared with biotechnology-leadership countries such as the USA and China.

Additionally, we notice that broad protection is the main strategy adopted by large companies in the development of GM plants. This strategy comprises the protection of invented genes/proteins and the derivatives of these inventions, such as gene constructs/cassettes, production methods, parts of GM plants, GM plant-derived products and seed mixtures. Moreover, many companies have protected agrobiotechnologies comprising stacked genes within the same technology, either related to the same trait (e.g., genes for insect resistance) or to distinct traits (e.g., genes for insect resistance and genes for herbicide tolerance). In the case of Brazil, despite more restrictive laws, broad protection may set the expectation that the law could be modified when documents are examined in the future.

Current status and perspectives

We must rethink the need for national investment in projects for biotechnological innovation and consolidate already existing initiatives, such as the National Policies for Biotechnology (Brasil, 2007), the legal mark of science technology and innovation (Law N° 13.243/2016 and Law Decree N°

Table 2 Brazilian patent requests related to insect-resistant GM plants (2010–2016).

| Item | Patent application number | Event/Gene/Product/Process | Target insects | Owner |
|------|---------------------------|--|-----------------------------------|---------------------|
| 1 | BR112012014772 | Vip3ab and Cry1fa proteins | Lepidoptera | Dow AgroSciences |
| 2 | BR112012015005 | Cry1ca-modified proteins | Lepidoptera | Dow AgroSciences |
| 3 | BR112012014700 | Cry1ab and Cry2aa proteins | Lepidoptera | Dow AgroSciences |
| 4 | BR112012014727 | Cry1da and Cry1be proteins | Lepidoptera | Dow AgroSciences |
| 5 | BR112012014702 | Protein stackings: Cry 1Fa-Cry2Aa and Cry 1I-Cry 1E | Lepidoptera | Dow AgroSciences |
| 6 | BR112012014803 | Vip3ab and Cry1ca proteins | Lepidoptera | Dow AgroSciences |
| 7 | BR112012014879 | Cry1be and Cry1f proteins | Lepidoptera | Dow AgroSciences |
| 8 | BR112012014801 | Cry1ca and Cry1fa proteins | Lepidoptera | Dow AgroSciences |
| 9 | BR112012014796 | Cry1ab and Cry1be proteins | Lepidoptera | Dow AgroSciences |
| 10 | BR112012014681 | Cry1da and Cry1ca proteins | Lepidoptera | Dow AgroSciences |
| 11 | BR102012019434 | Insect-resistant soybean event 9582.814.19.1 (CryI F and CryIAc (synpro) proteins) with tolerance to herbicide (PAT protein) | Lepidoptera | Dow AgroSciences |
| 12 | BR102012018662 | Insect-resistant soybean event pDAB9582.814.19.1::pDAB4468.04. 16.1 (CryIF and CryIAC (synpro) proteins) with tolerance to herbicide (AAD-12 and PAT proteins) | Lepidoptera | Dow AgroSciences |
| 13 | BR102015025537 | Gho/Sec24b2 and Sec24b1 silencing (RNAi) | Coleoptera and/or Hemiptera | Dow AgroSciences |
| 14 | BR102013032916 | Reptin silencing (RNAi) | Coleoptera | Dow AgroSciences |
| 15 | BR102012025724 | pp1-87b silencing (RNAi) | Coleoptera | Dow AgroSciences |
| 16 | PI1011950 | Cry and Dig-11 proteins | Coleoptera | Dow AgroSciences |
| 17 | PI1015333 | Dig-3 proteins | Lepidoptera | Dow AgroSciences |
| 18 | BR102012025759 | rpa70 silencing (RNAi) | Coleoptera | Dow AgroSciences |
| 19 | BR102014031844 | Opposite Ras silencing (RNAi) | Coleoptera and/or Hemiptera | Dow AgroSciences |
| 20 | BR102012025657 | rps6 silencing (RNAi) | Coleoptera | Dow AgroSciences |
| 21 | BR112012014746 | Cry1ca and Cry1ab proteins | Lepidoptera | Dow AgroSciences |
| 22 | BR112012027140 | Cry34ab/35ab and Cry3ba proteins | Coleoptera | Dow AgroSciences |
| 23 | BR112012027139 | Cry34ab/35ab and Cry6aa proteins | Coleoptera | Dow AgroSciences |

Table 2 (Continued)

| Item | Patent application number | Event/Gene/Product/Process | Target insects | Owner |
|------|---------------------------|--|-------------------------------------|---------------------|
| 24 | BR112012027218 | Cry34ab/35ab and Cry3aa proteins | Coleoptera | Dow AgroSciences |
| 25 | BR112012027208 | Cry3aa and Cry6aa proteins | Coleoptera | Dow AgroSciences |
| 26 | BR112012014804 | Cry1fa and Cry1ab proteins | Lepidoptera | Dow AgroSciences |
| 27 | BR112012014665 | Insect-resistant maize event dp-004114-3 (Cry1F, Cry34Ab1 and Cry35Ab1 proteins) with tolerance to herbicide (PAT protein) | Coleoptera | DuPont |
| 28 | PI0919339 | Bt toxins | Lepidoptera | Pionner DuPont |
| 29 | PI1007260 | Bt toxins | Lepidoptera | Pioneer DuPont |
| 30 | PI0918766 | Bt toxins | Coleoptera | Pioneer DuPont |
| 31 | PI0924154 | Bt toxins | Lepidoptera | Pioneer DuPont |
| 32 | PI0924153 | Bt toxins | Lepidoptera | Pioneer DuPont |
| 33 | PI0919336 | Bt toxins | Lepidoptera | Pioneer DuPont |
| 34 | BR112012030913 | Nezara viridula genes silencing (RNAi) | Hemiptera | Pioneer DuPont |
| 35 | PI0823184 | Cyt1, Cry4A, Cry4B, Cry10, Cry11, Cyt2 and Cry3 proteins | Coleoptera and/or Lepidoptera | EMBRAPA |
| 36 | PI1102841 | Alpha-amylase inhibitor mutants | Coleoptera | EMBRAPA |
| 37 | PI0906128 | Cry8Ha protein | Coleoptera | EMBRAPA |
| 38 | BR102012033506 | Laccase family genes silencing | Coleoptera | EMBRAPA |
| 39 | BR102012033542 | Cry1Ia12 protein | Lepidoptera | EMBRAPA |
| 40 | BR102013031014 | Control strategy against <i>Athetis lepigone</i> based on Cry1A protein | Lepidoptera | Dabei Nong |
| 41 | BR102013031821 | Control strategy against <i>Conogethes punctiferalis</i> based on Cry1F protein | Lepidoptera | Dabei Nong |
| 42 | BR102014003618 | Control strategy against <i>Sesamia inferens</i> based on Cry1B protein | Lepidoptera | Dabei Nong |
| 43 | BR102013031822 | Control strategy against <i>Sesamia inferens</i> based on Cry1F protein | Lepidoptera | Dabei Nong |
| 44 | BR102013031734 | Control strategy against <i>Athetis lepigone</i> based on Cry1F protein | Lepidoptera | Dabei Nong |
| 45 | BR102013018436 | PIC9-modified protein | Lepidoptera | Dabei Nong |
| 46 | BR102013018337 | PIC9-modified protein | Lepidoptera | Dabei Nong |
| 47 | BR112013000262 | Insect-resistant maize events MIR604, BT11 and MIR162 (Cry3A, Cry1Ab and Vip3Aa20 proteins) | Coleoptera | Syngenta |
| 48 | PI0922656 | Insect-resistant maize event 5307 (FR8 protein and PMI gene marker) | Coleoptera | Syngenta |

9.283/2018) (Brasil, 2016, 2018b) and EMBRAPAII (Brasil, 2013), which aim to expand partnerships with companies and encourage biotechnological development by national companies and institutes. These efforts will certainly boost innovation and increase the number of biotechnological

products in the market and that of patent applications, consequently strengthening the Brazilian economy.

Although examining patent applications in Brazil is a very slow process – which affected our analyses – INPI has created strategies to accelerate it, stimulating applicants to

apply for patents in Brazil. Some of these strategies include priority for applications involving *green patents* (INPI, 2016a), which comprise technologies for the environment such as alternative energy sources, energy conservation, the management of residues and sustainable agriculture.

The INPI considers sustainable agriculture to be as follows: (1) reforestation techniques; (2) alternative watering techniques; (3) alternative pesticides and (4) soil improvement (e.g., residue-derived organic fertilizers). GM plants are not included in this panel, despite the potential environmental benefits mentioned above; this is an important topic that should be taken into account by INPI when considering priority application. In parallel, the pilot projects for priority application of requested patents from institutes of science and technology (IST Patents) (INPI, 2017a), micro- and small-sized enterprises (MSE Patents) (INPI, 2016b; 2017b) and of Brazilian origin, with priority rights assured for application in different national or international patent offices (BR Priority) (INPI, 2015b; 2017c), have been initiatives undertaken by INPI to accelerate the examination process.

The protection strategies involving gene pyramiding are another alternative for use by Brazilian companies and institutions. Moreover, knowing the existing technologies may facilitate the development of new technologies and expand opportunities for partnerships with owners. Finally, to better promote the independence of national companies in this sector, using technologies within the public domain is also an alternative, provided that there is the constant monitoring of activity in corresponding patent databases.

Conflicts of interest

The authors declare that the research was performed in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Uncited reference

OECD (2005).

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Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at [doi:10.1016/j.biori.2019.04.003](https://doi.org/10.1016/j.biori.2019.04.003).

References

- Brasil. *Lei n° 9.279, de 14 de maio de 1996. Lei da propriedade industrial.* (1996). http://www.planalto.gov.br/ccivil_03/Leis/L9279.htm Accessed 6 August 2013
- Brasil. *Lei n° 9.456/97, de 25 de abril de 1997. Lei de Proteção de Cultivares.* (1997). http://www.planalto.gov.br/ccivil_03/leis/L9456.htm Accessed 9 June 2017.
- Brasil. (1998). *Ministério da Ciência e Tecnologia. Comunicado n° 54, de 29 de setembro de 1998. Solicita da CTNBio liberação comercial de soja geneticamente modificada tolerante ao herbicida Roundup Ready.* Brasília: DOU Diário Oficial da União. Publicado no D.O.U. n° 188, de 01 de outubro de 1998, seção 03, página 56.
- Brasil. (2005a). *Ministério da Ciência e Tecnologia. Parecer técnico prévio conclusivo n° 513/2005. Liberação comercial de algodão geneticamente modificado resistente às principais pragas da Ordem Lepidoptera.* Brasília: DOU Diário Oficial da União. Publicado no D.O.U. n° 57, de 24 de março de 2005, seção 01, página 02.
- Brasil. *Lei n° 11.105, de 24 de março de 2005.* (2005). http://www.planalto.gov.br/ccivil_03/_ato2004-2006/2005/lei/l11105.htm Accessed 6 August 2013.
- Brasil. *Decreto n° 6.041, de 08 de fevereiro de 2007.* (2007). http://www.planalto.gov.br/ccivil_03/_ato2007-2010/2007/decreto/d6041.htm Accessed 6 August 2017.
- Brasil. (2011). *Ministério da Ciência e Tecnologia. Extrato de parecer n° 3024/2011. Liberação Comercial de feijão geneticamente modificado.* Brasília: DOU Diário Oficial da União. Publicado no D.O.U. n° 179, de 15 de setembro de 2011.
- Brasil. *Decreto de 02 de setembro de 2013.* (2013). http://www.planalto.gov.br/ccivil_03/_ato2011-2014/2013/dsn/Dsn13662.htm Accessed 6 August 2017.
- Brasil. *Lei n° 13.243, de 11 de janeiro de 2016.* (2016). http://www.planalto.gov.br/ccivil_03/_ato2015-2018/2016/lei/l13243.htm Accessed 15 May 2018.
- Brasil. *Decreto n° 9.283, de 7 de fevereiro de 2018.* (2018a). http://www.planalto.gov.br/ccivil_03/_Ato2015-2018/2018/Decreto/D9283.htm Accessed 15 May 2018.
- Brasil. (2018b). *Ministério da Ciência e Tecnologia. Extrato de parecer n° 5767/2018. Liberação planejada no meio ambiente (RN6).* Brasília: DOU Diário Oficial da União. Publicado no D.O.U. n° 179, de 21 de fevereiro de 2018, seção 01, página 05.
- Bravo, A., Likitvatanavong, S., Gill, S. S., & Soberon, M. (2011). *Bacillus thuringiensis: a story of a successful bioinsecticide. Insect Biochemistry and Molecular Biology, 41, 423–431.*
- CTNBio. (2017). *Relatório Anual 2016 da Comissão Técnica Nacional de biossegurança – CTNBio.* Disponível em <<http://ctnbio.mcti.gov.br/relatorios-anuais>>. Acesso em 09 jun 2017.
- CTNBio. (2017). *Resumo Geral de Plantas GM aprovadas para Comercialização. Comissão Técnica Nacional de Biossegurança. Ministério da Ciência, Tecnologia, Inovações e Comunicação.* Disponível em: <http://ctnbio.mcti.gov.br/documents/566529/1684467/Tabela+de+Plantas.pdf/e9d66306-bc49-4595-bd8a-805b727e7750?version=1.0>. Acesso em 30 de ago 2017.
- FAO. *The state of food insecurity in the world. Meeting the 2015 international hunger targets: taking stock of uneven progress.*

- (2015). <http://www.fao.org/3/a-i4646e.pdf> Accessed 25 May 2018.
- Fraley, R. T., Rogers, S. G., Horsch, R. B., Sanders, P. R., Flick, J. S., Adams, S. P., et al. (1983). Expression of bacterial genes in plant cells. *Proceedings of the National Academy of Sciences*, 80, 4803–4807.
- Globo. *Cade aprova fusão de Dow e DuPont, condicionada à venda de ativos*. (2017). <http://g1.globo.com/economia/negocios/noticia/cade-aprova-fusao-de-dow-e-dupont-condicionada-a-venda-de-ativos.ghtml> Accessed 15 June 2017.
- INPI. *Resolução PR n° 144, de 12/03/2015*. (2015a). <http://www.inpi.gov.br/links-destaques/sobre/legislacao-1> Accessed 15 June 2017.
- INPI. *Resolução PR n° 153*. (2015b). <http://www.inpi.gov.br/sobre/legislacao-1> Accessed 15 June 2017.
- INPI. *Resolução PR n° 175*. (2016a). <http://www.inpi.gov.br/sobre/legislacao-1> Accessed 15 June 2017.
- INPI. *Resolução PR n° 160*. (2016b). <http://www.inpi.gov.br/sobre/legislacao-1> Accessed 15 June 2017.
- INPI. *Resolução PR n° 191*. (2017a). <http://www.inpi.gov.br/sobre/legislacao-1> Accessed 15 June 2017.
- INPI. *Resolução PR n° 181*. (2017b). <http://www.inpi.gov.br/sobre/legislacao-1> Accessed 15 June 2017.
- INPI. *Resolução PR n° 180*. (2017c). <http://www.inpi.gov.br/sobre/legislacao-1> Accessed 15 June 2017.
- ISAAA. *Global Status of Commercialized Biotech/GM Crops: 2016*. (2016). <http://cib.org.br/estudos-e-artigos/relatorio-isaaa-2016-sobre-a-situacao-global-das-lavouras-transgenicas/> Accessed 20 August 2017.
- MAPA. *CultivarWeb*. (2017). http://extranet.agricultura.gov.br/php/snpc/cultivarweb/cultivares_protegidas.php Accessed 10 July 2017.
- OECD. *A framework for Biotechnology statistics. Organisation for economic co-operation and development*. (2005). <http://www.oecd.org/science/inno/34935605.pdf> Accessed 20 August 2017.
- OECD. *Key Biotechnology Indicators. July 2015*. (2015). <http://oe.cd/kbi> Accessed 6 August 2017.
- Rodrigues, R. L., Lage, C. L. S., & Vasconcellos, A. G. (2011). Intellectual property rights related to the genetically modified glyphosate tolerant soybeans in Brazil. *Anais da Academia Brasileira de Ciências*, 83, 719–730.
- Verde, F. R. V., Dos Santos, P. R., & Guerrante, R. D. (2015). Radar tecnológico INPI - Biotecnologia de brasileiros (não saúde). *Rio de Janeiro: INPI*.
- Valor Econômico. *Compra da Monsanto pela Bayer deve ser finalizada este ano, diz CEO global*. (2017a). <http://www.valor.com.br/agro/4919916/compra-da-monsanto-pela-bayer-deve-ser-finalizada-este-ano-diz-ceo-global> Accessed 16 June 2017.
- Valor Econômico. *A mais suíça das empresas chinesas*. (2017b). <http://www.valor.com.br/agro/4961464/mais-suica-das-empresas-chinesas> Accessed 21 June 2017.
- Valor Econômico. *A ascensão dos barões dos alimentos*. (2017c). <http://www.valor.com.br/opiniao/5010238/ascensao-dos-baroes-dos-alimentos> Accessed 21 June 2017.
- WIPO. *WIPO IP Statistics Data Center*. (2017a). <https://www3.wipo.int/ipstats>. Accessed 20 June 2017.
- WIPO. *WIPO IP Statistics Data Center*. (2017b). <http://www.wipo.int/ipstats/en/index.html#resources>. Accessed 12 August 2017.
- WIPO. *International Patent Classification (IPC), version 2013.01*. (2013). <http://web2.wipo.int/ipcpub/#refresh=page>. Accessed 15 June 2017.