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## The Biotechnological Promises and Bottlenecks for a Sustainable and Intensified Agriculture in Africa\*

by

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KEYWORDS. — Biotechnology; GM Crops; Africa; Sustainable Agriculture; Agricultural Research.

SUMMARY. — On a continent that has to cope with high rates of poverty, disease, and malnutrition, the agricultural and farming practices have evolved only slightly for centuries. Of all the recently developed technologies, biotechnology represents an innovation toolbox that could significantly improve the livelihood of sub-Saharan populations. Its impact could be far beyond that reached in the industrialized world, in which agricultural biotechnology has become a multibillion industry. In sub-Saharan Africa, the absence of this foundation acts as a brake on the elaboration of a sustainable biotechnology-based industry. A critical element should be the development of a regionally innovative community in the field of agricultural biotechnology which focuses on crops relevant for Africa, agricultural practices, and economic needs, but which is also sensitive to public concerns about the use of genetic modifications.

TREFWOORDEN. — Biotechnologie; GG-gewassen; Afrika; Duurzame landbouw; Landbouwkundig onderzoek.

SAMENVATTING. — De beloftes en beperkingen van de biotechnologie voor een duurzame en intensieve landbouw in Afrika. — Tot op de dag van vandaag hebben de huidige landbouwpraktijken maar weinig impact op het Afrikaanse continent dat sinds eeuwen geteisterd wordt door honger, ziektes en ondervoeding. Van alle recent ontwikkelde technologieën is biotechnologie er één die de levensomstandigheden van de bevolking in Sub-Sahara-Afrika aanzienlijk kan verbeteren. De impact van biotechnologie kan in deze regio zelfs groter zijn dan deze die bereikt werd in de geïndustrialiseerde wereld, waar ze een miljardenindustrie geworden is. In Sub-Sahara-Afrika ontbreekt deze basis, wat een belangrijke belemmering vormt voor de ontwikkeling van een duurzame op biotechnologie-gebaseerde industrie. Voor de landbouw in Sub-Sahara-Afrika is het heel belangrijk om een lokale en innovatieve gemeenschap te ontwikkelen die zich wil inzetten voor de implementatie van biotechnologie in de landbouw, gefocused op gewassen die belangrijk zijn voor de Afrikaanse bevolking, die landbouwpraktijken optimaliseert alsook de economische impact ervan evalueert. Tenslotte dient ze ook een antwoord te bieden op de

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lokale belangen en bezorgdheden rond het gebruik van genetische modificaties in de landbouw.

Mots-clés. — Biotechnologie; OGM; Afrique; Agriculture durable; Recherche agronomique.

RÉSUMÉ. — Les promesses et contraintes de la biotechnologie pour une agriculture durable et intensive en Afrique. — Sur un continent fortement touché par la pauvreté, les maladies et la malnutrition, les pratiques agricoles n'ont que peu évolué depuis des siècles. De toutes les technologies récemment développées, la biotechnologie pourrait améliorer de manière significative les conditions de vie des populations en Afrique subsaharienne. Son impact pourrait dépasser celui atteint dans le monde industrialisé où l'industrie de la biotechnologie agricole s'est développée et vaut des milliards d'euros. En Afrique subsaharienne, l'absence de cette assise constitue un frein à l'extension d'une industrie biotechnologique durable. Un élément critique est le développement d'une communauté régionale innovatrice en matière de biotechnologie agricole qui se consacrerait à la culture de plantes africaines, aux pratiques agricoles et aux impératifs économiques tout en étant sensible aux préoccupations du consommateur quant à l'utilisation controversée des technologies de modifications génétiques.



For centuries, agriculture has been undergoing major changes through technical and scientific innovations and through the evolution of farming practices that have allowed farmers to increase yield and productivity. An important challenge of today's agriculture is the need to decrease the environmental footprint by reducing the required amount of natural resources, such as land, water, and inputs (*e.g.*, seeds, fertilizers, and machinery), while still meeting the world's increasing needs in food, fuel, and fibre.

Whereas agriculture in Western countries nowadays strongly focuses on optimizing production, land, and resource utilization by the application of more technological tools and data (known as precision farming), developing countries, and especially those of sub-Saharan Africa, are still fighting to get access to efficient germplasms and inputs that could already highly improve yield and production. In fact, over the last fifty years, thanks to the so-called "green revolution", many countries were able to increase their food production and reach an enhanced level of food security. This green revolution was represented by the determinant work of Norman Borlaug, an American agronomist, who instigated wheat variety breeding programmes and resulting lines. These new varieties combined with new agronomical methods were beneficial not only for American and South-American countries, but also for Asia, where the productivity, not only in wheat, but also in other cereals, was highly increased. In 1970, Norman Borlaug was honoured with the Nobel Prize for Peace because of his influential work in globally reducing hunger and poverty. The green revolution made the use of inorganic fertilizers and pesticides more methodical, with exponential yield increases as a consequence. However, these rationalized agricultural practices have been largely criticized as well, because in many cases they have led to an intensified, large-scale monoculture type of agriculture. Still, in that period, the cereal crop production tripled when the world population doubled, hence, overcoming repetitive food shortage (PINGALI 2012). The green revolution had mainly been initiated to boost the yield of staple crops cultivated under favourable conditions, thus neglecting areas with poor soil and marginal productivity, such as most of the African continent. Although enormous progress has been made, Africa still has to undergo its own green revolution to be able to reduce famine and indigence.

Meanwhile, research on crop improvement has undergone radical changes with the rise of new knowledge and technologies. The increased understanding of the molecular mechanisms behind plant growth and development has opened a whole range of possibilities. Biotechnology from "bio" (life), "technos" (tool) and "ology" (study of) is defined by the use of living organisms to develop new products or processes. Its current impact on agriculture results mainly from the introduction of ameliorated crops that belong to the class of genetically modified (GM) organisms (GMOs).

Plant biotechnology is considered to have started approximately eight thousand years ago, when humans began domesticating plants and selecting the most suitable ones for cultivation, such as those with large seeds, short ripening time, etc. This tedious process changed when fundamental scientific knowledge was applied. For example, the laws of heredity discovered by Mendel have been significant to set up systematic breeding programmes. From the 1930s, mutation breeding was implemented to enhance crop qualities. This technique, which uses mutagenic radiation or chemicals to induce mutations in plants, accelerates considerably the discovery of beneficial mutations. Mutagenesis has been applied for decades to improve the crops, vegetables and fruit that we eat today. The elaboration and application of molecular biology techniques have later generated new tools to amend breeding with, for example, the development of markerassisted selection which has provided a genetic basis for the selection of new traits.

The discovery of the genetic transformation technology enabled a big jump forward in the creation rate of new traits over the past three decades. Through genetic transformation specific pieces of DNA could be integrated into the genome of a plant, thus inserting required new functions, such as insect resistance. This DNA fragment could belong to any living organism, because all share the same genetic code. Genetic modification through transformation has led to the so-called GMOs. However, noteworthy, genetic modifications are the essence of variety amelioration, the final aim of which is to provide genetic diversity with new traits of interest for the farmers and/or the consumers, either through conventional breeding techniques, genetic transformation, or any other available technology. Hereby, two notions should be distinguished, namely transgenesis and cisgenesis. Transgenesis implies the integration into a crop genome of a DNA piece from another organism that is not sexually compatible, whereas cisgenesis integrates a DNA fragment that could have been obtained through successive crosses with sexually compatible organisms, such as, for instance, a resistance gene from a relative of wild potato (*Solanum tuberosum*) integrated into a commercial potato variety. Both cisgenesis and transgenesis allow the insertion of a (or multiple) new trait(s), still keeping all the genetic characteristics of the original variety.

The first genetic transformations were published in the early 1980s and the group of Marc Van Montagu and Jeff Schell at Ghent University were pioneers in the field. In fact, the soil bacterium, *Agrobacterium tumefaciens*, had been found to induce tumors in plants (SMITH & TOWNSEND 1907), of which later the tumor-inducing agent had been discovered to be a small piece of circular DNA, designated as tumor-inducing (Ti) plasmid (ZAENEN *et al.* 1974). By means of this Ti plasmid from *Agrobacterium* as a vector, new genes could be integrated into a plant by replacing the genes responsible for the tumor formation with genes of interest (HERRERA-ESTRELLA *et al.* 1983).

In the meantime, science has made tremendous progress. One of the most remarkable discoveries was the bacterial adaptive immunity to viruses and plasmids (DOUDNA & CHARPENTIER 2014). This research has led to the creation of a new genome-editing tool, clustered regularly interspaced short palindromic repeats (CRISPR)/CRISPR-associated protein 9 (Cas9), abbreviated as CRISPR-Cas9 (MOJICA *et al.* 2005, POURCEL *et al.* 2005), which is transforming the field of biology and biotechnology research (DOUDNA & CHARPENTIER 2014). This new technology together with other gene-editing techniques allow genetic modifications at a previously unknown precision level, marking the starting point of innovations that might well revolutionize the agricultural sector in the near future.

Technically, the CRISPR-Cas9 technology is based on a bacterial defence mechanism that can be adapted to target a specific DNA fragment. The technology consists of the Cas9 enzyme and a guide RNA (gRNA) responsible for directing the enzyme to the DNA. The gRNA binds to the Cas9 protein and, upon binding, induces a conformational change in the protein that converts the inactive protein into its active form. When the Cas9 protein/gRNA finds a potential target sequence, it achieves a "cut and paste" that allows minor modifications into a specific site of the genome of a living organism. This technology is more precise than previous genetic transformation techniques, because the modification site can be selected. Furthermore, plants can be obtained without any foreign DNA, only the required change in their own genome is kept. Traditionally, transgenic crop development has been reserved to a few big companies because of the tremendous cost for development and the related regulatory expenses. These high expenditures could only be paid back for a few major crops of global importance. Plants obtained with the more recent gene-editing technologies, such as CRISPR-Cas9, if not regulated in the same way as GMOs, could offer new opportunities for small and medium-sized enterprises and for the public sector to ameliorate crops that had been disregarded until now because of the low market value and little commercial interest. In other words, this technology could be a major game changer in the development of efficient crops in sub-Saharan Africa, where many crops are of regional or national interest, but represent only a small fraction of the international commodity trade.

Developing countries, especially in Africa, face a number of challenges that cannot be tackled and solved easily. In many African countries, a certain number of gaps need to be filled to reach a satisfactory level of agricultural efficiency, including, for example, development of the proper infrastructure to ensure collection, transportation, and storage of the harvest. A wide range of pests and diseases (favoured by climatic conditions) and a low access to high-quality inputs (seeds, fertilizers, and pesticides) further undermine productivity. In addition, strong policies supporting the implementation of a competitive agriculture are essential for the expansion and professionalization of the agricultural sector.

Moreover, when production and capacity can be increased, a fair access to markets, either regional, national, or international, is needed. The low industrialization level does currently not offer a sufficiently developed network to process the agricultural production, so that it has to be conveyed as raw material that also depends largely on the market seasonal fluctuations. Additionally, Africa lacks the added value and job opportunities provided by the processing of raw production. Food processing would add price stability for the producer on the local and regional markets, because processed products that can be preserved and easily stored, are less dependent on farming cycles and market prices, also allowing a superior access to international markets.

Nevertheless, science and innovation can propose solutions not to be ignored in hunger-suffering countries. Today, the African agricultural productivity for a whole range of crops is much lower than that in the European Union, North and South America, and Asia. According to the Food and Agricultural Organization, the average yield of maize (*Zea mays*) cultivation reached 10.7 tons/ha in the USA in 2014, whereas the average yield in Africa was only 2.1 tons/ha (http:// www.fao.org/faostat/en/#data). This difference is not only true for maize, but also for crops, such as banana (*Musa* sp.), cassava (*Manihot esculenta*), cowpea (*Vigna unguiculata*), rice (*Oryza sativa*), sorghum (*Sorghum bicolor*), and others. Together with an exponentially growing demography and all challenges mentioned above, this yield disparity leads to food shortage and societal crises with a worldwide impact that need to be taken into account with all the tools and support available, including biotechnology.

Although a wide variety of products obtained with the recent CRISPR-Cas9 technology are not yet available, the genetic engineering technology has already been used since the 1990s and it is still being adopted at an increasing rate. In

1996, in the initial phase for GMO commercialization, 1.7 Mha were grown with GMO crops versus 189.8 Mha worldwide in 2017 (ISAAA 2017). Remarkably, the spectrum of developers is also enlarging and involves developing and growing economies. The cultivated surface in these emerging economies (including Brazil, Russia, India, China, South Africa) is now more important than that in the traditionally high-tech agricultural regions of the USA and Europe (ISAAA 2017). Still, on this vast surface mainly a few crops of global economic importance are cultivated, such as soybean (*Glycine max*), maize, cotton (*Gossypium hirsutum*), and oilseed rape (*Brassica napus*). With the aim of developing a sustainable agriculture in Africa, one might wonder whether these approaches would really be game changers and whether they would be possibly applicable in the context of the African agricultural systems.

Although no GM product developed in Africa is on the market yet, the number of scientific research initiatives has augmented and the proportion of African products in the pipeline to commercialization is higher than that of other continents (PARISI *et al.* 2016). Currently, most, if not all, GM products are merchandized by a small number of actors that can afford the high developmental and regulatory costs and involve mainly a few major crops, such as maize, cotton, or soybean. However, an increasing number of research and field trial initiatives are being taken on so-called "orphan crops" that are potentially of high importance at a regional or national level and less relevant for international trade (ISAAA 2016).

Many of these orphan crops, especially on the African continent, are an essential part of local diets and economies, *e.g.*, cassava, yam (*Dioscorea* sp.), sweet potato (*Ipomoea batatas*), cowpea, and grass pea (*Lathyrus sativus*). Moreover, these crops are often better adapted to the local environment and agricultural practices and, thus, potentially more resilient to the biotic or abiotic consequences of climatic disturbances than other crops. Therefore, orphan crops or, more correctly, crops that have been neglected by the regional, national, and international research programmes are slowly gaining the attention they deserve by the biotechnology field. Until recently, these geographically limited markets had raised only a poor interest by the agrobiological industry, but the public sector through public or private partnerships is gradually performing research on these crops. This interest can even be amplified and extended to small and medium-sized enterprises, when cheaper and less effort-intense technologies, such as CRISPR-Cas9, can be applied for crop improvement.

Currently, regarding the GM traits in the Research and Development pipeline, for instance provitamin A-enriched crops, private as well as public initiatives seem to carry out research to develop consumer-beneficial crops in addition to the previous more farmer-oriented traits, such as herbicide resistance. Although these products might not always be relevant for the global industrial sector, they might potentially have a higher impact in developing regions and be more beneficial to small-holder farmers and consumers than in the industrialized countries.

The most mediatized example of such a GM crop developed by the public sector has probably been the so-called golden rice. Golden rice has been improved genetically to produce provitamin A (YE et al. 2000) and to become an additional tool in the fight against nutritional imbalances in the diet of the poorest, *i.e.*, a lack of provitamin A in a child's diet can cause immune deficiency with total blindness as a possible consequence. Although it has taken a long time to clear scientific and regulatory issues, golden rice is now approved in New Zealand, Australia, Canada, and lately by the Federal Drug Authority of the USA, but in the Philippines and Bangladesh, where it could have the greatest impact on the children's health, the approval is still pending. Importantly, this golden rice has no commercial purpose, only a humanitarian reason of existence. In this same humanitarian spirit, a series of projects are running on previously neglected crops, such as the development of virus resistance in cassava, biofortification in sorghum and banana, aflatoxin resistance in groundnut (Arachis hypogaea L.), and drought tolerance in tropical maize. All these projects could bring specific solutions to challenges faced locally or regionally by farmers and consumers.

For instance, the Queensland University of Technology (Australia) and the National Agriculture Research Organization (NARO) of Uganda coordinate a project funded by the Bill and Melinda Gates foundation to create a biofortified banana enriched in provitamin A. These banana plants have been successfully tested in the field at the NARO of Uganda (PAUL *et al.* 2018). Provitamin A-enriched bananas could have a strong impact for the Ugandan population that relies on cooking bananas as main starch source. In addition, the International Institute for Tropical Agriculture (IITA) is developing banana plants resistant to either bacterial wilt or nematodes. Diseases, such as bacterial wilt, can destroy a whole plantation and no direct alternative solutions are available besides the use of resistance genes from another plant, for instance, sweet pepper (*Capsicum annuum*) (TRIPATHI *et al.* 2017).

The impact of climate change on the African continent is predicted to be severe, especially because of the mostly rain-fed African agriculture. The climatic variability and especially droughts and extreme weather events can highly affect yield and crop quality. To address this challenge, two major initiatives have been undertaken on maize, which is a major food crop in African countries, namely the Drought-Tolerant Maize for Africa (DTMA) project implemented by the Centre for International Maize and Wheat Improvement (CIMMYT) and IITA and the Water-Efficient Maize for Africa (WEMA) project, a public/private partnership, in which the associated company provides the technology free of royalties (DE BUCK 2017). The latter project is currently testing varieties in field trials in several countries of the African continent.

Moreover, research and engineering of legume crops have been undertaken as well. As a leading example, cowpea is grown on more than eight million hectares in West and Central Africa, with Nigeria the largest producer with four million hectares (GÓMEZ 2004). Still, the demand is higher than the production. Hence,

the creation of a variety resistant to the pod borer would be a real solution for the yield losses the producers are facing because of this disease. This project is a private/public partnership funded by the United States Agency for International Development (USAID).

These initiatives are all at different developmental stages, namely from research to product release. Currently, only a few products, such as GM maize and GM cotton, are cultivated in South Africa and South Sudan, but still no products made in Africa for Africa are on the market, although as mentioned above several products are in the pipeline.

Among African countries, only two (South Africa and Sudan) currently grow GM crops for commercial purposes, but thirteen countries already host research programmes and/or conduct field trials with different GM crops produced by private companies or by public or public/private partnerships (ISAAA 2017). Ethiopia and Nigeria, the two most populated countries of Africa, have recently approved the commercialization of a GM cotton and, also in Nigeria, an insect-resistant (Bt) cowpea has been released in the meantime.

Not only for commercialization and cultivation of these GM crops, but also for research and their growth in field trials, an efficiently working regulatory framework is essential. Such a legal framework, which exists in most countries, regulates the use of GM technology-based products, *i.e.* from seed to final product. Since 2017, fourteen African countries have established a regulatory framework and have implemented it, for example, to control imports, laboratory research, field trials, or even commercialization and cultivation, as in South Africa and Sudan. In addition, eight other countries have a law in place that has never been applied before for field trials or other activities. However, of the fifty-four (fifty-five with western Sahara) African countries, only less than half of them have a legislation that allows the regulation of GMO research, cultivation, or import. Furthermore, in many of the countries that have a biosafety law, a capacity-building effort of the responsible governmental bodies needs to be strengthened. In fact, many countries use the African law model for biotechnological safety which was enacted by the African Union in 2007 as a base for the establishment of a national legislation. This step is essential to comply with the obligations under the Cartagena Protocol on Biosafety (Secretariat of the Convention of Biological Diversity 2000), which has been ratified by most African countries. Nevertheless, as any model law, it can serve only as a base and needs to be adapted to the national regulations to allow proper endorsement. Inaction from authorities leads to an unclear situation that frightens potential stakeholders willing to invest in the country. Such a reluctance has been noticed not only in Africa, but also in Europe, where the regulatory framework and the administrative implementation have not evolved in parallel with the technologies and the knowledge gained over the years. As a consequence, decisions are often taken without proper consideration of scientific facts.

In spite of regulatory frameworks installed or adopted after the appearance of any new major technological advance, each innovation has had its own acceptance history, some more tumultuous than others. Every day, new technical innovations are available and marketed. Some innovative technical products have a rather fast adoption rate by the public, such as, for example, the internet or the smartphone. In sharp contrast, GMOs have undergone a much more difficult and diverse acceptance rate, depending on region or country. The late Professor Calestous Juma illustrated very nicely in his last book how the adoption of some products, such as coffee, margarine, or even alternative electrical current, had faced struggles (JUMA 2016). Many parameters behind the fear for novelty are of social, cultural, and economic order and, thus, emotional parameters weigh potentially more than rational or scientific arguments (BLANCKE et al. 2015, 2016; COUÉE 2016). Although scientific facts in the case of GMOs might be not so straightforward to be understood by the broad public, the immediate benefits for the consumer might also be an important factor that can be evident, as, for example, in the case of the smartphone technology. Concerning the first GMOs on the market, the direct advantage was for the farmer, because initially the new traits were herbicide tolerance and insect resistance. Furthermore, in a period in which Europe was the scene of food safety scandals, such as the bovine spongiform encephalopathy and the dioxin crisis, GMO acceptance became problematic, but by adopting a precautionary principle to its extreme, development and implementation of new technologies can be blocked and, hence, prevent countries or regions from enjoying the benefits of a significant agricultural development. In Africa, in contrast to Europe or the USA, a large proportion of its population is still involved in agriculture with, for instance, 69 % farmers in Uganda versus only 1.3 % in Belgium (https://data.worldbank.org/indicator/ SL.AGR.EMPL.ZS?view=map). Consequently, the advantages of GM technology are more obvious and direct for a large population.

In other words, African countries have to weigh the risks and benefits in adopting technologies, such as GM or gene editing. This evaluation to make strong political decisions can be done only when the decision-makers have access to well-balanced and science-based information.

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