

# Photoactive TiO<sub>2</sub> antibacterial coating on surgical external fixation pins for clinical application

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**Abstract:** External fixation is a method of osteosynthesis currently used in traumatology and orthopedic surgery. Pin tract infection is a common problem in clinical practice. Infection occurs after bacterial colonization of the pin due to its contact with skin and the local environment. One way to prevent such local contamination is to create a specific coating that could be applied in the medical field. In this work, we developed a surface coating for external fixator pins based on the photocatalytic properties of titanium dioxide, producing a bactericidal effect with sufficient mechanical strength to be compatible with surgical use. The morphology and structure of the sol-gel coating layers were characterized using, respectively, scanning electron microscopy and X-ray diffraction. The resistance properties of the coating were investigated by mechanical testing. Photodegradation of acid orange 7 in aqueous solution was used as a probe to assess the photocatalytic activity of the titanium dioxide layers under ultraviolet irradiation. The bactericidal effect induced by the process was evaluated against two strains, ie, *Staphylococcus aureus* and multiresistant *Staphylococcus epidermidis*. The coated pins showed good mechanical strength and an efficient antibacterial effect after 1 hour of ultraviolet irradiation.

**Keywords:** hybrid sol-gel, external pin fixation, titanium dioxide, antibacterial effect, mechanical strength, ultraviolet photoactivity

## Introduction

External fixation is a method of osteosynthesis commonly used by surgeons in the fields of orthopedic surgery and trauma. The main feature of this method is that the implant of the assembly, called the tube fixator, is located outside the body and connected to bone by transcutaneous pins.<sup>1-3</sup> Complications associated with these pins, mainly pin tract infection, are an important source of morbidity that increase the complexity of care, with additional surgical procedures, resulting in longer hospital stays<sup>4</sup> and higher socioeconomic costs.<sup>5</sup>

Although pin tract infection is a common problem in clinical practice, there is no consensus regarding its definition, which is reflected in the wide ranges of incidence (0%–80%) reported in the literature.<sup>6-9</sup> Infection occurs after bacterial colonization of the pin, due to its contact with the skin and local environment, and phenomena brought about by surgery, such as a decrease in the local immune response, soft tissue trauma, and bone necrosis.<sup>10,11</sup> Infection can develop locally, but may also spread to adjacent soft tissue and bone, leading to osteomyelitis and loss of mechanical anchorage of the pin.

Diagnosis of pin tract infection should be based on nonspecific clinical arguments such as local inflammation, presence of discharge or pus, and pain as described in the Checketts–Otterburn classification.<sup>12</sup> Unfortunately, there is no objective test or

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standardized tool to confirm the diagnosis, and only evolution and time will do so. In a study of 214 implanted pins in 42 patients, Mahan et al<sup>13</sup> found a bacterial colonization rate of 74.8%, mainly involving three species, ie, *Staphylococcus epidermidis* (90.6%), *Staphylococcus aureus* (37.5%), and *Escherichia coli* (9.4%). *S. aureus* was more frequently associated with local infection and osteolysis than *S. epidermidis*. Consequently, since the 1970s, clinicians have tried to solve the problem of pin tract infection using a number of methods,<sup>14</sup> including use of alloys to manufacture pins or coating the pin with antibiotics or chemical substances (such as hydroxyapatite), silver particles, and zinc or titanium oxide, and applying external electromagnetic fields. However, none of the latter experiments settled the issue.

Since a publication by Matsunaga et al<sup>15</sup> in 1985, the bactericidal effect induced by titanium dioxide (TiO<sub>2</sub>) exposed to ultraviolet irradiation has been used successfully in many areas<sup>16–19</sup> like disinfection of water and textiles and in other cleaning processes. TiO<sub>2</sub> is indeed one of the most effective photocatalysts currently in use due to its strong oxidizing power, lack of toxicity, and long-term physical and chemical stability. It has been widely used for the decomposition of organic compounds and to destroy microbial organisms, such as cancer cells, viruses, and bacteria, and has potential application in the sterilization of medical devices and air-conditioning filters because of its self-sterilizing ability.<sup>20–23</sup> When irradiated with near ultraviolet light, TiO<sub>2</sub> has strong bactericidal activity.<sup>24</sup>

Other antibacterial materials available include silver-based compounds, which have been widely investigated as antibacterial agents because of their potent broad-spectrum antibacterial activity.<sup>25–29</sup> However, the antibacterial mechanism of silver species raises concerns about their potential cytotoxicity.<sup>30–32</sup> It is suggested that diffusing silver may act as a Trojan horse by entering cells and then releasing silver ions that damage intracellular function.<sup>32</sup> Development of inorganic antibacterial agents with high antibacterial activity, biosafety, and hopefully osteoconductivity is necessary. TiO<sub>2</sub> is a good candidate because of its efficient photoactivity and lack of toxicity.

In this work, we hypothesized that the photoactive properties of TiO<sub>2</sub> could be used to prevent pin tract infection. Scanning electron microscopy and X-ray diffraction were used to characterize the morphology and structure of the coating layers. Photodegradation tests of acid orange 7 (AO7) in aqueous solution were used to assess the photocatalytic activity of TiO<sub>2</sub> coatings exposed to ultraviolet irradiation. Mechanical tests were performed using a fresh bovine femur

to evaluate the strength of the coating. Finally, we evaluated the bactericidal effect of the coating using *S. aureus* and *S. epidermidis* strains.

## Materials and methods

### External fixator pin, metal disk, and quartz substrates

Stainless steel pins (FeCrNiMo18-15-3; ~ AISI 316 low carbon vacuum melt), 150 mm in length and 4 mm in diameter, and produced by L Klein (Bienne, Switzerland) were provided by Biomet, Inc (Warsaw, IN, USA). For technical reasons, it was necessary to use substrates of a suitable format. Stainless steel disks (AISI 316 FeCrNiMo18-10-3) 15 mm in diameter and 0.25 mm in thickness were provided by Goodfellow SARL (Lille, France). The coatings on the quartz substrates were specifically designed for use in X-ray analysis.

### Titanium sol formulation

In this work, the precursor solution was a mixture of titanium isopropoxide 97% (Sigma-Aldrich, St Louis, MO, USA) and ethyl alcohol 99.5% (Prolabo, VWR International, Radnor, PA, USA).

In order to moderate alkoxide reactivity, an organic ligand, acetylacetonate (99.5%, Fluka, Sigma-Aldrich), was added to the precursor solution. A drop of deionized water was used to start the hydrolysis–condensation reactions. The chemical composition of the starting alkoxide sol was Ti(OC<sub>3</sub>H<sub>7</sub>)<sub>3</sub>:C<sub>5</sub>H<sub>8</sub>O<sub>2</sub>:C<sub>2</sub>H<sub>5</sub>OH:H<sub>2</sub>O (molar ratio 1:1:20:1). An organic spacer, polyethylene glycol (Sigma-Aldrich, number average molecular weight 600), was added to increase the porosity of the layer (volume equal to 20% of the weight).

### Cleaning and dip coating of substrates

The substrates were degreased by successive sonication in trichloroethylene (98%, Prolabo), acetone (99%, Fisher Scientific, Waltham, MA, USA), and methanol (99.5%, UCB, Brussels, Belgium), followed by rinsing with deionized water and blowing dry with nitrogen. Samples were immersed in the sol for 1 minute. The withdrawing speed was set to 31 cm per minute to obtain a suitable coating thickness.

### Annealing treatments

Annealing treatments were carried out in an RS 80/750/11 Nabertherm tubular furnace equipped with a P300 processor. Coatings were treated in air, using a ramp of 5°C per minute to 500°C and then isothermal treatment at 500°C for

1 hour. Samples were then allowed to cool down to room temperature.

### X-ray diffraction analysis

The crystalline structure of the TiO<sub>2</sub> layers was determined by X-ray diffraction, using a Philips X'pert pro diffractometer with Cu K $\alpha$  radiation.

### Field emission scanning electron microscopy

The morphology of the samples was characterized using a Zeiss Supra 55VP field emission scanning electron microscope (FESEM) equipped with a Gemini column. To limit the effects of charge, a low voltage (3 kV) was used. Pictures were obtained with the secondary in-lens electron detector. Samples were observed on a flat view or with a various tilt angle.

### Photodegradation of AO7

Photodegradation of AO7 was used as a probe to assess the photoactivity of the TiO<sub>2</sub> layers. Photocatalytic experiments were conducted using an aqueous solution of AO7 (also known as Orange II; Acros Organics, Geel, Belgium) at a concentration of  $5.0 \times 10^{-5}$  mol/L, placed in a cylindrical glass reactor equipped with a magnetic stirrer. The glass reactor was irradiated with a polychromatic fluorescent ultraviolet lamp (Philips TDL 8 W; total optical power, 1.3 W), in a configuration providing about 0.35 mW/cm<sup>2</sup> at the sample surface. The photodegradation kinetics were recorded (in triplicate) by assaying the AO7 solution exposed to different ultraviolet irradiation times using a PerkinElmer Lambda 35 ultraviolet spectrophotometer. Quartz glass cells with an optical pathway of 1 cm were used. Deionized water was taken as the reference. After photodegradation of the dye, we monitored the decrease in absorbance of the solution at 483 nm (strong absorption band of AO7).

### Mechanical tests

The mechanical tests were performed by an orthopedic surgeon, and consisted of screwing and unscrewing a coated pin in a fresh cow femur, precisely as it would be done in a human patient (with a dedicated surgical ancillary) when making a unilateral external fixation. The self-drilling and self-tapping pin was introduced using a T-handle with the chuck tightened at maximum strength. The pin was screwed into the bone to a depth of about 1 cm after threading. A tube was then connected to the pin with connecting rods. After these tests, the coating behavior on the pin was investigated by FESEM.

### Bacterial culture

The bacterial strains used were *S. aureus* (ATCC 6538) and a clinical multiresistant strain of *S. epidermidis* (isolated in our university hospital). The strains were cultured for 18 hours in trypticase soy medium. Next, 50  $\mu$ L of each inoculum with a concentration of  $4 \times 10^8$  colony forming units (CFU)/mL were placed on each coated or uncoated disk, previously sterilized by autoclaving. The sample was incubated for 18 hours at room temperature in a dry atmosphere. Each disk with its bacterial load was then exposed to ultraviolet light. Irradiation was provided by a Philips polychromatic fluorescent ultraviolet lamp (Cleo Performance) with a total power of 40 W, in a configuration delivering about 5 mW/cm<sup>2</sup> at the surface for 15, 30, 45, 60, and 90 minutes of single exposure. The viable bacteria count on the disks was measured after disintegration of the biofilm by sonication in 1 mL of saline solution. The number of CFU in the suspension was then determined after dilution and plating on trypticase soy agar, incubated at 37°C for 18 hours. Non-viable bacteria on the coated disks were counted after 90 minutes of ultraviolet irradiation. All tests were performed in triplicate.

### Statistical analysis

Bacteriological data were subjected to the statistical analyses described below. As usual, bacterial concentrations were log-transformed to avoid the extreme skewness effect of the original concentration. The TiO<sub>2</sub> coating and ultraviolet exposure were defined as binary variables. Bacterial concentrations were determined at six time points, ie, 0, 15, 30, 45, 60, and 90 minutes. For each observation, bacterial concentrations were counted on three different disks with a double reading. The reliability of the two bacterial concentration readings and the three disk counts was almost perfect, with all intraclass correlation coefficients being above 0.9. The effects of TiO<sub>2</sub> coating, duration of ultraviolet exposure, and the *staphylococcus* strain used were tested in a generalized linear model, using bacterial concentration as the response variable, which included some first-order interactions, notably between the TiO<sub>2</sub> coating and ultraviolet exposure and between time variable and each of the aforementioned factors. Finally, a second-order interaction was evaluated between TiO<sub>2</sub> coating, ultraviolet exposure and time. On condition of significant effect, details were provided, performing Tukey's post hoc multiple comparisons procedure. All analyses were performed using SAS version 9.4 (SAS Institute Inc, Cary, NC, USA) and conducted setting the type I error rate (*P*) at 0.05.

## Results

### X-ray analysis

The X-ray characterizations of the TiO<sub>2</sub> layers on the quartz substrate before and after annealing at 500°C in air for 1 hour are presented in Figure 1. The non-annealed TiO<sub>2</sub> layer was amorphous. In order to convert the TiO<sub>2</sub> layer to a crystalline phase, the sample was annealed in air at 500°C, using a ramp rate of 5°C per minute. We observed the characteristic line of anatase (1 0 1). Crystallite size was calculated using the Scherrer formula:

$$D = 0.94\lambda / \beta \cos\theta$$

where  $\lambda$  is the wavelength of the CuK $\alpha_1$  line,  $\theta$  is the Bragg diffraction angle, and  $\beta$  is the full-width at half maximum of a peak. We calculated the crystallite size by using the full-width at half maximum of the anatase (1 0 1). The average crystal size was about 64 nm.

### Sample surface morphology

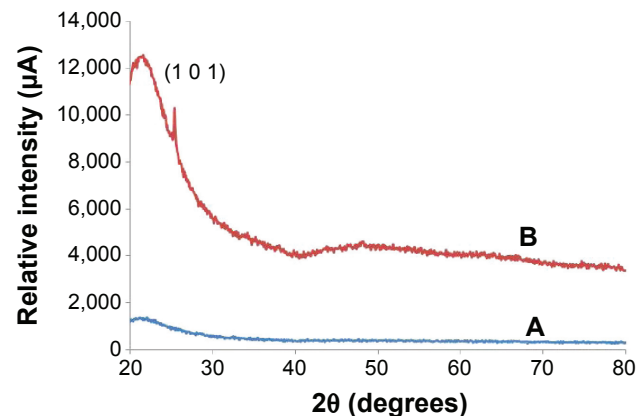
#### Initial surface images of disk and pin fixator

FESEM images of the iron disk and pin fixator are presented in Figure 2. The surface of both samples was rough and scratched. There was no marked difference between the initial surface of the disk and pin fixer.

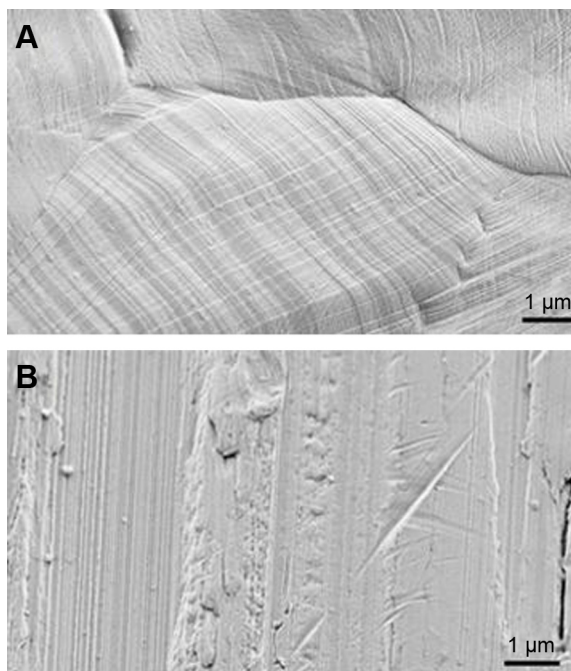
#### Surface images of coated disk and pin fixator

The sol formula, the dip-coating process parameters, and the annealing treatment allowed full coverage of the substrates for both the iron disk and pin fixator. Some scratches were detected on the layer (Figure 3). These punctual defects did not affect the overall strength of the cover layer.

The thickness of the oxide layer ranged between 300 and 500 nm (measured by FESEM) depending on the nature of



**Figure 1** X-ray diffraction patterns of TiO<sub>2</sub> layers on quartz substrate. **Notes:** (A) As grown and (B) annealed at 500°C.



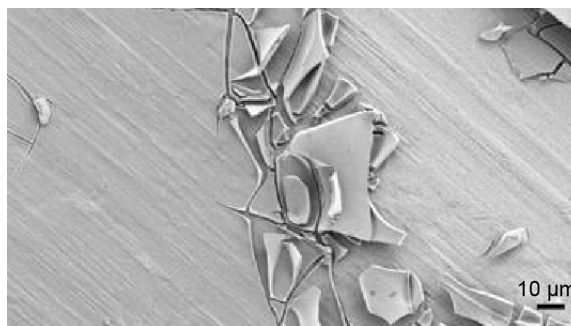
**Figure 2** Field emission scanning electron micrographs of iron disk (A) and pin fixator (B).

the substrates. Using high magnification at the nanoscale, we highlighted the grain morphology of the layer (Figure 4).

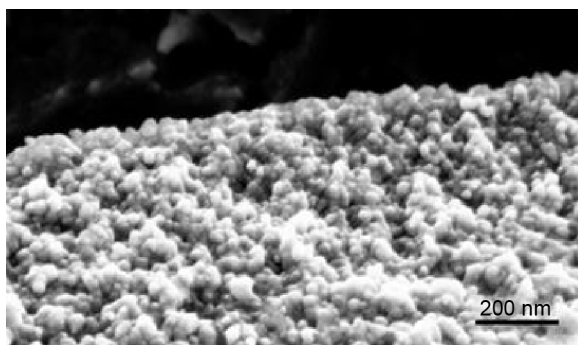
### Mechanical testing

Unilateral external fixation was performed on a fresh cow femur (Figure 5). Mechanical experiments were done in “extreme conditions” rather than classical clinical practice. There was no detectable change in the mechanical properties of the covered pin when compared to the uncovered pin. Thus, resistance of this layer to scratching would be preserved in the clinical setting.

An FESEM image showing the contact area after tightening of the pin with a T-handle and chuck used for the screwing and unscrewing experiments is given in Figure 6. Damaged areas on the coating can be seen, consisting of



**Figure 3** Field emission scanning electron micrograph of a punctual scratch on a cover layer.



**Figure 4** Field emission scanning electron micrograph showing the granular structure of the oxide layer at nanoscale.

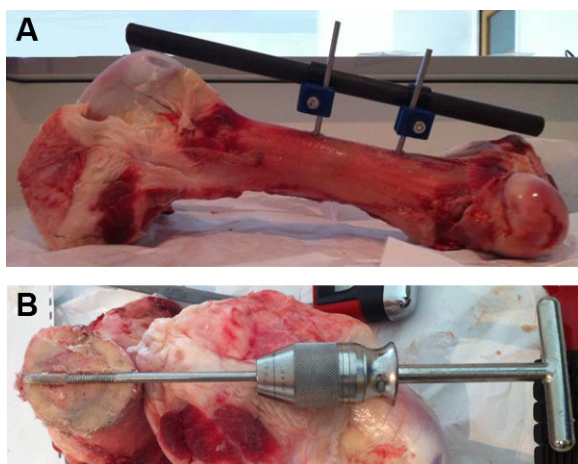
scratches and crunches in the area of contact between the grip and the coated pin.

Concerning the area of contact between the coated pin and bone, simulating overscrewing, we observed a crushing effect on the layer in the area of contact, resulting in some scratches at the interface (Figure 7).

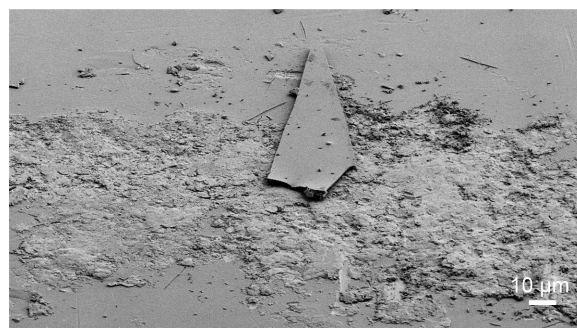
### Kinetics of photodegradation

AO7, which contains an azo bond, is a model molecule commonly used in photocatalytic tests to simulate azo dyes in wastewater pollutants coming from industry.<sup>33</sup> We did not find significant adsorption of the dye in the first step of the experiment, ie, immersion of the coated sample in AO7 solution under stirring in the dark. However, under ultraviolet irradiation, we observed a gradual discoloration of the solution, consistent with the decrease in absorbance recorded versus irradiation time.

Figure 8 shows the photodegradation kinetics of the organic dye versus irradiation time. After 8 hours of cumulative



**Figure 5** Illustrations of the mechanical test performed.  
**Notes:** (A) Image of a unilateral fixator assembly performed on a cow femur. (B) Visualization of the insertion of the pin into the bone on a horizontal section.



**Figure 6** Field emission scanning electron micrograph of the contact area between the grip and the coated pin.

ultraviolet irradiation, about 64% of the initial AO7 molecular species was degraded. This result confirmed the photoactivity of the coated sample, which was the consequence of crystallization of the TiO<sub>2</sub> layer during annealing. The anatase phase, previously detected in the X-ray experiments, was responsible for induction of ultraviolet-assisted degradation of the molecular dye species.

### Antibacterial effect

The inactivation kinetics for the two bacterial strains were found to be similar overall, and are presented in Figure 9 (*S. aureus*) and Figure 10 (*S. epidermidis*). For this reason, the results are analyzed and interpreted together. After incubation (for 18 hours), the inoculum concentration was recorded to be about 10<sup>6</sup> CFU/mL for *S. aureus*, but was significantly ( $P < 0.001$ ) higher for *S. epidermidis* (about 10<sup>7</sup> CFU/mL), without any significant difference between the coated and uncoated disks ( $P = 0.979$ ). Concerning disks kept in the dark (control group), no significant changes ( $P = 0.981$ ) in bacterial concentration were noted throughout the procedure.

After 15 minutes of ultraviolet irradiation, we noticed a significant decrease ( $P < 0.003$ ) in bacterial concentration (about 2 log CFU/mL) for both the coated and uncoated disks. Until the end of the experiment, ultraviolet irradiation alone proved to have no more effect on bacterial concentration (“uncoated disk with ultraviolet” group); however, after 60 minutes of ultraviolet irradiation, an additional significant decrease ( $P = 0.0014$ ) was noticed with the TiO<sub>2</sub>-coated disks. This observation was confirmed and amplified after 90 minutes of ultraviolet irradiation ( $P = 0.001$ ; about 2 log CFU/mL).

Our results indicate a rapid and lethal effect of ultraviolet light on bacteria during the first step of the experiment. Importantly, they show a further cumulative detrimental effect of the photoactive TiO<sub>2</sub> coating and ultraviolet irradiation on survival of bacteria. The role of TiO<sub>2</sub> is to promote



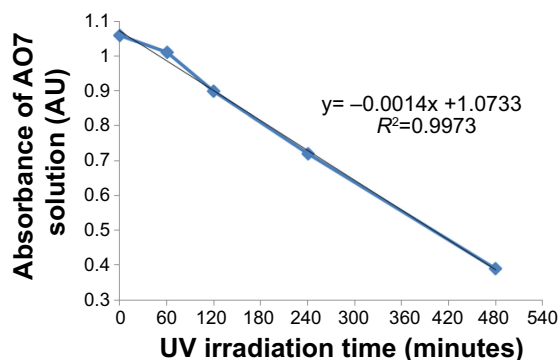
**Figure 7** Field emission scanning electron micrograph of the contact area between bone and the coated pin after explantation.

and accelerate inactivation of bacteria at lower levels of ultraviolet exposure.

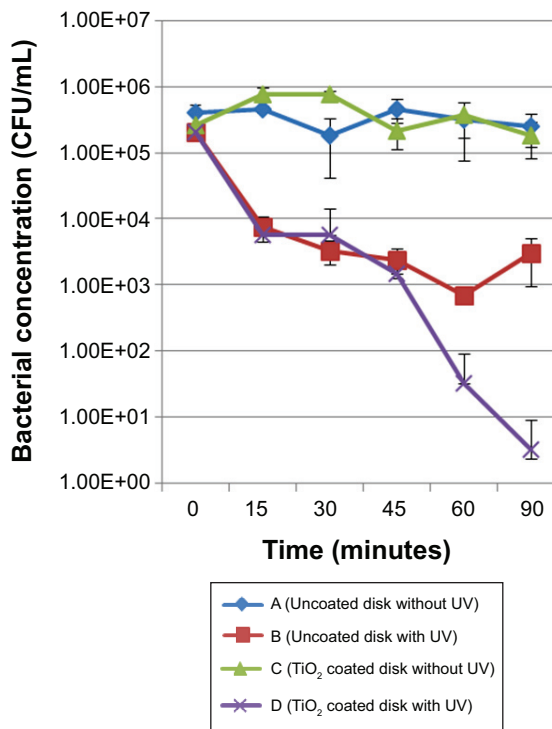
### Discussion

The use of TiO<sub>2</sub> coating on orthopedic implants has been well researched.<sup>34,35</sup> However, in this study, we used sol-gel and dip-coating processes for coating rather than the plasma ion implantation techniques used in other studies. Our study entailed application of a TiO<sub>2</sub> coating on a specific stainless steel (FeCrNiMo18-15-3; ~ AISI 316 low carbon vacuum melt), which is used commercially to create the pin. This technique has the advantages of being simple and cost-effective for production on an industrial scale. The novelty of our work is that mechanical tests were done to assess the feasibility of including this layer in real-life conditions, which has rarely been included in other studies. Finally, we found comparable results for bacterial inactivation, including resistant nosocomial strains that are problematic in everyday practice.

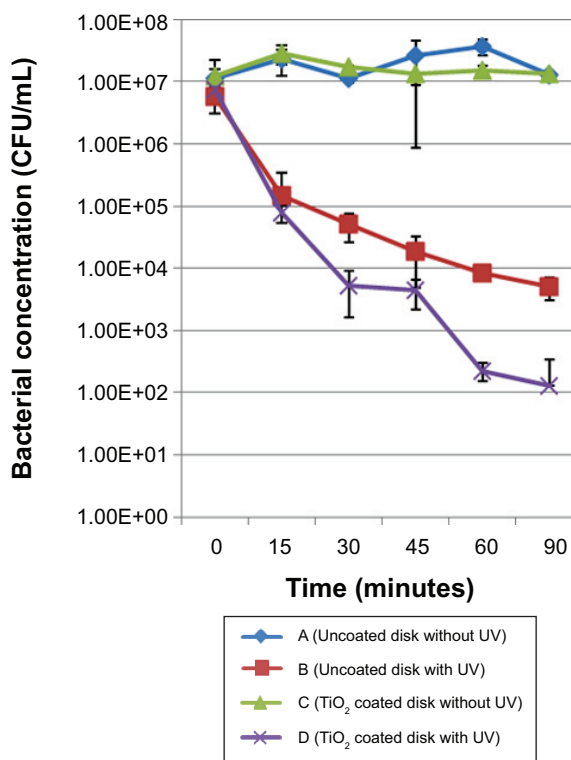
Our hypothesis was that the photoactive properties of TiO<sub>2</sub> could be used to prevent pin tract infection, and our study results show that the TiO<sub>2</sub> coating had bactericidal effects relevant to current problematic strains. Thus, we are able to confirm the feasibility of using this new coating for external fixator pins. However, a lot of additional work will have to be done to be able to transfer our findings to clinical practice.



**Figure 8** Photodegradation of AO7 solution in the presence of TiO<sub>2</sub>-coated disk versus UV irradiation time, as measured by the solution’s absorbance at λ =483 nm. **Abbreviations:** AO7, acid orange 7; AU, absorbance units; UV, ultraviolet.



**Figure 9** Inactivation kinetics for the *Staphylococcus aureus* strain. Comparison between coated and uncoated samples submitted to the same UV treatment duration. **Abbreviations:** CFU, colony forming units; UV, ultraviolet.



**Figure 10** Inactivation kinetics for the *Staphylococcus epidermidis* strain. Comparison between coated and uncoated samples subjected to the same UV treatment duration. **Abbreviations:** CFU, colony forming units; UV, ultraviolet.

TiO<sub>2</sub> seems to be safe or at least have controllable side effects, with the only potential risk being diffusion of free TiO<sub>2</sub> molecules within the body.<sup>36–39</sup> However, in this case, free TiO<sub>2</sub> particles cannot diffuse into the body because the pin thread is not coated. There is no contact between the coating and the patient's tissues. However, toxicity studies (eg, fibroblast cultures in presence of the substrate) are needed to confirm the safety of this technique. Ultraviolet irradiation is limited to the TiO<sub>2</sub> coating (ie, the patient's skin would not be exposed to ultraviolet rays). Exposure of patients to ultraviolet irradiation is indeed the key issue, and should be kept to a minimum. In our opinion, chemical and physical protection (eg, a custom-made cover applied all around the external fixator) would be necessary to avoid the adverse effects of ultraviolet irradiation, principally on the skin.

Sol-gel and dip-coating processes were used to create the coating because they are simple and cost-effective. The FESEM analysis of morphology showed full coverage of the substrate, with two superimposed layers showing cracks and scales. Mechanical tests confirmed the limitations of the under extreme conditions; nevertheless, the deeper layer was almost intact. To prevent this weakness, optimization of the layer could entail exchanging the sol formulation for a hardening element like zirconium, or using a modified annealing protocol.

Photodegradation of an AO7 solution confirmed the photocatalytic activity of the coating. However, the reaction was slow, with a 64% decrease in absorbance after 8 hours of cumulative ultraviolet irradiation. This was possibly due to the low power (0.35 mW/cm<sup>2</sup>) and the nonspecific spectrum (350–400 nm) of the ultraviolet lamp. Although our experimental conditions are different, our results are similar to those found in the literature.<sup>40,41</sup>

The bacterial parameters were chosen to simulate pin tract infection involving the most common causative organism (*S. aureus*)<sup>13</sup> and the most common source of hospital-acquired infection (multiresistant *S. epidermidis*).<sup>42</sup> The conditions chosen (ie, concentration, trypticase soy medium, incubation phase) allowed us to carry out bacterial survival and monitor development of biofilm.

The concentration of *S. epidermidis* was significantly higher ( $P<0.001$ ) than that of *S. aureus* after the incubation phase, probably because of the greater ability of *S. epidermidis* to colonize biomaterial and produce biofilm.<sup>43,44</sup> There was no significant difference ( $P=0.979$ ) in this regard between the uncoated and coated disks.

After 15 minutes of ultraviolet irradiation, the concentrations of *S. aureus* and *S. epidermidis* decreased significantly

( $P<0.003$ ) in every group exposed to UV irradiation, indicating the lethal DNA damage produced by ultraviolet irradiation in cells.<sup>45,46</sup> However, it should be noted that our ultraviolet lamp power (5 mW/cm<sup>2</sup>) was within the range commonly found in the literature (0.5–20 mW/cm<sup>2</sup>).<sup>47</sup> The duration of ultraviolet exposure is a real problem. Such long exposure is not compatible with everyday practice. We are attempting to dope the titanium sol with a metallic compound in order to expand the layer absorbance spectrum to an ultraviolet-visible wavelength. Another way to improve antibacterial efficiency is to work on the crystallinity of the layer.

We found a bactericidal effect due to the photoactive TiO<sub>2</sub> coating after 1 hour of ultraviolet irradiation. This was indicated by a significant 5 log decrease in bacterial concentration for the two strains at the end of the test ( $P<0.001$ ). Complete bacterial inactivation was not achieved, probably because of the high initial bacterial concentration (4×10<sup>8</sup> CFU/mL) chosen for the study. Our results confirm that this process is also efficient for multi-resistant bacteria,<sup>19,48–50</sup> with only small differences between the two tested strains.

A number of facts could account for the delayed effect:

- The incubation phase enhanced the formation of biofilm, and in that state bacteria are less sensitive to reactive oxygen species.<sup>51</sup>
- The trypticase soy medium that promotes *staphylococcus* survival and inhibits bactericidal TiO<sub>2</sub> photoactive process by competitive interaction between free radical and organic materials, by absorption of ultraviolet radiation, and by decreasing the amount of direct contact between bacteria and TiO<sub>2</sub> molecules.<sup>36,52–55</sup>
- The high initial bacterial concentration (4×10<sup>8</sup> CFU/mL) may decrease the catalytic effect<sup>56,57</sup> because agglomeration of numerous bacterial cells with TiO<sub>2</sub> particles takes place.
- The tested bacterial strains were Gram-positive bacteria, which are known to be more resistant to the photoactive effect of TiO<sub>2</sub> than Gram-negative bacteria, due to their thick and complex cell wall.<sup>58,59</sup>
- Inadequate photocatalytic properties of our TiO<sub>2</sub> photoactive coating in regard to degradation tests using AO7 solution.

## Conclusion

Our study enabled us to develop an antibacterial coating for stainless steel commonly used in surgical practice. The process using photoactive TiO<sub>2</sub> exposed to ultraviolet irradiation

is actually well known and applied widely in disinfection procedures. Our model was effective against the two main bacterial strains involved in pin tract infections. Mechanical tests confirmed the ability of the coating to resist important stresses. Moreover, this type of coating is created by sol-gel dip-coating techniques that are not expensive and easily scalable for industrial application. We hope that this new option could prevent pin tract infection, even if heavy optimization work is yet to be done in order to amplify its bactericidal properties. The future research plan for this project is to improve the mechanical properties of the coating (eg, by adding of zirconium alkoxide) and to increase the bactericidal properties with adjunction of metallic ions (TiO<sub>2</sub> layer doping) in order to expand the layer absorbance spectrum to an ultraviolet-visible wavelength. Work will then be done on the annealing parameters in order to create a crack-free layer with suitable crystallinity to ensure better antibacterial efficiency. Finally, if the results are positive, in vivo tests would be envisaged.

## Acknowledgments

The authors thank Joel Cellier for the X-ray diffraction measurements and Lemlih Ouchchane for the statistical analysis.

## Disclosure

The authors report no conflicts of interest in this work.

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